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TABLE OF CONTENTS

HEADER	1
ABSTRACT	1
PLAIN LANGUAGE SUMMARY	2
SUMMARY OF FINDINGS	4
BACKGROUND	8
OBJECTIVES	9
METHODS	9
Figure 1.	11
RESULTS	13
DISCUSSION	15
AUTHORS' CONCLUSIONS	16
ACKNOWLEDGEMENTS	17
REFERENCES	18
CHARACTERISTICS OF STUDIES	22
DATA AND ANALYSES	27
Analysis 1.1. Comparison 1 Perinatal death review and audit as part of an intervention package including an educational outreach visit and morbidity/mortality conferences compared with no intervention, Outcome 1 Perinatal mortality rate.	28
Analysis 1.2. Comparison 1 Perinatal death review and audit as part of an intervention package including an educational outreach visit and morbidity/mortality conferences compared with no intervention, Outcome 2 Perinatal mortality related to suboptimal care.	28
Analysis 1.3. Comparison 1 Perinatal death review and audit as part of an intervention package including an educational outreach visit and morbidity/mortality conferences compared with no intervention, Outcome 3 Perinatal morbidity related to suboptimal care.	28
Analysis 2.1. Comparison 2 Maternal death review and audit as part of an intervention package including the ALARM course and training audit committees compared with no intervention, Outcome 1 Change in inpatient maternal mortality rate.	29
Analysis 2.2. Comparison 2 Maternal death review and audit as part of an intervention package including the ALARM course and training audit committees compared with no intervention, Outcome 2 Change in inpatient stillbirth rate.	29
Analysis 2.3. Comparison 2 Maternal death review and audit as part of an intervention package including the ALARM course and training audit committees compared with no intervention, Outcome 3 Change in inpatient neonatal mortality rate before 24 hours.	29
Analysis 2.4. Comparison 2 Maternal death review and audit as part of an intervention package including the ALARM course and training audit committees compared with no intervention, Outcome 4 Change in inpatient neonatal mortality rate after 24 hours.	30
ADDITIONAL TABLES	30
APPENDICES	33
WHAT'S NEW	48
CONTRIBUTIONS OF AUTHORS	48
DECLARATIONS OF INTEREST	48
SOURCES OF SUPPORT	49
DIFFERENCES BETWEEN PROTOCOL AND REVIEW	49

[Intervention Review]

Death audits and reviews for reducing maternal, perinatal and child mortality

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ABSTRACT

Background

The United Nations' Sustainable Development Goals (SDGs) include reducing the global maternal mortality rate to less than 70 per 100,000 live births and ending preventable deaths of newborns and children under five years of age, in every country, by 2030. Maternal and perinatal death audit and review is widely recommended as an intervention to reduce maternal and perinatal mortality, and to improve quality of care, and could be key to attaining the SDGs. However, there is uncertainty over the most cost-effective way of auditing and reviewing deaths: community-based audit (verbal and social autopsy), facility-based audits (significant event analysis (SEA)) or a combination of both (confidential enquiry).

Objectives

To assess the impact and cost-effectiveness of different types of death audits and reviews in reducing maternal, perinatal and child mortality.

Search methods

We searched the following from inception to 16 January 2019: CENTRAL, Ovid MEDLINE, Embase OvidSP, and five other databases. We identified ongoing studies using ClinicalTrials.gov and the World Health Organization (WHO) International Clinical Trials Registry Platform, and searched reference lists of included articles.

Selection criteria

Cluster-randomised trials, cluster non-randomised trials, controlled before-and-after studies and interrupted time series studies of any form of death audit or review that involved reviewing individual cases of maternal, perinatal or child deaths, identifying avoidable factors,

Death audits and reviews for reducing maternal, perinatal and child mortality (Review)

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and making recommendations. To be included in the review, a study needed to report at least one of the following outcomes: perinatal mortality rate; stillbirth rate; neonatal mortality rate; mortality rate in children under five years of age or maternal mortality rate.

Data collection and analysis

We used standard Cochrane Effective Practice and Organisation of Care (EPOC) group methodological procedures. Two review authors independently extracted data, assessed risk of bias and assessed the certainty of the evidence using GRADE. We planned to perform a meta-analysis using a random-effects model but included studies were not homogeneous enough to make pooling their results meaningful.

Main results

We included two cluster-randomised trials. Both introduced death review and audit as part of a multicomponent intervention, and compared this to current care. The QUARITE study (QUALity of care, Risk management, and TEchnology) concerned maternal death reviews in hospitals in West Africa, which had very high maternal and perinatal mortality rates. In contrast, the OPERA trial studied perinatal morbidity/mortality conferences (MMCs) in maternity units in France, which already had very low perinatal mortality rates at baseline.

The OPERA intervention in France started with an outreach visit to brief obstetricians, midwives and anaesthetists on the national guidelines on morbidity/mortality case management, and was followed by a series of perinatal MMCs. Half of the intervention units were randomised to receive additional support from a clinical psychologist during these meetings. The OPERA intervention may make little or no difference to overall perinatal mortality (low certainty evidence), however we are uncertain about the effect of the intervention on perinatal mortality related to suboptimal care (very low certainty evidence). The intervention probably reduces perinatal morbidity related to suboptimal care (unadjusted odds ratio (OR) 0.62, 95% confidence interval (CI) 0.40 to 0.95; 165,353 births; moderate-certainty evidence). The effect of the intervention on stillbirth rate, neonatal mortality, mortality rate in children under five years of age, maternal mortality or adverse effects was not reported.

The QUARITE intervention in West Africa focused on training leaders of hospital obstetric teams using the ALARM (Advances in Labour And Risk Management) course, which included one day of training about conducting maternal death reviews. The leaders returned to their hospitals, established a multidisciplinary committee and started auditing maternal deaths, with the support of external facilitators. The intervention probably reduces inpatient maternal deaths (adjusted OR 0.85, 95% CI 0.73 to 0.98; 191,167 deliveries; moderate certainty evidence) and probably also reduces inpatient neonatal mortality within 24 hours following birth (adjusted OR 0.74, 95% CI 0.61 to 0.90; moderate certainty evidence). However, QUARITE probably makes little or no difference to the inpatient stillbirth rate (moderate certainty evidence) and may make little or no difference to the inpatient neonatal mortality rate after 24 hours, although the 95% confidence interval includes both benefit and harm (low certainty evidence). The QUARITE intervention probably increases the percent of women receiving high quality of care (OR 1.87, 95% CI 1.35 - 2.57, moderate-certainty evidence). The effect of the intervention on perinatal mortality, mortality rate in children under five years of age, or adverse effects was not reported.

We did not find any studies that evaluated child death audit and review or community-based death reviews or costs.

Authors' conclusions

A complex intervention including maternal death audit and review, as well as development of local leadership and training, probably reduces inpatient maternal mortality in low-income country district hospitals, and probably slightly improves quality of care. Perinatal death audit and review, as part of a complex intervention with training, probably improves quality of care, as measured by perinatal morbidity related to suboptimal care, in a high-income setting where mortality was already very low.

The WHO recommends that maternal and perinatal death reviews should be conducted in all hospitals globally. However, conducting death reviews in isolation may not be sufficient to achieve the reductions in mortality observed in the QUARITE trial. This review suggests that maternal death audit and review may need to be implemented as part of an intervention package which also includes elements such as training of a leading doctor and midwife in each hospital, annual recertification, and quarterly outreach visits by external facilitators to provide supervision and mentorship. The same may also apply to perinatal and child death reviews. More operational research is needed on the most cost-effective ways of implementing maternal, perinatal and paediatric death reviews in low- and middle-income countries.

PLAIN LANGUAGE SUMMARY

Reviewing deaths to prevent mothers and children from dying in the future

What was the aim of this review?

This Cochrane Review aimed to assess if 'death audits and reviews' (exploring why people have died and what could have been done to prevent these deaths) can prevent mothers and children from dying. The review authors collected and analysed all relevant studies to answer this question and found two studies.

Key messages

In a study from West African hospitals, where death rates among women and babies were high, reviewing deaths probably led to fewer deaths among pregnant women, new mothers and newborn babies. In French hospitals, where death rates among babies were low, it may have made little or no difference to death rates among newborn babies.

What did the review study?

Every year, millions of babies and children die. Many women also die while they are pregnant or giving birth, or shortly afterwards. More than half of these deaths happen in sub-Saharan Africa.

In many settings, health facilities or communities carry out 'death audits and reviews'. Here, people explore why a person died, what could have been done to avoid this death and what could be done better in the future.

Death audits and reviews could potentially help improve the quality of care and prevent new deaths among mothers and children. But they could also cost money, be based on wrong information and take health workers away from other important tasks. If they are done badly, they could also make health workers feel blamed and humiliated, which could lead to poorer care. We need to find out if audits and reviews work and which approach works best.

The review authors searched for studies where people from health facilities or the community carried out audits or reviews of deaths of pregnant women, women who had recently given birth, newborn babies or children under five years of age. The studies had to compare places or times where death audits and reviews were used to places or times where they were not.

What were the main results of the review?

The review authors found two relevant studies. Both studies assessed death audits at health facilities.

The first study took place in West African hospitals with high death rates among women and babies. In this study, doctors and midwives were given extra training in pregnancy and childbirth care. This included one day of training in how to carry out death audits of women who had died during pregnancy or childbirth. They then returned to their hospitals and held audits at monthly meetings, with support from an expert from a different hospital. These hospitals were compared to hospitals without the training and audit meetings. For mothers and babies who were in hospital, this approach:

- probably led to fewer pregnant women and new mothers dying, and probably led to slightly better care for mothers;
- probably led to fewer babies dying during the first 24 hours. However, it may have made no difference to the number of babies who died after their first 24 hours, although the range where the actual effect may be (the "margin of error") includes both an increase and a decrease in the number of babies who died.
- probably made no difference to the number of stillbirths.

The second study took place in French hospitals that already had very few deaths among newborns. In this study, doctors and midwives were given information about pregnancy and childbirth guidelines. They then held audit meetings in their hospitals where they discussed stillbirths and newborn babies who had become sick or died. These hospitals were compared to hospitals without the information and the meetings. This approach:

- may have made little or no difference to the number of babies who died during their first week
- probably reduced the number of babies who were sick because they received poor quality care.

We don't know what the effect was on stillbirths or on the number of mothers or older babies and children who died because the study did not measure this.

How up-to-date was this review?

The review authors searched for studies that had been published up to 16 January 2019.

SUMMARY OF FINDINGS

Summary of findings for the main comparison. Perinatal death review and audit as part of an intervention package including an educational outreach visit and morbidity/mortality conferences compared with no intervention

Perinatal death review and audit as part of an intervention package including an educational outreach visit and morbidity/mortality conferences compared with no intervention

Patient or population: births

Settings: maternity units in France

Intervention: perinatal death review and audit as part of an intervention package including an educational outreach visit and morbidity/mortality conferences (OPERA trial)

Comparison: no intervention

Outcomes	Illustrative comparative risks* (95% CI)		Relative effect (95% CI)	No of participants (studies)	Certainty of the evidence (GRADE)	Results in words
	Assumed risk	Corresponding risk				
	Comparison	With death reviews				
Perinatal mortality rate (overall) (at month 12–26)	4.7 stillbirths or deaths per 1000 total births	4.9 stillbirths or deaths per 1000 total births (from 4 to 6 stillbirths or deaths per 1000)	OR 1.05 (0.91 to 1.21)	95 maternity units, 165353 births (1 study ^a)	⊕⊕⊕⊕ Low^b	The intervention may make little or no difference to perinatal mortality.
Proportion of perinatal deaths related to suboptimal care^c (at months 12–26)	85 per 1000 stillbirths or deaths whose quality of care could be scored	90 per 1000 stillbirths or deaths whose quality of care could be scored (from 49 to 181 stillbirths or deaths per 1000)	OR 1.14 (0.55 to 2.37)	95 maternity units, 759 stillbirths or deaths whose quality of care could be scored (1 study ^a)	⊕⊕⊕⊕ Very Low	We are uncertain about the effect of the intervention on perinatal mortality related to suboptimal care
Stillbirth rate	—	—	—	—	—	Not reported

Neonatal mortality rate	—	—	—	—	—	Not reported
Mortality rate in children < 5 years of age	—	—	—	—	—	Not reported
Maternal mortality rate	—	—	—	—	—	Not reported
Proportion of perinatal morbidity cases related to suboptimal care^c (at months 12–26)	115 per 1000 morbidity cases whose quality of care could be scored	76 per 1000 morbidity cases whose quality of care could be scored (from 49 to 110 per 1000 morbidity cases)	OR 0.62 (0.40 to 0.95)	95 maternity units, 1640 cases of morbidity whose quality of care could be scored (1 study ^d)	⊕⊕⊕⊖ Moderate^e	The intervention probably reduces perinatal morbidity related to suboptimal care.
Adverse effects	-	-	-	-	-	Not reported

*The basis for the **assumed risk** (e.g. the median control group risk across studies) is provided in footnotes. The **corresponding risk** (and its 95% confidence interval) is based on the assumed risk in the comparison group and the **relative effect** of the intervention (and its 95% CI).

CI: confidence interval; **OR:** odds ratio.

GRADE Working Group grades of evidence

High certainty: further research is very unlikely to change our confidence in the estimate of effect.

Moderate certainty: further research is likely to have an important impact on our confidence in the estimate of effect and may change the estimate.

Low certainty: further research is very likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate.

Very low certainty: we are very uncertain about the estimate.

^aDupont 2017 (cluster randomised trial).

^bDowngraded two levels due to limitations in study design and execution and imprecise estimate. The 95% CI included both slight harm and appreciable benefit.

^cThe proportion here refers to the proportion of cases related to suboptimal care.

^dDowngraded three levels due to limitations in study design and very imprecise estimate. The 95% CI included both appreciable harm and appreciable benefit.

^eDowngraded one level due to limitations in study design and execution.

Summary of findings 2. Maternal death review and audit as part of an intervention package including the ALARM course and training audit committees compared with no intervention

Maternal death review and audit as part of an intervention package including the ALARM course and training audit committees compared with no intervention

Patient or population: mothers delivering in the hospitals

Settings: district and regional referral hospitals in West Africa

Intervention: complex intervention to develop local leadership and empower obstetric teams to conduct maternal death audits (QUARITE trial)

Comparison: no intervention

Outcomes	Anticipated absolute effects ^{a,f}		Relative effect (95% CI)	No of participants (studies)	Certainty of the evidence (GRADE)	Results in words
	Assumed risk	Corresponding risk				
	With no intervention	With death reviews				
Perinatal mortality rate	—	—	—	—	—	Not reported
Inpatient stillbirth rate	94 stillbirths per 1000 total births	98 stillbirths per 1000 total births (from 86 to 112 stillbirths per 1000)	AOR^a 1.05 (0.91 to 1.22)	46 hospitals, 197,336 participants (1 study ^b)	⊕⊕⊕⊖ Moderate^c	The intervention probably makes little or no difference to inpatient stillbirth rate.
Inpatient neonatal mortality rate – before 24 hours	11 neonatal deaths per 1000 live births	8 neonatal deaths per 1000 live births (from 7 to 10 deaths per 1000)	AOR^a 0.74 (0.61 to 0.90)	46 hospitals, 197,336 participants (1 study ^b)	⊕⊕⊕⊖ Moderate^c	The intervention probably reduces inpatient neonatal mortality rate before 24 hours.
Inpatient neonatal mortality rate – after 24 hours	2 neonatal deaths per 1000 live births	2 neonatal deaths per 1000 live births (from 1 to 3 deaths per 1000)	AOR^a 0.88 (0.62 to 1.24)	46 hospitals, 197,336 participants (1 study ^b)	⊕⊕⊕⊖ Low^d	The intervention may make little or no difference to inpatient neonatal mortality rate after 24 hours. However, the 95% confidence interval indicates that the intervention may reduce or increase inpatient neonatal mortality rate after 24 hours.
Mortality rate in children < 5 years of age	—	—	—	—	—	Not reported
Inpatient maternal mortality rate	711 maternal deaths per 100000 pregnant women	605 maternal deaths per 100000 pregnant women (from 520 to 697 deaths per 100000) ^g	AOR^a 0.85 (0.73 to 0.98)	46 hospitals, 197,336 participants (1 study ^b)	⊕⊕⊕⊖ Moderate^c	The intervention probably reduces inpatient maternal mortality.

Quality of care (Proportion of women receiving high quality care ^d)	298 women per 1000 pregnant women received high quality care	442 women per 1000 pregnant women received high quality care (from 364 to 522 women per 1000)	OR 1.87 (1.35 - 2.57)	32 hospitals, 658 consecutive participants (1 study ^b)	⊕⊕⊕⊖ Moderate^c	The intervention probably increases the proportion of women receiving high quality of care.
Quality of care for women with complications (Proportion of women with eclampsia or postpartum haemorrhage receiving high quality care ^d)	377 women per 1000 pregnant women with complications received high quality care	503 women per 1000 pregnant women with complications received high quality care (from 368 to 638 women per 1000)	OR 1.67 (0.96 - 2.91)	32 hospitals, 209 complicated participants (1 study ^b)	⊕⊕⊖⊖ Low^e	The intervention may increase the proportion of women with complications who receive high quality of care. However, the 95% confidence interval includes no effect.
Adverse effects	-	-	-	-	-	Not reported

*The basis for the **assumed risk** (e.g. the median control group risk across studies) is provided in footnotes. The **corresponding risk** (and its 95% confidence interval) is based on the assumed risk in the comparison group and the **relative effect** of the intervention (and its 95% CI).

AOR: adjusted odds ratio; **CBCA:** criterion-based clinical audit; **CI:** confidence interval.

GRADE Working Group grades of evidence

High certainty: further research is very unlikely to change our confidence in the estimate of effect.

Moderate certainty: further research is likely to have an important impact on our confidence in the estimate of effect and may change the estimate.

Low certainty: further research is very likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate.

Very low certainty: we are very uncertain about the estimate.

^a Adjusted for the two stratification variables: hospital type and country, as well as for variables selected a priori as potential risk factors for hospital-based mortality, including both (a) baseline (year 1) characteristics of hospitals (availability of adult intensive care unit, blood bank, anaesthetist, and gynaecologist-obstetrician) and (b) characteristics of individual women (residence, age, parity, previous caesarean delivery, any pathology during pregnancy, prenatal visit attendance, multiple pregnancy, referral from another health facility, antepartum or postpartum haemorrhage, pre-eclampsia or eclampsia, prolonged or obstructed labour, uterine rupture, and puerperal infection or sepsis).

^b [Dumont 2013](#) (cluster randomised trial). Perinatal outcomes were assessed for singletons only, excluding multiple pregnancies from the analyses.

^c Downgraded one level for indirectness because this is based on a single study with a relatively small number of events. For a complex intervention such as this, the effect of the intervention may be modified by setting, or may work differently in a different setting.

^d Defined as CBCA score of >70%

^e Downgraded two levels due to imprecision of the estimate, and for indirectness because this is based on a single study with a small number of events.

^f These anticipated absolute effects are based on the numbers of events and participants in the year 4 outcome assessment for the trial (see [Table 1](#)).

^g Note that the denominator here is pregnant women and not the number of live births.

BACKGROUND

Description of the condition

The United Nations' Sustainable Development Goals include reducing the global maternal mortality rate to fewer than 70 per 100,000 live births and ending preventable deaths of newborns and children under five years of age. All countries aim to reduce neonatal mortality to at least 12 per 1000 live births and mortality in children aged less than five years to at least 25 per 1000 live births, by 2030 (UN 2017). Although progress is being made towards achieving these goals, it is not fast enough, especially in low-income countries (Wang 2014; WHO 2014). The absolute numbers and rates of maternal, child and perinatal deaths are higher in Africa than in any other region. In 2015, there were an estimated 303,000 maternal deaths globally, 99% of which were in low- and middle-income countries, and 66% in sub-Saharan Africa alone (WHO 2015). In 2016, there were an estimated 5,642,000 child deaths globally, more than half of which occurred in sub-Saharan Africa (UNICEF 2017).

For this review, we used the following definitions.

- 'Maternal mortality': death of a woman during pregnancy or within 42 days of delivery or termination of pregnancy, irrespective of the duration and site of the pregnancy, from any cause related to or aggravated by the pregnancy or its management, but not from accidental or incidental causes (WHO 2004).
- 'Perinatal mortality': stillbirth or death of a newborn baby within the first seven days of life (WHO 2006).
- 'Neonatal mortality': death of a newborn baby at any time from birth to 28 days (WHO 2006).
- 'Child mortality': death of a child under the age of five years (UNICEF 2015).
- Maternal mortality ratio: number of maternal deaths per 100,000 live births (UNICEF 2015; WHO 2014).
- Perinatal mortality rate: number of stillbirths and deaths in the first week of life per 1000 total births (WHO 2006).
- Neonatal mortality rate: number of babies who die from 0 to 28 days per 1000 live births (WHO 2006).
- Child mortality rate: number of deaths in children under age five years per 1000 live births (WHO 2006).

There is some overlap between the 'perinatal' and 'neonatal' categories, which both include babies aged zero to seven days, and between the 'neonatal' and 'child' categories, which both include babies aged zero to 28 days.

Description of the intervention

'Death audit and review' is a broad term intended to include every different method of reviewing deaths, that not only identifies the medical cause of death, but also attempts to identify avoidable factors that contributed to the death and make recommendations for avoiding such deaths in the future. The principal methods used are community-based audit (verbal and social autopsy), facility-based audits such as significant event analysis (SEA) or morbidity and mortality conferences (MMCs), and a combination of both (e.g. through a 'confidential enquiry').

In low-income countries without comprehensive death registration, deaths in the community are often investigated using verbal autopsy. The family of the deceased is interviewed according to a standard questionnaire (developed by WHO 2007), and the information is then interpreted by physicians or by a computer to ascertain the most likely medical cause of death (Waiswa 2010). However, there is usually no attempt to identify avoidable factors as it is assumed that it is already known which interventions are needed to tackle each disease. Verbal autopsy has been incorporated into wider health and demographic surveillance strategies (Adazu 2005), although its accuracy has been questioned due to the non-specific nature of signs and symptoms that may not be easily observed or remembered at interview (Butler 2010; Sloan 2001; Waiswa 2010). Social autopsy was designed as an add-on to verbal autopsy, and indeed the two are sometimes combined as a 'verbal and social autopsy' (VASA) (Kalter 2011). The aim is to make a 'social diagnosis', identifying avoidable factors prior to death in the home, community and within health facilities at different stages of the patient pathway (e.g. parents do not recognise the severity of the illness; parents delay seeking care or seek care from an inappropriate provider; there are delays in reaching the health facility because of transport problems; after arriving at the facility the patient does not receive adequate treatment or has to wait excessively). In India, this has been used in a participatory manner, which has been termed social audit for community action (SACA) (Nandan 2005). In this method, the community is asked to identify causes of death and avoidable factors. In this review, we included combined VASA studies, but not stand-alone verbal autopsy (whether conducted by a physician or a computer) with the intention of only identifying the medical cause of death.

Death audits in health facilities are usually based on MMCs or SEA. Cases are usually discussed in a multidisciplinary team meeting (Hussein 2007). After discussing the details of the case, health workers identify avoidable factors and learning needs, and propose actions to be taken and changes to be made. The process does not intend to place blame, but the names of staff involved are not kept confidential. Indeed it is argued that "non-confidential straightforwardness and open-mindedness" are vital for a successful strategy (Supratikto 2002). A similar process occurs in 'mortality meetings', 'root cause analysis' meetings and, indeed, 'serious case reviews' (in child protection cases). Most mortality meetings take place at secondary healthcare facilities, drawing upon medical records to identify the diagnosis and key management interventions. Severe morbidity or near-miss reviews are similarly used to learn lessons; these examine cases in which an individual almost died. In the UK, participation in SEAs are an important part of revalidation for doctors, and the Royal College of General Practitioners recommends that "SEA team discussions should be a routine part of your practice's quality improvement and clinical governance" (RCGP 2014).

Confidential enquiry is the most comprehensive method by which to investigate deaths. It considers the diagnosis and treatment in health facilities, and the entire course of an illness and treatment-seeking pathway, identifying avoidable factors and recommending changes at every level of the health and social care system and beyond, in order to prevent future deaths. This is particularly important in low-income countries where the majority of child deaths occur outside of health facilities (Bremner 2001). A key feature of such enquiries is that the names of the individuals and any health workers involved are kept confidential, so that blame is

avoided. These enquiries were pioneered in high-income countries, based entirely on written (usually medical) records examined by a multidisciplinary panel of experts, which includes health workers and other professionals such as social services and the police (Lewis 2011; Pearson 2008a). Such confidential enquiries have been useful for evaluating gaps in health care in the UK (Pearson 2008a; WHO 2004), but are not yet widely used in low-income countries (Hussein 2007). The expert analysis involves both quantitative and qualitative elements. In the UK, all the included deaths are analysed quantitatively for basic information such as age, sex, socioeconomic status, location of death, time (seasonality) of death and cause of death. Further detailed investigations are carried out on all maternal deaths and a subset of child deaths. A multidisciplinary panel reviews each of these cases and identifies avoidable factors. These are analysed thematically, illustrated by cases, and were used to generate recommendations as to how deaths might be avoided in the future (Pearson 2008b; Knight 2019).

How the intervention might work

Participation by communities in death audits is a strong basis for collective action to reduce mortality. In health facilities, significant event audit is a potentially powerful intervention to enable staff to learn from their mistakes and to institute important changes to procedures within their institution; the key mechanism is believed to be recommendation, followed by implementation of the proposed solutions (Pattinson 2009). The confidential enquiry approach is designed to identify avoidable factors at every step of the treatment-seeking pathway, and to make recommendations to improve the health system and to address avoidable factors outside of health facilities. Case review meetings, followed by the dissemination of recommendations to health workers, communities, or both, are essentially aiming to change clinician and patient behaviour.

Although there are many frameworks for classifying behaviour change interventions, the only comprehensive and conceptually coherent one is the 'theoretical domains framework', which consists of 14 domains (Cane 2012; Michie 2014). Many of these domains are addressed by death audit and review. Those participating in the death review meetings gain knowledge about avoidable factors. The recommendations often set goals, and progress towards these can be audited. Repetition of similar recommendations may help clinicians to recall guidelines, whereas social pressure may encourage better adherence. Death review meetings may also change health workers' beliefs about the consequences of their actions: the knowledge that deaths will be investigated and reviewed may motivate them to avoid poor practice. Discussing deaths, especially of mothers and children, often evokes an emotional response, which usually motivates health workers and parents to do all they can to prevent such deaths.

Death reviews may conceivably have some adverse effects. First, there is a cost (time and financial) to conducting death reviews. In the community, field workers need to be employed to investigate cases. In health facilities, staff are taken away from frontline duties to review cases, which may have an adverse impact on the delivery of care. It has been argued that these resources should instead be spent directly on implementing interventions that are known to be effective (Koblinsky 2017). Second, if death reviews are not handled sensitively, they may lead to blaming, humiliation and demotivation of staff, which may in turn lead

to poorer quality of care. Third, focus on only one level of care (such as a district hospital) may lead to the diversion of resources away from other levels of care (such as primary care facilities). Fourth, there is the potential for errors – reviews based on indirect information (especially at the community level) may be incomplete or inadequate at diagnosing the likely cause of death, because the information available may be insufficient or inaccurate (or both), or the people discussing the cases may be inexperienced, or a combination of these.

Why it is important to do this review

The World Health Organization (WHO) recommends that health facilities should conduct maternal and perinatal death reviews (MPDRs) (WHO 2013; WHO 2016). In general, there is an underlying assumption that death reviews are useful and will impact on mortality but there is little robust evidence to support this (Pattinson 2005). It would be useful for policy-makers to understand which type of death review has the greatest impact on maternal, perinatal and child death rates, and what the essential features of an effective death review process are. Although confidential enquiry seems to be the most comprehensive method for addressing a wide range of avoidable factors, and hence has the potential for the greatest impact, it is unclear whether it could be adapted, whether it would be feasible or whether it would be effective in reducing mortality in low-income countries. There is no comprehensive systematic review in the literature examining the impact of these methods of investigating deaths.

OBJECTIVES

To assess the impact and cost-effectiveness of different types of death audits and reviews in reducing maternal, perinatal and child mortality.

METHODS

Criteria for considering studies for this review

Types of studies

We included cluster randomised trials. However, as these are expensive and difficult to conduct, and large sample sizes are needed to measure impact on mortality, we anticipated that we would find very few. Therefore, we also included cluster non-randomised trials, studies with a step-wedge design, controlled before-and-after studies and interrupted time series studies.

For cluster randomised trials, cluster non-randomised trials and controlled before-and-after studies, we used the EPOC criteria (EPOC 2017a), and excluded studies with only one intervention or control site. For interrupted time series studies, we excluded studies that did not have a clearly defined point in time when the intervention occurred and at least three data points before and three after the intervention.

Types of participants

Participants receiving the intervention (audits and reviews of maternal, perinatal or child deaths) could be involved with health facilities of any level or the wider community, such as subdistricts or districts, or both. Participants who should benefit from the intervention were pregnant women giving birth, their neonates and their children under five years old at the study sites during the study period in which the outcomes are measured.

Types of interventions

We included any form of death audit or review that involved studying individual cases of maternal, perinatal or child deaths, identifying avoidable factors and making recommendations. We classified the interventions as verbal and social autopsy, facility-based death audit and SEA, or confidential enquiry. We excluded verbal autopsy studies that evaluated only causes of death and not avoidable factors. We included studies of maternal, perinatal, newborn and child deaths, alone or in combination. We excluded severe morbidity or near-miss reviews. We included comparisons of the same population before introduction of the death review, or other comparable communities in which the death review was not implemented.

Types of outcome measures

We planned to include studies in the review irrespective of whether measured outcome data were reported in a 'usable' way.

Main outcomes

To be included in the review, a study needed to report at least one of the following outcomes:

- perinatal mortality rate;
- stillbirth rate;
- neonatal mortality rate;
- mortality rate in children under five years of age;
- maternal mortality rate.

Secondary outcomes

For included studies, we also considered other outcomes:

- outcomes relating to maternal severe morbidity, such as maternal near miss or as defined by study authors;
- outcomes relating to quality of care in participating facilities;
- perinatal morbidity related to suboptimal care;
- cost per death averted.

Search methods for identification of studies

Electronic searches

We searched the following databases from database inception to 16 January 2019:

- Cochrane Central Register of Controlled Trials (CENTRAL; in the Cochrane Library; 2019, Issue 1 of 12);

- Ovid MEDLINE(R) Epub Ahead of Print, In-Process & Other Non-Indexed Citations, Ovid MEDLINE(R) Daily, Ovid MEDLINE and Versions(R) (OvidSP; from 1946);
- Embase (OvidSP; from 1974);
- Global Health (OvidSP; from 1973);
- Global Health Library – Regional Indexes, WHO; (www.globalhealthlibrary.net/php/index.php);
- NHS Economic Evaluation Database (NHS EED; in the Cochrane Library; 2015, Issue 2 of 4);
- Popline, K4Health (www.popline.org/advancedsearch);
- Cumulative Index to Nursing and Allied Health Literature Host (CINAHL EBSCOHost; from 1982);
- Science Citation Index & Conference Proceedings Citation Index – Science (Web of Science Core Collection, Thomson Reuters; from 1945).

There were no language or publication date limits. See [Appendix 1](#) for search strategies.

Searching other resources

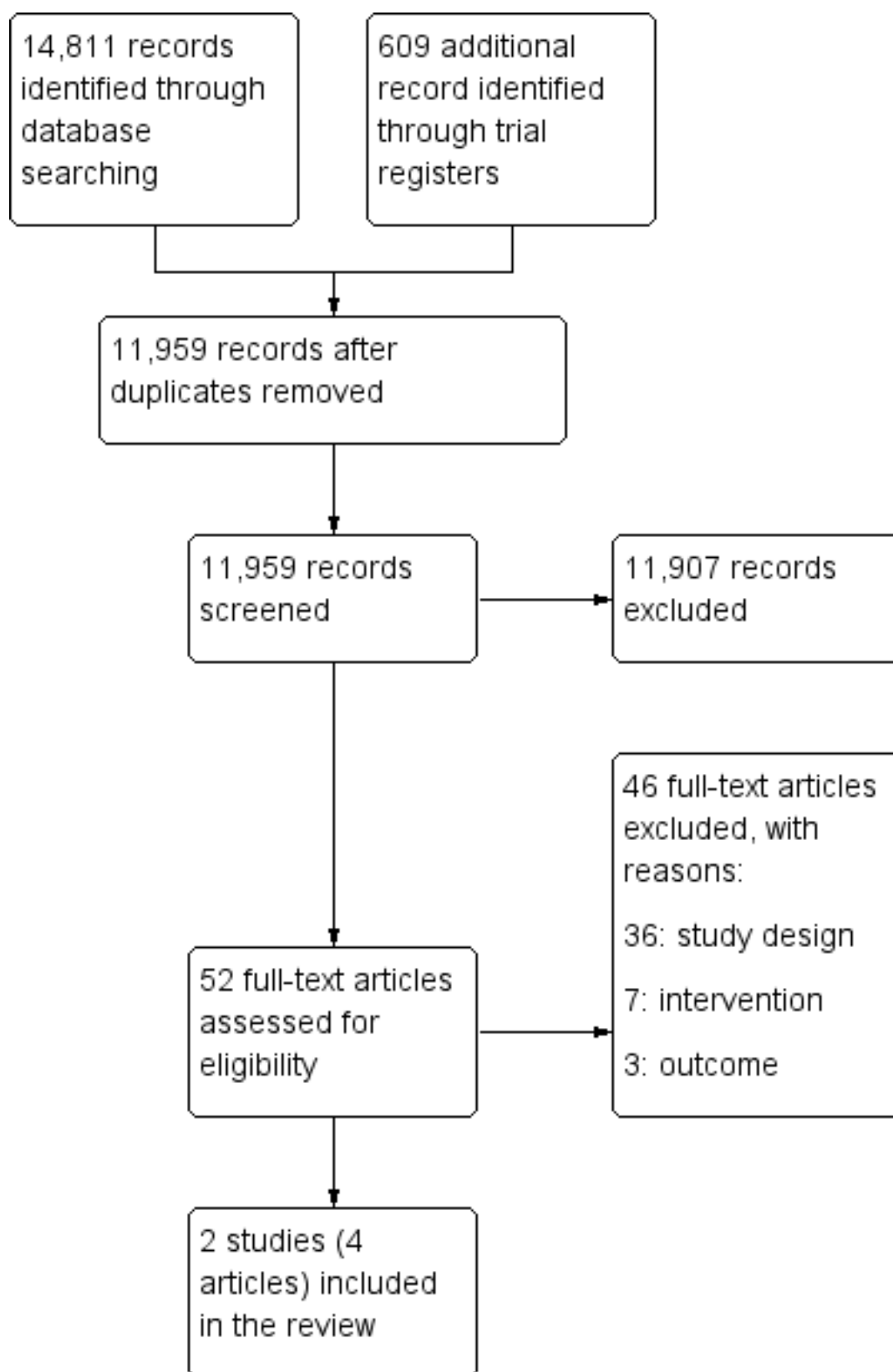
We identified ongoing studies through searches of ClinicalTrials.gov (clinicaltrials.gov/) and the WHO International Clinical Trials Registry Platform (www.who.int/ictpr/en/). We screened the reference lists of relevant articles found in these searches. We contacted experts in the field to advise us of unpublished or grey literature.

Data collection and analysis

Selection of studies

We downloaded all titles and abstracts retrieved by electronic searching to a reference management database and removed duplicates. Two review authors independently screened titles and abstracts for inclusion. We retrieved the full-text study reports/publications and two review authors independently screened the full text, identified studies for inclusion and recorded the reasons for exclusion of ineligible studies. We resolved any disagreements through discussion and when required, we consulted a third review author. We listed studies that initially appeared to meet the inclusion criteria but were later excluded, together with reasons for exclusion, in the [Characteristics of excluded studies](#) table. We collated multiple reports of the same study so that each study, rather than each report, was the unit of interest in the review. We intended to provide any information we could obtain about ongoing studies. We recorded the selection process in a PRISMA flow diagram ([Figure 1](#)) ([Liberati 2009](#)) and the [Characteristics of excluded studies](#) table.

Figure 1. Study flow diagram.



Data extraction and management

We used a standard data collection form, adapted from the Cochrane good practice data collection form, for study characteristics and outcome data. We piloted this on one study in the review.

Two review authors (MW, JP) independently extracted the following study characteristics from the included studies.

- Methods: study design, number of study centres and location, study setting, withdrawals, date of study, follow-up.
- Participating health facilities: number, inclusion criteria, exclusion criteria, other relevant characteristics.
- Interventions: intervention components, comparison, fidelity assessment.
- Outcomes: main and other outcomes specified and collected, time points reported.
- Notes: funding for trial, notable conflicts of interest of trial authors, ethical approval.

Two review authors (MW and JP) independently extracted outcome data from included studies. We noted in the [Characteristics of included studies](#) table whether outcome data were reported in an unusable way. We resolved disagreements by consensus.

Assessment of risk of bias in included studies

Two review authors (MW and JP) independently assessed the risk of bias for each study using the criteria outlined in the *Cochrane Handbook for Systematic Reviews of Interventions* ([Higgins 2011](#)), and the guidance from the EPOC group ([EPOC 2017b](#)). We contacted the authors of each study for clarification when the publication lacked clarity on whether a criterion was met. We resolved any disagreements by discussion. We assessed the risk of bias according to the following domains.

Cluster randomised trial/cluster non-randomised trial/controlled before-and-after study criteria:

- random sequence generation;
- allocation concealment;
- blinding of participants and personnel;
- blinding of outcome assessment;
- incomplete outcome data;
- selective outcome reporting;
- baseline outcomes measurement;
- baseline characteristics;
- other bias.

We judged each potential source of bias as high, low or unclear, and provided a quote from the study report together with a justification for our judgement in the 'Risk of bias' table. We summarised our 'Risk of bias' judgements across different studies for each of the domains listed. We considered blinding separately for different key outcomes where necessary (e.g. for unblinded outcome assessments, risk of bias for all-cause mortality may be very different from a participant-reported pain scale). Where information on risk of bias related to unpublished data or correspondence with a trialist, we noted this in the 'Risk of bias' table. We did not exclude studies on the grounds of their risk of bias,

but clearly reported the risk of bias when presenting the results of the studies.

When considering treatment effects, we took into account the risk of bias for the studies that contributed to that outcome.

Assessment of bias in conducting the systematic review

We conducted the review according to the published protocol ([Willcox 2018](#)), and reported any deviations from it in the [Differences between protocol and review](#) section.

Measures of treatment effect

We estimated the effect of the intervention using odds ratios (OR) for dichotomous data, together with the appropriate associated 95% confidence intervals (CI). For the Summary of Findings Tables, illustrative comparative risks were calculated using GRADEPro software.

We planned to estimate mean differences (for studies using the same scale) or standardised mean differences (for studies using different scales) for continuous data, together with the 95% appropriate associated CIs, but we found no studies reporting continuous data. For interrupted time series studies, we planned to estimate a standardised effect size for each study by dividing the level by the slope and the standard error by the standard deviation of the preintervention slope, but we found no interrupted time series studies.

Unit of analysis issues

For cluster randomised trials, we conducted the analysis at the same level as the allocation using a summary measure from each cluster.

Dealing with missing data

We contacted study authors to request missing data. We used intention-to-treat analyses by including all participants who were supposed to have received a particular intervention. We planned to perform sensitivity analyses by excluding studies with high rates of loss to follow-up.

Assessment of heterogeneity

We planned to assess statistical heterogeneity in each meta-analysis visually and using the I^2 and χ^2 statistics, regarding heterogeneity as substantial if the I^2 statistic was greater than 60% or if there was a low P value (less than 0.10) in the χ^2 test for heterogeneity. However, we performed no meta-analyses.

Assessment of reporting biases

If there were 10 or more studies in the meta-analysis, we planned to investigate reporting biases (such as publication bias) using funnel plots. However, only found two studies. We contacted authors to clarify points which were not explicit in their publications.

Data synthesis

While we planned to perform meta-analysis following Cochrane and EPOC guidance (see the protocol for this review for full details; [Willcox 2018](#)), we judged that this was not possible and did not perform meta-analyses. Therefore, we summarised results extracted from the included studies narratively.

Summary of findings

We summarised the findings of the main intervention comparison for the most important outcomes:

- perinatal mortality rate;
- stillbirth rate;
- neonatal mortality rate;
- mortality rate in children under five years of age;
- maternal mortality rate;
- outcomes relating to quality of care in participating facilities;
- perinatal morbidity related to suboptimal care.

We present these in 'Summary of findings' tables to draw conclusions about the certainty of the evidence within the text of the review. Where a study presented a change in rates over time, we calculated the illustrative effect of the intervention by applying the adjusted odds ratio for the intervention to the rates observed in the control group at follow-up, using GRADEpro software.

Two review authors independently assessed the certainty of the evidence (high, moderate, low or very low) using the five GRADE considerations (study limitations, consistency of effect, imprecision, indirectness and publication bias). We used the methods and recommendations described in Section 8.5 and Chapter 12 of the *Cochrane Handbook for Systematic Reviews of Interventions* (Higgins 2011) and the EPOC worksheets (EPOC 2017c), and using GRADEpro software (GRADEpro GDT). We resolved disagreements on certainty ratings by discussion, providing justification for decisions to downgrade or upgrade the ratings using footnotes in the table, and made comments to aid readers' understanding of the review where necessary. We used plain language statements to report these findings in the review.

As it was not possible to meta-analyse the data, we summarised the results in the text.

Subgroup analysis and investigation of heterogeneity

We planned to carry out the following subgroup analyses:

- type of country (low- versus middle- versus high-income, according to World Bank classification at the time of the study);
- type of death review (verbal and social autopsy versus SEA versus confidential enquiry);
- setting: facility-based versus community-based.

The following outcomes would be used in subgroup analysis:

- perinatal mortality rate;
- stillbirth rate;
- neonatal mortality rate;
- mortality rate in children under five years of age;
- maternal mortality rate.

It was not possible to undertake these analyses due to the decision not to pool data across the included studies.

Sensitivity analysis

We planned to perform sensitivity analysis defined a priori to assess the robustness of our conclusions and explore its impact on effect sizes. This was to involve:

- restricting the analysis to published studies;
- restricting the analysis to studies with a low risk of bias (i.e. high-quality randomised trials).

It was not possible to undertake these analyses due to the decision not to pool data across the included studies.

RESULTS

Description of studies

Results of the search

We identified 11,959 articles from electronic and supplementary searches, after removing duplicates. We excluded 11,907 articles following a review of titles and abstracts and retrieved and assessed the full text of 52 articles (Figure 1). We excluded 46 full-text articles; 36 of these did not meet our criteria for study design (most were uncontrolled before-and-after studies, see Table 2); seven did not include death review or audit as part of the intervention; and three did not measure impact on maternal, perinatal or child mortality. Two cluster-randomised trials (one of them reported in three separate articles) met the inclusion criteria. We have presented the study flow diagram in Figure 1.

Included studies

Only two studies (four articles) met all the inclusion criteria and are described in the [Characteristics of included studies](#) table (Dumont 2013; Dupont 2017). Both introduced death review and audit as part of a complex intervention, which had different components. The QUARITE study (QUALity of care, Risk management, and TEchnology) was conducted in hospitals in Mali and Senegal (Dumont 2013), which had very high maternal and perinatal mortality rates. The intervention focused on training leaders of obstetric teams in capital, regional and district hospitals using the ALARM (Advances in Labour And Risk Management) international course, which included one day about conducting maternal death reviews. The leaders then returned to their hospitals where they established a multidisciplinary committee and started auditing maternal deaths, with the support of external facilitators every quarter. In contrast, the OPERA trial was conducted in maternity units in France, which already had very low perinatal mortality rates at baseline (Dupont 2017). The intervention started with an outreach visit to brief obstetricians, midwives and anaesthetists on the national guidelines on morbidity/mortality case management, and was followed by a series of morbidity/mortality conferences (MMCs). Half of the intervention units were randomised to receive additional support from a clinical psychologist during these meetings.

Excluded studies

Twenty-six of the excluded full-text articles were uncontrolled before-and-after studies. Eight were excluded because they did not include participatory death review meetings.

Risk of bias in included studies

We assessed the risk of bias of both trials (Dumont 2013; Dupont 2017). Because of the nature of the intervention, it was not possible to blind the participating health facilities or their staff, but patients were blinded to the allocation of the hospital they were attending in the QUARITE study (Dumont 2013). In both cases, the outcome

assessors were blinded. Overall, the QUARITE trial had a low risk of bias whereas the OPERA trial had a moderate risk of bias.

Allocation

There was a low risk of selection bias in both studies because both used stratified randomisation to allocate health facilities to intervention or control groups. In the OPERA trial, the intervention and control groups had similar characteristics at baseline (Dupont 2017). In the QUARITE trial, there were some important differences between intervention and control groups at baseline, but these were accounted for in the analysis, which measured change in mortality rates from baseline (Dumont 2013).

Blinding

Blinding of health facilities and their staff was not possible due to the nature of the intervention. In the QUARITE trial, it is stated that patients were blinded to allocation of the facility (Dumont 2013). In both trials, the outcome assessors were blinded. In the OPERA trial, the outcome assessors had previously been involved in the mortality meetings, but because the outcomes were assessed from anonymised medical records one year after the meetings, and only a sample of cases were discussed at meetings, it is unlikely that the assessors would remember whether cases came from intervention or control hospitals, so we judged this at low risk of bias (Dupont 2017).

Incomplete outcome data

There was low risk of attrition bias in the QUARITE trial because no hospitals were lost to follow-up (Dumont 2013). In the OPERA trial, 2/97 private hospitals withdrew, but this was prior to randomisation (low risk of attrition bias; Dupont 2017).

Selective reporting

The protocol of the QUARITE trial was published and the trial reported the outcomes specified in the protocol, so there was no selective reporting (Dumont 2013). The authors of the OPERA confirmed that there were no outcomes in the protocol which were not reported in the final trial report (Dupont 2017).

Other potential sources of bias

In the OPERA trial, six units randomised to the intervention group did not implement the intervention and were transferred to the control group. Therefore, the analysis was per protocol rather than intention to treat (Dupont 2017).

Effects of interventions

See: [Summary of findings for the main comparison Perinatal death review and audit as part of an intervention package including an educational outreach visit and morbidity/mortality conferences compared with no intervention](#); [Summary of findings 2 Maternal death review and audit as part of an intervention package including the ALARM course and training audit committees compared with no intervention](#)

Comparison 1: Perinatal death review and audit as part of an intervention package including an educational outreach

visit and morbidity/mortality conferences compared with no intervention

One included study (the OPERA trial) assessed a complex intervention to promote guidelines and MMCs compared with no intervention and reported perinatal mortality and quality of care (Dupont 2017). There were no data on stillbirth rate, neonatal mortality, mortality rate in children under five years of age, maternal mortality, maternal morbidity or cost effectiveness. No undesirable effects of the intervention were reported. ([Summary of findings for the main comparison](#); [Table 3](#)).

Primary outcomes

Perinatal mortality

The intervention may have made little or no difference to overall perinatal mortality in this setting (OR 1.05, 95% CI 0.91 to 1.21; low certainty evidence). We are uncertain of the effect of the intervention on perinatal mortality specifically related to suboptimal care as the certainty of the evidence is very low (OR 1.14, 95% CI 0.55 to 2.37; very low-certainty evidence) ([Analysis 1.1](#); [Analysis 1.2](#)) (Dupont 2017).

Secondary outcomes

Perinatal morbidity related to suboptimal care

The OPERA intervention probably reduced perinatal morbidity which is specifically related to suboptimal care (OR 0.62, 95% CI 0.40 to 0.95; moderate-certainty evidence; [Analysis 1.3](#)).

Comparison 2: Maternal death review and audit as part of an intervention package including the ALARM course and training audit committees compared with no intervention

One included study (the QUARITE trial) assessed a complex intervention to develop local leadership and empower obstetric teams to conduct maternal death reviews compared with no intervention. It reported maternal mortality, neonatal mortality, perinatal morbidity, stillbirths and quality of care (Dumont 2013). There were no data on perinatal mortality, mortality rate in children under five years of age, maternal morbidity or cost-effectiveness. No undesirable effects of the intervention were reported. ([Summary of findings 2](#); [Table 1](#)).

Primary outcomes

Stillbirths

The QUARITE intervention probably made little or no difference to the inpatient stillbirth rate (adjusted OR 1.05, 95% CI 0.91 to 1.22; moderate certainty evidence; [Analysis 2.2](#)).

Neonatal mortality

The QUARITE intervention probably reduced inpatient neonatal mortality within 24 hours of birth (adjusted OR 0.74, 95% CI 0.61 to 0.90; moderate certainty evidence; [Analysis 2.3](#)). Neonatal mortality within 24 hours dropped in the study from 11.6 to 9.7 deaths per 1000 births in hospitals receiving the QUARITE intervention, compared to an increase from 9.0 to 10.7 deaths per 1000 births in the comparison hospitals (overall unadjusted reduction of 3.6 deaths per 1000 live births).

The intervention may have made little or no difference to inpatient neonatal mortality after 24 hours up to discharge. However, the 95% confidence interval indicates that the intervention may

reduce or increase inpatient neonatal mortality rate after 24 hours (adjusted OR 0.88, 95% CI 0.62 to 1.24; low certainty evidence; [Analysis 2.4](#)).

Maternal mortality

The QUARITE intervention probably reduced inpatient maternal mortality (adjusted OR 0.85, 95% CI 0.73 to 0.98; moderate certainty evidence; [Analysis 2.1](#)). The number of maternal deaths per 1000 women in the study went from 10.3 to 6.8 in hospitals receiving the QUARITE intervention, compared with a reduction from 8.1 to 7.1 per 1000 women in the comparison hospitals (overall reduction of 2.5 maternal deaths per 1000 women, 95% CI 0.9 to 4.2 deaths).

In subgroup analyses, the effect was larger in district hospitals (adjusted OR 0.65, 95% CI 0.55 to 0.77), but the intervention appeared to have had little or no effect in regional referral hospitals (adjusted OR 1.02, 95% CI 0.79 to 1.31). This may have been because of other interventions introduced during the study period in the regional referral hospitals in both the intervention and control sites ([Dumont 2013](#)). A further analysis showed that the effect of the intervention was larger for women delivering by caesarean section (adjusted OR 0.71, 95% CI 0.58 to 0.82) compared to women delivering vaginally (adjusted OR 0.87, 95% CI 0.69 to 1.11). The authors suggested this was because intrapartum caesarean delivery is associated with two- to six-fold higher risks of hospital-based maternal and neonatal mortality compared to spontaneous vaginal delivery, related to delays in seeking and receiving care, and so there is a greater potential to reduce mortality.

Secondary outcomes

Quality of care

The QUARITE intervention probably increases the proportion of women receiving high quality of care (OR 1.87, 95% CI 1.35 - 2.57; moderate-certainty evidence). For women with eclampsia or postpartum haemorrhage the intervention may increase the proportion of women with complications who receive high quality of care. However, the 95% confidence interval includes no effect (OR 1.67, 95% CI 0.96 - 2.91; low certainty evidence).

Quality of care was measured from medical records using criterion-based clinical audit (CBCA). The audit tool included 26 unweighted criteria that measured 5 dimensions of care: patient history, clinical examination, laboratory examinations, labour management (partograph), delivery care and postpartum monitoring. These criteria were applied for all women. An additional 7 items were scored only for women with severe pre-eclampsia/eclampsia and another 7 items only for women with postpartum haemorrhage. Each criterion was given one point and the overall score was calculated as a proportion of the relevant denominator (26 for women without severe complications, 33 for women with severe pre-eclampsia / eclampsia or postpartum haemorrhage). "Good quality care" was defined as a score of 70% or higher. Further analyses of the CBCA tool showed that low scores (less than 70%) predict perinatal mortality, which indicates construct validity ([Pirkle 2012](#)). Therefore we chose to present this binary outcome.

DISCUSSION

Summary of main results

This review only identified two cluster-randomised trials examining the effectiveness of death audit and review for reduction of maternal and perinatal mortality. There were many other studies of death audits and reviews, but none met the methodological criteria for inclusion in this review; almost all were uncontrolled before-and-after studies.

Both studies included death audits as the central component of a complex intervention, and these interventions were sufficiently different that we considered meta-analysis not to be useful. Furthermore, the settings were very different: QUARITE was conducted in countries with very high levels of maternal and perinatal mortality, whereas OPERA was conducted in France, which has very low levels of maternal and perinatal mortality. This may explain why there was no reduction in overall mortality in the OPERA trial (low certainty evidence), since it was very low in both control and intervention groups. However, the intervention probably reduced perinatal morbidity related to suboptimal care (moderate certainty evidence). The QUARITE intervention probably reduced inpatient maternal mortality, especially in women delivering by caesarean section, and in district hospitals (moderate certainty evidence), and probably also increased the proportion of women receiving high quality of care (moderate certainty evidence).

Overall completeness and applicability of evidence

Evidence from the QUARITE trial is likely applicable to maternal death audit and review in other low-income countries, as the intervention is well described in the protocol and could be replicated elsewhere. Senegal and Mali are among the world's poorest countries, so if it was possible to achieve a reduction in maternal mortality there within three years of intervention, it should be possible in most similar countries (although some contextual differences may affect the impact – for example the financial and legal). However, it is hard to assess the applicability of this evidence in middle- and high-income countries, where differences in the settings could modify the effect of the intervention.

The lack of impact on perinatal mortality in the OPERA trial can be explained by the already very low perinatal mortality rates in France. However, it did show that even in this relatively well-resourced setting, quality of perinatal care can be improved. This evidence may be applicable to other high-income countries with similar health system arrangements. We did not find any evidence on the effectiveness of perinatal death audit in low- and middle-income countries. Although the QUARITE intervention may have made little or no difference to inpatient stillbirth rates or neonatal mortality rates after 24 hours, the intervention involved audit and review of only maternal deaths (not perinatal deaths). Therefore, one cannot draw any conclusions from the QUARITE trial about the effectiveness of reviewing perinatal, neonatal or child deaths. Furthermore, the review process for perinatal and child deaths is different to that for maternal deaths.

Both of the included studies evaluated complex interventions, including death reviews as the main component and also training. The effects shown may be partly due to these training

components of the interventions. Another review has shown that continuing education meetings and workshops in general lead to a mean improvement in compliance with desired practice of 6% (interquartile range 1.8% to 15.9%) (Forsetlund 2009). However, there are almost no high-quality randomised controlled trials evaluating the impact on maternal or perinatal mortality of continuing education or training courses in emergency obstetric care (Ameh 2019). While it is impossible to ascertain separately the effects of the training and death review components of the intervention in the included studies, the effect observed in West Africa could be considered to be larger than might have been expected from training alone. The training component may also have indirect positive effects – for example, training may facilitate the implementation of audit and feedback, which may be resisted by clinicians in settings where this is a novel practice. However, in order to confirm these ideas, it would be necessary to conduct a randomised controlled trial comparing training alone, versus training plus death reviews.

There are important gaps in the evidence. We found no studies of death audits or reviews in isolation, or of late neonatal or child deaths. We also found no studies of death audit or review outside of hospitals (e.g. investigating deaths in the community). In addition, we identified no eligible studies conducted in middle-income countries or studies measuring cost-effectiveness. Neither of the included studies reported measures of maternal severe morbidity or cost per death averted. Because of the very limited number of included studies, we were unable to conduct any of the subgroup analyses that we had planned.

It is likely that few comparative studies have been conducted because showing an impact on maternal mortality (or even perinatal mortality) requires a large number of hospitals and women to take part, which is difficult and expensive to run. It can also be challenging to find study sites where no other interventions are being implemented that could also have an impact on maternal and perinatal mortality. In addition, many funders assume that research on MPDR is not a priority because in many countries there already a government policy that MPDR should be conducted, and it is already recommended by WHO (WHO 2013; WHO 2016). This makes it difficult to find control areas where MPDR is not being conducted in some form. A survey in 2015 found that 56/62 (90%) of countries had a policy on maternal death review (Bandali 2016). Therefore we are in the process of conducting a systematic review of uncontrolled before-and-after studies.

Quality of the evidence

The certainty of the evidence ranged from very low to moderate for the outcomes considered in this review. For both studies, there were issues with blinding because it is not possible to blind hospitals to whether or not they received the intervention. However, it was possible to blind those analysing the outcomes and both trials did this. Although we assessed the QUARITE study to be at low risk of bias overall, the certainty of evidence for the second comparison was downgraded due to indirectness. This was because the results from the QUARITE study included in this comparison are from a limited range of settings with very high maternal and perinatal mortality rates. The effects of this complex intervention may be modified by setting, or may work differently in a different setting. The OPERA trial had only one other shortcoming, namely that it did not use intention-to-treat analysis. In a future update, we may attempt to apply an 'intention to treat' analysis

to these data. Some of the results had wide CIs and it is therefore likely that further research would have an important impact on our confidence in the estimates of effect and may change the estimates, so the overall certainty of evidence was downgraded for this reason.

Potential biases in the review process

Our search strategy was designed to maximise sensitivity (detecting relevant research) at the expense of specificity (excluding irrelevant research). It is possible that some studies may have been missed if they were in the grey literature but we did consult widely with experts and all reports we found in the grey literature did not meet our methodological inclusion criteria. We conducted this review according to Cochrane standards.

Agreements and disagreements with other studies or reviews

Several other reviews have studied the effect of audit and feedback. One Cochrane Review found that audit and feedback was associated with a mean 4.3% improvement in health professionals' compliance with guidelines (Ivers 2012). The effect is small overall, but it appeared to be greater when baseline performance was low, when feedback came from a supervisor or colleague and was provided more than once, and when it included both explicit targets and an action plan. These circumstances were all present in the QUARITE trial, which showed a similar level of improvement in the quality of care measure (5.2%). In the OPERA trial, baseline performance was already good so there was probably little room for improvement. Interventions specifically tailored to overcome identified barriers to change led on average to an absolute 9.5% improvement (OR 1.52, 95% CI 1.27 to 1.82) in desired professional practice (Baker 2010). Death review should also incorporate a discussion of barriers to change, which is taken into consideration when making and implementing recommendations. One frequent recommendation is training of health professionals, and both studies in our review included an element of training on guidelines and best practice. Another Cochrane Review has shown that in-service training can improve performance of appropriate neonatal resuscitation, although impact on mortality was inconclusive (Opiyo 2015). Pattinson 2005 specifically aimed to evaluate the effectiveness of critical incident audit and feedback on maternal and perinatal mortality and morbidity, but found no trials meeting the inclusion criteria – this concurs with our search which only found two studies, both published after the last update for this review. The same author led a non-Cochrane review of perinatal mortality audits in low- and middle-income countries, which found that there was a mean reduction in perinatal mortality of 30% (95% CI 21% to 38%), but this is based on uncontrolled before-and-after studies which did not meet the inclusion criteria for this review (Pattinson 2009).

AUTHORS' CONCLUSIONS

Implications for practice

Maternal, perinatal and child mortality are a priority globally, especially in low-income countries, where much progress is needed to reach the United Nations' Sustainable Development Goals by 2030 (UN 2017). This review provides evidence that a complex intervention including maternal death audit and review, as well as development of local leadership and training, led to a 35% reduction in inpatient maternal mortality in district hospitals of

low-income countries, and probably slightly improved quality of care. There is also some evidence to support a complex intervention including perinatal death audit and review as well as training: the only cluster-randomised trial was conducted in a high-income country where mortality was already very low, but it still showed that the intervention probably improved quality of care, as measured by perinatal morbidity related to suboptimal care. However, there is currently no high-quality evidence on the effects of paediatric death audit and review as neither of the included studies investigated this. Neither did we find any high-quality evidence on the impact of community-based death reviews, or any evaluation of cost-effectiveness. Although other reviews have shown the general impacts of interventions such as audit and feedback and educational meetings (Ivers 2012; Forsetlund 2009), implementation of paediatric death audit and review and community-based death reviews should probably be done in the context of rigorous evaluation to allow evidence on their impacts to be collected.

The World Health Organization recommends that maternal and perinatal death reviews should be conducted in all hospitals globally. However, conducting death reviews in isolation may not be sufficient to achieve the reductions in mortality observed in the QUARITE trial. Evidence from this review suggests that maternal death audit and review may need to be implemented as part of an intervention package which also includes training of a leading doctor and midwife in each hospital (not only on emergency obstetric care but also on social barriers, health rights and adult education methods), annual recertification and quarterly outreach visits by external facilitators to provide supervision and mentorship. The same may also apply to perinatal and child death reviews. No undesirable effects of the intervention were documented in the trials we included. In the included studies, the intervention was clearly 'no-blame' and confidential, which avoided the potential undesirable effect of health workers feeling blamed and then disengaging from the process. Implementers should consider this aspect when planning and implementing maternal and perinatal death reviews.

Implications for research

More evidence is needed from randomised trials, particularly about the effectiveness of perinatal death review in low-income settings, including perinatal mortality. This research is needed as it may not be possible to generalise results from maternal death reviews to perinatal deaths, because these are different in some important aspects. For example, there are far fewer maternal deaths than perinatal deaths and, therefore, it may not be possible to audit all perinatal deaths in the same way (even in France in the OPERA study, only a sample of perinatal deaths was discussed in meetings). Furthermore, perinatal death audits would require more multidisciplinary collaboration involving not only obstetricians and midwives but also paediatricians and paediatric nurses. Death audit and review also has the potential to reduce mortality in other groups such as children, and deaths occurring

outside of health facilities – none of these have been evaluated in a randomised controlled trial. Rigorous evaluation of cost-effectiveness is required because the numbers of deaths, and costs of investigation in the community, are likely to be much greater than for hospital-based audit and review. Future trials should consider measuring some of the outcomes for which we found no evidence (such as child mortality, cost effectiveness and maternal morbidity). Rigorous research on this topic in low-income countries is difficult because of the lack of comprehensive birth and death registration systems, so study teams would need to establish their own systems, in order to measure outcomes accurately.

A number of questions remain about the implementation of death audits and reviews in different contexts. Although it is possible to have reasonable confidence that a complex intervention including training and facility-based maternal death reviews is more likely to lead to a significant reduction in inpatient maternal mortality than usual care in low-income countries, further well-planned studies are required to determine how the death audits and reviews approach should be designed to maximise effectiveness in different contexts, and how the approach compares to other quality improvement strategies (such as training alone).

Alongside research on effectiveness we also need operational research on the most cost-effective ways of implementing maternal and perinatal death review in low- and middle-income countries. The process of conducting reviews is time-consuming and has important opportunity costs (such as taking front-line staff away from their duties). It may not be necessary to audit and review all perinatal deaths in order to achieve important reductions in mortality and improvements in quality of care, particularly in low-income settings with very high mortality rates. Strategies for maximising impact and minimising costs (financial and time) need to be developed and evaluated.

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CHARACTERISTICS OF STUDIES

Characteristics of included studies [ordered by study ID]

Dumont 2013

Methods	Cluster-randomised trial QUARITE (QUALity of care, Risk management, and TEchnology) trial
Participants	Mothers who delivered in 46 first-level and second-level public referral hospitals in Mali and Senegal, with > 800 deliveries per year that had a functional operating room and had not previously conducted maternal death reviews.

Death audits and reviews for reducing maternal, perinatal and child mortality (Review)

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Dumont 2013 (Continued)

Interventions	<p>Intervention: a complex intervention to develop local leadership and empower obstetric teams.</p> <p>6-day training for 1 doctor and 1 midwife from each hospital: using the ALARM (Advances in Labour and Risk Management) international course. Consisted of 3 days of training in best practices in emergency obstetric care; 1 day of training in maternal death reviews; 1 day of awareness training related to economic, sociocultural and ethical barriers (including sexual and reproductive rights); and 1 day of training in adult education methods. The trainees were recertified on an annual basis. Following the initial training, a multidisciplinary audit committee was formed in each site and trained in the process of undertaking maternal death reviews. The audit cycle and onsite training were then launched with the support of external facilitators (certified instructors) during their quarterly educational outreach visits.</p> <p>Control group: no intervention. Administrators of these hospitals were informed that the 6-day training workshop would be provided at the end of the trial.</p>	
Outcomes	<p>Primary outcome</p> <ul style="list-style-type: none">• Change in inpatient maternal mortality rate. <p>Secondary outcomes</p> <ul style="list-style-type: none">• Change in inpatient perinatal mortality rate (divided into stillbirths, neonatal deaths before 24 hours and neonatal deaths after 24 hours but prior to discharge).• Resource availability, quantified by the hospital complexity index.• Medical practice for emergency obstetric care, assessed through the following essential obstetric interventions, considered effective in reducing maternal and perinatal mortality: assisted delivery (forceps and vacuum extraction), caesarean section, transfusion and hysterectomy, or transfer to another, more specialised health facility.• Proportion of women receiving good quality of care (defined as a score of 70% or more on a criterion-based clinical audit).	
Notes	Funded by Canadian Institutes of Health Research.	
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	The participating hospitals were in 6 strata corresponding to the combination of 2 countries (Mali and Senegal) and 3 hospital types: hospitals in the capital, regional hospitals and district hospitals outside the capital. Investigators attempted to ensure optimal balance between the hospitals assigned to the intervention and the control groups in terms of their number and size (number of deliveries per year). Therefore, within each stratum, they first ranked the hospitals with respect to size, and then used blocked randomisation, with each block of size 2 containing 2 hospitals with adjacent ranks, i.e. of similar size.
Allocation concealment (selection bias)	Low risk	All participating hospitals were randomised simultaneously, after their list was provided, which eliminated any risk of allocation bias.
Blinding of participants and personnel (performance bias) All outcomes	High risk	It was not possible to blind participants and personnel.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	The data collection and the implementation of the intervention were undertaken by different and independent organisations in each country. The organisations were not blinded with respect to randomisation but they were not involved in the assessment of the outcome. The statisticians assessing outcome

Dumont 2013 (Continued)

		were blinded to randomisation of each group (Dumont, personal communication).
Incomplete outcome data (attrition bias) All outcomes	Low risk	No hospitals were lost to follow-up. To include all eligible women in the intention-to-treat analyses, missing data for individual characteristics were imputed based on their distributions in the study population. In sensitivity analysis, women who died before labour were excluded because they usually sought care only after developing severe complications at home.
Selective reporting (reporting bias)	Low risk	Protocol was available. Predefined outcomes measurements were reported.
Contamination	Low risk	Randomisation by hospital, and hospitals were in different areas.

Dupont 2017

Methods	Cluster randomised trial OPERA trial
Participants	Births at 95 maternity units in France; 50% had 500–1499 deliveries per year, 40% had > 1499 deliveries per year, and 10% had < 500 deliveries per year.
Interventions	<p>Intervention: single limited complex intervention to promote national guidelines on antenatal care and peripartum practices; perinatal MMCs in presence or absence of a CP.</p> <p>The first component was an outreach visit to brief obstetricians, midwives and anaesthetists on the national guidelines on morbidity/mortality case management. Precisely, the guidelines were about monitoring of normal pregnancy, management of intrauterine growth-restricted fetuses during labour, interpretation of fetal heart rate during labour and safe practice in instrumental vaginal delivery. During the visit, the co-ordinators discussed the scientific and medical validity of the guidelines with the medical staff to support their implementation.</p> <p>The second component was a series of MMCs dedicated to review a selected sample of perinatal morbidity and mortality cases with all staff members who managed them. The MMCs were held in the maternity units concerned and were led by the investigators, who invited all the staff to attend. MMC + CP sessions included analyses of the staff decision-making processes and explored the role of psychological factors in these processes. 3–4 MMCs were held in maternity units with > 1500 deliveries, 2–3 in units with 500–1500 deliveries, and 2 in units with < 500 deliveries per year.</p> <p>Control group: continued current care – no intervention.</p>
Outcomes	<p>Primary outcome</p> <ul style="list-style-type: none"> Rate of suboptimal care among cases of perinatal morbidity or mortality (stillbirths and neonatal). <p><i>[Optimal care was defined as the management of a morbidity or mortality case in perfect accordance with the guidelines. In each case, the quality of care was jointly examined by two reviewers who had to agree on classifying it according to a 7-point scale, where 7 = optimal, 6 = nearly optimal, 5 = satisfactory, 4 = nearly satisfactory, 3 = possibly suboptimal, 2 = certainly suboptimal, and 1 = not classifiable. Suboptimal care was defined as score 3 or 2 (i.e. not compliant with the guidelines).]</i></p> <p>Secondary outcomes</p> <ul style="list-style-type: none"> Rate of suboptimal care among morbidity cases. Rate of suboptimal care among mortality cases.

Death audits and reviews for reducing maternal, perinatal and child mortality (Review)

Dupont 2017 (Continued)

- Rate of avoidable morbidity cases.
- Rate of avoidable mortality cases.
- Incidence of morbidity (excluding intrauterine growth restriction).
- Incidence of mortality.

Notes	Maternal mortality and morbidity were not evaluated. Funded by the French Ministry of Health under its Clinical Research Hospital Program (contract no. 27-41).
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Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer-generated randomisation was stratified by network, institutional status (public university-related, public university-unrelated and private) and annual number of deliveries (< 500, 500–1499 and 1500). A second randomisation within the intervention group generated two subgroups assigned to have MMCs in presence or absence of a CP.
Allocation concealment (selection bias)	Low risk	The health units could not know in advance to which group they would be randomised (Dupont, personal communication).
Blinding of participants and personnel (performance bias) All outcomes	High risk	Impossible to blind participants and personnel. Personnel may behave differently if they are aware that they are receiving an intervention.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	The investigators who evaluated the outcome on suboptimal care were blinded as to whether the case was from an intervention or control site. The same investigators took part in mortality meetings at intervention sites but only a small number of cases were discussed and the outcome evaluation was conducted 1 year later so they could not remember the cases (Dupont, personal communication).
Incomplete outcome data (attrition bias) All outcomes	Low risk	2 private units withdrew (prior to randomisation). All 95 units completed the trial.
Selective reporting (reporting bias)	Low risk	There were no outcomes in the protocol which were not reported in the publication (Dupont, personal communication).
Other bias	High risk	Six units randomised to the intervention group did not implement the intervention and were transferred to the control group and analysed per protocol instead of intention to treat. It is impossible to know whether the transferred units are completely similar to those that retained their original randomisation.
Contamination	Low risk	Although it is possible that an individual health worker could work in both an intervention and a control hospital, it is very unlikely that this would apply to the whole team.

CP: clinical psychologist; MMC: morbidity/mortality conference; OR: odds ratio.

Characteristics of excluded studies [ordered by study ID]

Death audits and reviews for reducing maternal, perinatal and child mortality (Review)

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Study	Reason for exclusion
Allanson 2015	Study design
Biswas 2014	Study design
Biswas 2018	Outcome
Bugalho 1993	Study design
Cahyanti 2018	Outcome
Crofts 2015	Intervention
Dooley 2014	Study design
Dumont 2005	Study design
Dumont 2006	Study design
Eskes 2014	Study design
Gaunt 2010	Study design
Gebrehiwot 2014	Study design
Incekoy Girgin 2018	Intervention
Kaharuza 2012	Study design
Kongnyuy 2008	Study design
Kwast 1995	Study design
Main 2018	Study design
Maresh 1998	Study design
Mbaruku 1995	Study design
Min 2017	Intervention
Moodley 2014	Study design
Nakibuuka 2012	Study design
Okonofua 2017	Study design
Papiernik 2005	Study design
Patrick 2007	Study design
Pattinson 1995	Study design
Pattinson 2006	Study design
Pattinson 2009	Study design

Study	Reason for exclusion
Pattinson 2011	Study design
Persson 2013	Intervention did not include death audit or review.
Ravichandran 2014	Study design
Reiffenstuhl 1982	Intervention did not include death audit or review.
Santos 2006	Study design
Serbanescu 2017	Study design
Shrestha 2006	Study design
Srofenyoh 2016	Intervention
Stratulat 2012	Study design
Supratikto 2002	Outcome
Tette 2016	Intervention did not include death audit and review.
Thomas 1985	Study design
van den Akker 2011	Study design
van Roosmalen 1989	Study design
Ward 1995	Study design
Wilkinson 1991	Study design
Wilkinson 1997	Study design
Willcox 2018	Study design

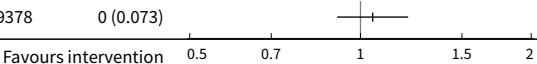
DATA AND ANALYSES

Comparison 1. Perinatal death review and audit as part of an intervention package including an educational outreach visit and morbidity/mortality conferences compared with no intervention

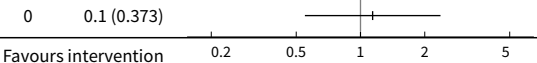
Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Perinatal mortality rate	1		Odds Ratio (Fixed, 95% CI)	Subtotals only
2 Perinatal mortality related to suboptimal care	1		Odds Ratio (Fixed, 95% CI)	Subtotals only

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
3 Perinatal morbidity related to suboptimal care	1		Odds Ratio (Fixed, 95% CI)	Subtotals only

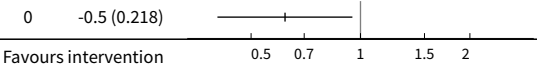
Analysis 1.1. Comparison 1 Perinatal death review and audit as part of an intervention package including an educational outreach visit and morbidity/mortality conferences compared with no intervention, Outcome 1 Perinatal mortality rate.

Study or subgroup	Intervention N	Control N	log[Odds Ratio] (SE)	Odds Ratio IV, Fixed, 95% CI	Weight	Odds Ratio IV, Fixed, 95% CI
Dupont 2017	95975	69378	0 (0.073)		0%	1.05[0.91,1.21]
Favours intervention				0.5	0.7	1
				1.5	2	Favours control

Analysis 1.2. Comparison 1 Perinatal death review and audit as part of an intervention package including an educational outreach visit and morbidity/mortality conferences compared with no intervention, Outcome 2 Perinatal mortality related to suboptimal care.

Study or subgroup	Intervention N	Control N	log[Odds Ratio] (SE)	Odds Ratio IV, Fixed, 95% CI	Weight	Odds Ratio IV, Fixed, 95% CI
Dupont 2017	0	0	0.1 (0.373)		0%	1.14[0.55,2.37]
Favours intervention				0.2	0.5	1
				2	5	Favours control

Analysis 1.3. Comparison 1 Perinatal death review and audit as part of an intervention package including an educational outreach visit and morbidity/mortality conferences compared with no intervention, Outcome 3 Perinatal morbidity related to suboptimal care.


Study or subgroup	Intervention N	Control N	log[Odds Ratio] (SE)	Odds Ratio IV, Fixed, 95% CI	Weight	Odds Ratio IV, Fixed, 95% CI
Dupont 2017	0	0	-0.5 (0.218)		0%	0.62[0.4,0.95]
Favours intervention				0.5	0.7	1
				1.5	2	Favours control

Comparison 2. Maternal death review and audit as part of an intervention package including the ALARM course and training audit committees compared with no intervention

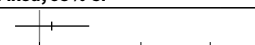
Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
1 Change in inpatient maternal mortality rate	1		Odds Ratio (Fixed, 95% CI)	Subtotals only

Outcome or subgroup title	No. of studies	No. of participants	Statistical method	Effect size
2 Change in inpatient stillbirth rate	1		Odds Ratio (Fixed, 95% CI)	Subtotals only
3 Change in inpatient neonatal mortality rate before 24 hours	1		Odds Ratio (Fixed, 95% CI)	Subtotals only
4 Change in inpatient neonatal mortality rate after 24 hours	1		Odds Ratio (Fixed, 95% CI)	Subtotals only

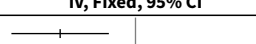
Analysis 2.1. Comparison 2 Maternal death review and audit as part of an intervention package including the ALARM course and training audit committees compared with no intervention, Outcome 1 Change in inpatient maternal mortality rate.

Study or subgroup	Intervention N	Control N	log[Odds Ratio] (SE)	Odds Ratio IV, Fixed, 95% CI	Weight	Odds Ratio IV, Fixed, 95% CI
Dumont 2013	0	0	-0.2 (0.075)		0%	0.85[0.73,0.98]
				Favours intervention 0.5 0.7 1 1.5 2 Favours control		

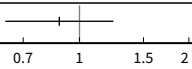
Analysis 2.2. Comparison 2 Maternal death review and audit as part of an intervention package including the ALARM course and training audit committees compared with no intervention, Outcome 2 Change in inpatient stillbirth rate.

Study or subgroup	Intervention N	Control N	log[Odds Ratio] (SE)	Odds Ratio IV, Fixed, 95% CI	Weight	Odds Ratio IV, Fixed, 95% CI
Dumont 2013	0	0	0 (0.075)		0%	1.05[0.91,1.22]
				Favours intervention 0.5 0.7 1 1.5 2 Favours control		

Analysis 2.3. Comparison 2 Maternal death review and audit as part of an intervention package including the ALARM course and training audit committees compared with no intervention, Outcome 3 Change in inpatient neonatal mortality rate before 24 hours.

Study or subgroup	Intervention N	Control N	log[Odds Ratio] (SE)	Odds Ratio IV, Fixed, 95% CI	Weight	Odds Ratio IV, Fixed, 95% CI
Dumont 2013	0	0	-0.3 (0.099)		0%	0.74[0.61,0.9]
				Favours intervention 0.5 0.7 1 1.5 2 Favours control		

Analysis 2.4. Comparison 2 Maternal death review and audit as part of an intervention package including the ALARM course and training audit committees compared with no intervention, Outcome 4 Change in inpatient neonatal mortality rate after 24 hours.

Study or subgroup	Inter- vention N	Control N	log[Odds Ratio] (SE)	Odds Ratio IV, Fixed, 95% CI	Weight	Odds Ratio IV, Fixed, 95% CI
Dumont 2013	0	0	-0.1 (0.175)		0%	0.88[0.62,1.24]

ADDITIONAL TABLES

Table 1. Number of events and participants in the QUARITE trial (Dumont 2013)

Outcome	Comparison group					Intervention group				
	Baseline		Year 4			Baseline		Year 4		
	Num-ber of event	Num-ber of par-tici-pants	Num-ber of event	Num-ber of par-tici-pants	Rate	Num-ber of event	Num-ber of par-tici-pants	Num-ber of event	Num-ber of par-tici-pants	Rate
Inpatient stillbirth rate (per 1000 total births)	3441	39992	86.0	27051	324 83.2 (decreased by 2.8 stillbirths from baseline to year 4)	3883	41368	93.9	23850	426 84.0 (decreased by 9.9 stillbirths from baseline to year 4)
Inpatient neonatal mortality rate - before 24 hours (per 1000 live births)	332	36551	9.0	50547	054 10.7 (increased by 1.7 neonatal deaths from baseline to year 4)	434	37485	11.6	44646	188 9.7 (decreased by 1.9 neonatal deaths from baseline to year 4)
Inpatient neonatal mortality rate - after 24 hours (per 1000 live births)	99	36551	2.7	9947	054 2.1 (decreased by 0.6 neonatal deaths from baseline to year 4)	232	37485	6.2	18546	188 4.0 (decreased by 2.2 neonatal deaths from baseline to year 4)
Inpatient maternal mortality rate (per 100,000 pregnant women)	337	41655	809	38153	581 711 (decreased by 98 maternal deaths from baseline to year 4)	445	43269	102	85656	2662 676 (decreased by 352 maternal deaths from baseline to year 4)
Quality of care: Proportion of women receiving high quality care (per 1000 pregnant women)	-	-	101	339	298	-	-	141	319	442
Quality of care: Proportion of women with eclampsia or postpartum haemorrhage receiving high quality care (per 1000 pregnant women with complications)	-	-	43	114	377	-	-	48	95	505

Table 2. Uncontrolled before-and-after studies of death reviews

Study ID	Intervention description	Outcomes assessed	Country	Setting/s where implemented
Allanson 2015	Perinatal Problem Identification Programme	Hospital-based perinatal mortality	South Africa	163 health facilities (29 community health centres, 105 district hospitals, 4 national central hospitals, 22 regional hospitals and 3 provincial tertiary hospitals)
Bugalho 1993	In-facility case review of perinatal deaths and maternal deaths	Hospital-based perinatal mortality	Mozambique	1 national referral hospital
Dumont 2006	In-facility case review of maternal deaths	Hospital-based maternal mortality	Senegal	1 district hospital
Eskes 2014	In-facility case review of term perinatal deaths	Perinatal mortality	Netherlands	90 Dutch hospitals with obstetric/ paediatric departments
Gaunt 2010	In-facility case review of perinatal deaths	Hospital-based perinatal mortality	South Africa	1 district hospital
Kongnyuy 2008	In-facility case review of maternal deaths and criterion-based clinical audit	Hospital-based maternal mortality	Malawi	13 hospitals and 60 health centres
Mbaruku 1995	Retrospective case review of in-facility maternal deaths 1984–1986, followed by prospective case reviews 1987–1991	Hospital-based maternal mortality	Tanzania	1 regional referral hospital
Moodley 2014	Confidential enquiry into maternal deaths	Maternal mortality	South Africa	National level
Mussell et al (unpublished - reference in Pattinson 2009)	Perinatal Problem Identification Programme	Hospital-based perinatal mortality	Bangladesh	1 hospital
Nakibuuka 2012	In-facility case review of perinatal deaths	Hospital-based perinatal mortality	Uganda	1 private referral hospital
Okonofua 2017	In-facility case review of maternal deaths	Hospital-based maternal mortality	Nigeria	3 referral hospitals in Lagos
Papiernik 2000, Papiernik 2005	In-facility case review of perinatal deaths	Perinatal mortality	France	17 private maternity units, 5 secondary care hospitals, 4 referral hospitals
Patrick 2007	In-facility case review of perinatal deaths: retrospective 1995–1996 and prospective 1996–2000	Hospital-based perinatal mortality	South Africa	1 referral hospital
Pattinson 1995	Perinatal Problem Identification Programme	Perinatal mortality	South Africa	1 hospital

Death audits and reviews for reducing maternal, perinatal and child mortality (Review)

32

Table 2. Uncontrolled before-and-after studies of death reviews *(Continued)*

Pattinson 2006	In-facility case review of maternal deaths and severe acute maternal morbidity	Hospital-based maternal mortality	South Africa	2 district and 2 academic hospitals
Pattinson 2011	Audit of perinatal deaths	Facility-based perinatal mortality	South Africa	6 midwife obstetric units, 24 district hospitals, 5 regional hospitals
Shrestha 2006	Audit of perinatal deaths	Hospital-based perinatal mortality	Nepal	1 tertiary hospital
Stratulat 2012	Confidential enquiry into perinatal deaths	Perinatal deaths among term newborns	Moldova	National level
Thomas 1985	Confidential enquiry into perinatal deaths	Perinatal mortality	Wales	1 county
van den Akker 2011	In-facility case review of maternal deaths and severe acute maternal morbidity	Maternal mortality	Malawi	1 district hospital and 28 smaller health facilities
van Roosmalen 1989	In-facility case review of perinatal deaths and retrospective audit of stillbirths	Hospital-based perinatal mortality	Tanzania	1 district hospital
Ward 1995	In-facility case review and audit of perinatal deaths	Hospital-based perinatal mortality	South Africa	1 referral hospital
Wilkinson 1991	In-facility case review of perinatal deaths	Facility-based perinatal mortality	South Africa	1 district hospital + surrounding clinics
Wilkinson 1997	Audit of perinatal deaths	Facility-based perinatal mortality	South Africa	1 district
Willcox 2018	Community-based confidential enquiry into child deaths	Under-5 mortality	Uganda and Mali	5 subdistricts/subcounties in each country

Table 3. Numbers of events and participants in the OPERA trial (Dupont 2017)

Outcome	Comparison group		Intervention group	
	Number of events	Number of participants	Number of events	Number of participants
Perinatal mortality rate (overall)	448	95975	340	69378
% of perinatal deaths related to suboptimal care	37	435	29	324
% of perinatal morbidity cases related to suboptimal care	116	1007	48	633

APPENDICES

Appendix 1. Search strategies

MEDLINE, Ovid

Death audits and reviews for reducing maternal, perinatal and child mortality (Review)

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# ▼	Searches	Results
1	(child mortality/ or fetal mortality/ or infant mortality/ or maternal mortality/ or perinatal mortality/) and (clinical audit/ or medical audit/)	310
2	(Pregnant Women/ or exp Child/) and ("cause of death"/ or Mortality/) and (clinical audit/ or medical audit/)	54
3	(Pregnancy Complications/mo or Stillbirth/ or Sudden Infant Death/) and (clinical audit/ or medical audit/)	73
4	((maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r) adj3 (mortality or death?)).ti,ab. and (clinical audit/ or medical audit/)	395
5	(stillbirth? or sudden infant death? or sids or cot death? or crib death?).ti,ab. and (clinical audit/ or medical audit/)	59
6	(child mortality/ or fetal mortality/ or infant mortality/ or maternal mortality/ or perinatal mortality/) and (review* or audit* or meeting? or enquir* or inquir*).ti,ab.	4241
7	(Pregnant Women/ or exp Child/) and ("cause of death"/ or Mortality/) and (review* or audit* or meeting? or enquir* or inquir*).ti,ab.	1612
8	(Pregnancy Complications/mo or Stillbirth/ or Sudden Infant Death/) and (review* or audit* or meeting? or enquir* or inquir*).ti,ab.	961
9	((death? or mortality) adj3 (review* or audit* or meeting? or enquir* or inquir*)) and (maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r).ti,ab.	2039
10	((stillbirth? or sudden infant death? or sids or cot death? or crib death?) adj5 (review* or audit* or meeting? or enquir* or inquir*).ti,ab.	333
11	((confidential enquir* or confidential inquir*) and ((maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r) adj3 (mortality or death?))).ti,ab.	319
12	((confidential enquir* or confidential inquir*) and (stillbirth? or sudden infant death? or sids or cot death? or crib death?).ti,ab.	57
13	(cemach or cmace or cemd or cmde).ti,ab.	62
14	(saving mothers lives or making pregnancy safer or making childbirth safer).ti,ab.	43
15	((verbal autops* or social autops*) adj5 (maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r)).ti,ab.	114
16	((near miss* or significant event* or critical event* or critical incident?) and (maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r) and (review* or audit* or meeting? or enquir* or inquir*).ti,ab.	360
17	1 or 2 or 3 or 4 or 5 or 6 or 7 or 8 or 9 or 10 or 11 or 12 or 13 or 14 or 15 or 16	8057

(Continued)

18	exp Animals/ not humans/	4,439,627
19	17 not 18	8026
20	(review or meta analysis or news or comment or editorial).pt. or cochrane database of systematic reviews.jn. or comment on.cm. or (systematic review or literature review).ti.	3,599,131
21	19 not 20	5682
22	randomized controlled trial.pt.	456,938
23	controlled clinical trial.pt.	92,283
24	randomized.ab.	406,694
25	placebo.ab.	187,539
26	drug therapy.fs.	2,004,674
27	randomly.ab.	287,478
28	trial.ab.	422,400
29	groups.ab.	1,777,916
30	22 or 23 or 24 or 25 or 26 or 27 or 28 or 29	4,169,464
31	21 and 30	924
32	multicenter study.pt.	230,766
33	pragmatic clinical trial.pt.	709
34	(randomis* or randomiz*).ti,ab.	533,911
35	(trial or multicenter or multi center or multicentre or multi centre).ti.	212,507
36	(intervention? or effect? or impact? or controlled or control group? or (before adj5 after) or (pre adj5 post) or ((pretest or pre test) and (posttest or post test)) or quasiexperiment* or quasi experiment* or evaluat* or time series or time point? or repeated measur*).ti,ab.	8,366,295
37	32 or 33 or 34 or 35 or 36	8,561,470
38	37 not 30	6,215,306
39	21 and 38	1760
40	exp "Costs and Cost Analysis"/	213,468
41	economics/ or exp economics, hospital/ or exp economics, medical/	62,797
42	"Value of Life"/	5583
43	quality-adjusted life years/	9946
44	Decision Trees/	10,131

(Continued)

45	economic evaluation*.ti,ab.	9722
46	(Cost* adj2 (Effective* or analysis* or Utility* or Benefit* or Minimi*)).ti,ab.	138,444
47	(pharmacoeconomic* or pharmaco-economic*).ti,ab.	3720
48	economic*.ti.	40,888
49	("Value of life" or "quality adjusted life year*" or qaly* or qald* or qale* or "disability adjusted life year*" or daly).ti,ab.	14,304
50	(sf6 or short form 6 or shortform6 or euroqol or euro quality of life or eq5d or eq-5d).ti,ab.	9323
51	(hye or health* year equivalent*).ti,ab.	53
52	(health utilit* or disutilit*).ti,ab.	1971
53	"willingness to pay".ti,ab.	4119
54	standard gamble.ti,ab.	774
55	(time trade off or time tradeoff or tto).ti,ab.	1588
56	(vas or visual analog*).ti,ab.	64,886
57	((economic adj2 model*) or markov or monte carlo method).ti,ab.	23,587
58	(decision* adj (tree* or model* or analysis)).ti,ab.	12,137
59	(resource* adj (use* or utilisation)).ti,ab.	9248
60	((healthcare or health care or direct service or hospital or drug*) adj cost*).ti,ab.	30,459
61	40 or 41 or 42 or 43 or 44 or 45 or 46 or 47 or 48 or 49 or 50 or 51 or 52 or 53 or 54 or 55 or 56 or 57 or 58 or 59 or 60	495,867
62	61 not (30 or 37)	239,344
63	21 and 62	100
64	31 or 39 or 63	2784

Embase, Ovid

# ▼	Searches	Results
1	(childhood mortality/ or fetus mortality/ or infant mortality/ or maternal mortality/ or exp perinatal mortality/ or prenatal mortality/) and medical audit/	682
2	(pregnant woman/ or child/ or exp infant/ or preschool child/ or toddler/) and ("cause of death"/ or Mortality/) and medical audit/	449

(Continued)

3	((*pregnancy complication/ and ("cause of death"/ or *mortality/)) or *child death/ or *newborn death/ or *sudden infant death syndrome/ or exp *fetus death/) and medical audit/	90
4	((maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r) adj3 (mortality or death?)).ti,ab. and medical audit/	794
5	(stillbirth? or sudden infant death? or sids or cot death? or crib death?).ti,ab. and medical audit/	187
6	(*childhood mortality/ or *fetus mortality/ or *infant mortality/ or *maternal mortality/ or exp *perinatal mortality/ or *prenatal mortality/) and (review* or audit* or meeting? or enquir* or inquir*).ti,ab.	2838
7	(pregnant woman/ or child/ or exp infant/ or preschool child/ or toddler/) and ("cause of death"/ or *Mortality/) and (review* or audit* or meeting? or enquir* or inquir*).ti,ab.	1156
8	((*pregnancy complication/ and ("cause of death"/ or *mortality/)) or *child death/ or *newborn death/ or *sudden infant death syndrome/ or exp *fetus death/) and (review* or audit* or meeting? or enquir* or inquir*).ti,ab.	2084
9	((death? or mortality) adj3 (review* or audit* or meeting? or enquir* or inquir*)) and (maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r)).ti,ab.	2867
10	((stillbirth? or sudden infant death? or sids or cot death? or crib death?) adj5 (review* or audit* or meeting? or enquir* or inquir*).ti,ab.	496
11	((confidential enquir* or confidential inquir*) and ((maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r) adj3 (mortality or death?))).ti,ab.	551
12	((confidential enquir* or confidential inquir*) and (stillbirth? or sudden infant death? or sids or cot death? or crib death?)).ti,ab.	85
13	(cemach or cmace or cemd or cmde).ti,ab.	318
14	(saving mothers lives or making pregnancy safer or making childbirth safer).ti,ab.	71
15	((verbal autops* or social autops*) adj5 (maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r)).ti,ab.	138
16	((near miss* or significant event* or critical event* or critical incident?) and (maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r) and (review* or audit* or meeting? or enquir* or inquir*).ti,ab.	570
17	1 or 2 or 3 or 4 or 5 or 6 or 7 or 8 or 9 or 10 or 11 or 12 or 13 or 14 or 15 or 16	9225
18	(editorial or letter or note or "review").pt. or cochrane database of systematic reviews.jn. or (systematic review or literature review).ti.	4,703,611
19	17 not 18	7115
20	randomized controlled trial/	496,811

Death audits and reviews for reducing maternal, perinatal and child mortality (Review)

37

(Continued)

21	controlled clinical trial/	459,202
22	single blind procedure/ or double blind procedure/	177,789
23	crossover procedure/	55,009
24	random*.tw.	1,289,179
25	placebo*.tw.	272,054
26	((singl* or doubl*) adj (blind* or mask*)).tw.	211,276
27	(crossover or cross over or factorial* or latin square).tw.	127,797
28	(assign* or allocat* or volunteer*).tw.	677,956
29	20 or 21 or 22 or 23 or 24 or 25 or 26 or 27 or 28	2,128,969
30	19 and 29	498
31	clinical trial/	968,030
32	multicenter study/	181,820
33	(randomis* or randomiz*).ti,ab.	757,379
34	(trial or multicenter or multi center or multicentre or multi centre).ti.	297,401
35	(intervention? or effect? or impact? or controlled or control group? or (before adj5 after) or (pre adj5 post) or ((pretest or pre test) and (posttest or post test)) or quasiexperiment* or quasi experiment* or evaluat* or time series or time point? or repeated measur*).ti,ab.	10,710,433
36	31 or 32 or 33 or 34 or 35	11,251,708
37	36 not 29	9,568,756
38	19 and 37	2578
39	health economics/ or pharmacoeconomics/	43,371
40	exp health care costs/	260,766
41	exp economic evaluation/	271,749
42	quality adjusted life year/	20,849
43	economics/ or economic aspect/	333,969
44	economic evaluation*.ti,ab.	13,630
45	(Cost* adj2 (Effective* or analysis* or Utility* or Benefit* or Minimi*)).ti,ab.	188,360
46	(pharmacoeconomic* or pharmaco-economic*).ti,ab.	7843
47	economic*.ti.	51,747

(Continued)

48	("Value of life" or "quality adjusted life year*" or qaly* or qald* or qale* or "disability adjusted life year*" or daly).ti,ab.	23,054
49	(sf6 or short form 6 or shortform6 or euroqol or euro quality of life or eq5d or eq-5d).ti,ab.	16,536
50	(hye or health* year equivalent*).ti,ab.	107
51	(health utilit* or disutilit*).ti,ab.	3308
52	"willingness to pay".ti,ab.	6465
53	standard gamble.ti,ab.	994
54	(time trade off or time tradeoff or tto).ti,ab.	2290
55	(vas or visual analog*).ti,ab.	97,572
56	((economic adj2 model*) or markov or monte carlo method).ti,ab.	29,329
57	(decision* adj (tree* or model* or analysis)).ti,ab.	17,213
58	(resource* adj (use* or utilisation)).ti,ab.	14,022
59	((healthcare or health care or direct service or hospital or drug*) adj cost*).ti,ab.	46,977
60	39 or 40 or 41 or 42 or 43 or 44 or 45 or 46 or 47 or 48 or 49 or 50 or 51 or 52 or 53 or 54 or 55 or 56 or 57 or 58 or 59	978,150
61	60 not (29 or 36)	556,447
62	19 and 61	86
63	30 or 38 or 62	3162

Global Health, Ovid

# ▼	Searches	Results
1	(infant mortality/ or maternal mortality/ or neonatal mortality/ or perinatal mortality/ or stillbirths/) and (review* or audit* or meeting? or enquir* or inquir*).ti,ab.	1730
2	(children/ or preschool children/ or infants/) and ("cause of death"/ or Mortality/) and (review* or audit* or meeting? or enquir* or inquir*).ti,ab.	2698
3	((pregnancy complications/ and ("causes of death"/ or mortality/)) or sudden infant death syndrome/) and (review* or audit* or meeting? or enquir* or inquir*).ti,ab.	330
4	((death? or mortality) adj3 (review* or audit* or meeting? or enquir* or inquir*)) and (maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r).ti,ab.	669
5	((stillbirth? or sudden infant death? or sids or cot death? or crib death?) adj5 (review* or audit* or meeting? or enquir* or inquir*).ti,ab.	99

Death audits and reviews for reducing maternal, perinatal and child mortality (Review)

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(Continued)

6	((confidential enquir* or confidential inquir*) and ((maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r) adj3 (mortality or death?))).ti,ab.	87
7	((confidential enquir* or confidential inquir*) and (stillbirth? or sudden infant death? or sids or cot death? or crib death?)).ti,ab.	10
8	(cemach or cmace or cemd or cmde).ti,ab.	5
9	(saving mothers lives or making pregnancy safer or making childbirth safer).ti,ab.	16
10	((verbal autops* or social autops*) adj5 (maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r)).ti,ab.	92
11	((near miss* or significant event* or critical event* or critical incident?) and (maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r) and (review* or audit* or meeting? or enquir* or inquir*)).ti,ab.	96
12	1 or 2 or 3 or 4 or 5 or 6 or 7 or 8 or 9 or 10 or 11	4573
13	(random* or blind* or allocat* or assign* or trial* or placebo* or crossover* or cross-over*).ti,ab.	280,984
14	12 and 13	635
15	(multicenter or multi center or multicentre or multi centre).ti.	5213
16	(intervention? or effect? or impact? or controlled or control group? or (before adj5 after) or (pre adj5 post) or ((pretest or pre test) and (posttest or post test)) or quasiexperiment* or quasi experiment* or evaluat* or time series or time point? or repeated measur*).ti,ab.	1,302,529
17	15 or 16	1,304,574
18	17 not 13	1,108,174
19	12 and 18	1757
20	economics/ or economic evaluation/	8105
21	economic analysis/ or "cost analysis"/ or "cost benefit analysis"/ or "cost effectiveness analysis"/	12,451
22	(Cost* adj2 (Effective* or analysis* or Utility* or Benefit* or Minimi*)).ti,ab.	25,527
23	(pharmacoeconomic* or pharmaco-economic*).ti,ab.	348
24	economic*.ti.	11,263
25	economic evaluation*.ti,ab.	1620
26	("Value of life" or "quality adjusted life year*" or qaly* or qald* or qale* or "disability adjusted life year*" or daly).ti,ab.	3740
27	(sf6 or short form 6 or shortform6 or euroqol or euro quality of life or eq5d or eq-5d).ti,ab.	673

(Continued)

28	(hye or health* year equivalent*).ti,ab.	1
29	(health utilit* or disutilit*).ti,ab.	215
30	"willingness to pay".ti,ab.	1630
31	standard gamble.ti,ab.	34
32	(time trade off or time tradeoff or tto).ti,ab.	191
33	(vas or visual analog*).ti,ab.	3901
34	((economic adj2 model*) or markov or monte carlo method).ti,ab.	2861
35	(decision* adj (tree* or model* or analysis)).ti,ab.	1810
36	(resource* adj (use* or utilisation)).ti,ab.	1304
37	((healthcare or health care or direct service or hospital or drug*) adj cost*).ti,ab.	4359
38	20 or 21 or 22 or 23 or 24 or 25 or 26 or 27 or 28 or 29 or 30 or 31 or 32 or 33 or 34 or 35 or 36 or 37	56,861
39	38 not (14 or 19)	56,715
40	12 and 39	39
41	14 or 19 or 40	2431

Popline, K4Health

1	TI: (death OR deaths OR mortality) AND TI: (audit OR audits OR enquiry OR enquiries OR inquiry OR inquiries OR meeting OR meetings)	87
2	TI: death review OR mortality review OR death reviews OR mortality reviews	29
3	TI: (confidential enquiry OR confidential enquiries OR confidential inquiry OR confidential inquiries OR cemach OR cmace OR cemd OR cmde) AND (death OR deaths OR mortality)	16
4	TI: verbal autopsy OR verbal autopsies OR social autopsy OR social autopsies	29
5	TI: (near miss OR significant events OR critical events OR critical incidents OR significant event OR critical event OR critical incident) AND (audit OR audits OR enquiry OR enquiries OR inquiry OR inquiries OR meeting OR meetings)	0
6	1 or 2 or 3 or 4 or 5	161

Global Health Library, WHO

1	(maternal OR mother OR mothers OR maternity OR child OR children OR childhood OR infant OR infants OR pediatric OR paediatric OR fetal OR foetal OR perinatal OR pregnant OR pregnancy OR childbirth OR birth OR labor OR labour) AND (death OR deaths OR mortality) AND (audit OR audits OR enquiry OR enquiries OR inquiry OR inquiries OR meeting OR meetings)	299
2	(death review OR mortality review OR death reviews OR mortality reviews) AND (maternal OR mother OR mothers OR maternity OR child OR children OR childhood OR infant OR infants OR pediatric OR paediatric OR fetal OR foetal OR perinatal OR pregnant OR pregnancy OR childbirth OR birth OR labor OR labour)	83
3	(stillbirth OR stillbirths OR sudden infant death OR sudden infant deaths OR sids OR cot death OR cot deaths OR crib death OR crib deaths) AND (audit OR audits OR enquiry OR enquiries OR inquiry OR inquiries OR meeting OR meetings)	34
4	confidential enquiry OR confidential enquiries OR "confidential inquiry" OR "confidential inquiries" OR cemach OR cmace OR cemd OR cmde OR saving mothers lives OR making pregnancy safer OR making childbirth safer	0
5	(verbal autopsy OR verbal autopsies OR social autopsy OR social autopsies) AND (maternal OR mother OR mothers OR maternity OR child OR children OR childhood OR infant OR infants OR pediatric OR paediatric OR fetal OR foetal OR perinatal OR pregnant OR pregnancy OR childbirth OR birth OR labor OR labour)	22
6	(near miss OR significant events OR critical events OR critical incidents OR significant event OR critical event OR critical incident) AND (audit OR audits OR enquiry OR enquiries OR inquiry OR inquiries OR meeting OR meetings) AND (maternal OR mother OR mothers OR maternity OR child OR children OR childhood OR infant OR infants OR pediatric OR paediatric OR fetal OR foetal OR perinatal OR pregnant OR pregnancy OR childbirth OR birth OR labor OR labour)	0
7	1 OR 2 OR 3 OR 4 OR 5 OR 6	438

Web of Science Core Collection, Thomson Reuters

Set	Results	
# 19	2634	#18 OR #15 OR #12
# 18	141	#17 AND #10
# 17	999,282	#16 NOT (#11 OR #13)
# 16	1,931,491	TS=(economic* OR cost* OR pharmacoeconomic* OR pharmaco-economic* OR "value of life" OR "quality adjusted life year*" OR qaly* OR qald* OR qale* OR "disability adjusted life year*" OR daly* OR sfg OR "short form 6" OR shortform6 OR "quality of life" OR euro-qol OR "euro quality" OR eq5d OR eq-5d OR hye OR "health year equivalent*" OR "healthy year equivalent*" OR "willingness to pay" OR "standard gamble" OR "time trade off" OR "time tradeoff" OR tto OR vas OR "visual analog" OR "resource use" OR "resource utilisation" OR "decision tree*" OR "decision model*" OR "decision analysis")
# 15	1500	#14 AND #10
# 14	10,439,530	#13 NOT #11

Death audits and reviews for reducing maternal, perinatal and child mortality (Review)

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(Continued)

# 13	11,998,514	TS=(intervention? or effect? or impact? or controlled or control group? or (before SAME after) or (pre SAME post) or ((pretest or "pre test") and (posttest or "post test"))) or quasi-experiment* or "quasi experiment*" or evaluat* or "time series" or "time point?" or "repeated measur*")
# 12	993	#11 AND #10
# 11	3,030,368	TS=(random* or blind* or allocat* or assign* or trial* or placebo* or crossover* or cross-over*)
# 10	4704	#9 OR #8 OR #7 OR #6 OR #5 OR #4 OR #3 OR #2 OR #1
# 9	280	TS=((("near miss*" or "significant event*" or "critical event*" or "critical incident?") and (maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r) and (review* or audit* or meeting? or enquir* or inquir*)))
# 8	543	TS=((("verbal autops*" or "social autops*") SAME (maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r)))
# 7	37	TS=("saving mothers lives" or "making pregnancy safer" or "making childbirth safer")
# 6	101	TS=(cemach or cmace or cemd or cmde)
# 5	66	TS=(confidential enquir* or confidential inquir*) AND TS=(stillbirth? or sudden infant death? or sids or cot death? or crib death?)
# 4	286	TS=(confidential enquir* or confidential inquir*) AND TS=((maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r) NEAR/3 (mortality or death?)))
# 3	1254	TS=((stillbirth? or sudden infant death? or sids or cot death? or crib death?) SAME (review* or audit* or meeting? or enquir* or inquir*)))
# 2	1247	TS=((death? or mortality) NEAR/3 (review* or audit* or meeting? or enquir* or inquir*))) AND TS=(maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r)
# 1	1916	TS=((maternal or mother* or maternity or child* or infan* or p?ediatric* or fetal or foetal or perinatal or pregnan* or childbirth or birth or labo?r) NEAR/3 (mortality or death?))) AND TI=(review* or audit* or meeting? or enquir* or inquir*)

CENTRAL & NHSEED, Cochrane Library, Wiley

ID	Search
#1	MeSH descriptor: [Child Mortality] explode all trees
#2	MeSH descriptor: [Fetal Mortality] explode all trees
#3	MeSH descriptor: [Perinatal Mortality] explode all trees

Death audits and reviews for reducing maternal, perinatal and child mortality (Review)

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(Continued)

#4	MeSH descriptor: [Pregnancy Complications] this term only and with qualifier(s): [Mortality - MO]
#5	MeSH descriptor: [Stillbirth] explode all trees
#6	MeSH descriptor: [Sudden Infant Death] explode all trees
#7	MeSH descriptor: [Pregnant Women] explode all trees
#8	MeSH descriptor: [Child] explode all trees
#9	MeSH descriptor: [Infant] explode all trees
#10	#7 or #8 or #9
#11	MeSH descriptor: [Mortality] this term only
#12	MeSH descriptor: [Cause of Death] explode all trees
#13	#11 or #12
#14	#10 and #13
#15	((maternal or mother* or maternity or child* or infan* or pediatric* or paediatric* fetal or foetal or perinatal or pregnan* or childbirth or birth or labor or labour) near (mortality or death*)):ti,ab,kw
#16	stillbirth* or "sudden infant death*" or sids or "cot death*" or "crib death*":ti,ab,kw
#17	#1 or #2 or #3 or #4 or #5 or #6 or #14 or #15 or #16
#18	MeSH descriptor: [Medical Audit] explode all trees
#19	MeSH descriptor: [Clinical Audit] this term only
#20	(review* or audit* or meeting* or enquir* or inquir*):ti,ab,kw
#21	#18 or #19 or #20
#22	#17 and #21
#23	((((death* or mortality) near (review* or audit* or meeting* or enquir* or inquir*)) and (maternal or mother* or maternity or child* or infan* or pediatric* or paediatric* fetal or foetal or perinatal or pregnan* or childbirth or birth or labor or labour)):ti,ab,kw
#24	((stillbirth* or "sudden infant death*" or sids or "cot death*" or "crib death*") near (review* or audit* or meeting* or enquir* or inquir*)):ti,ab,kw
#25	((confidential enquir* or confidential inquir*) and ((maternal or mother* or maternity or child* or infan* or pediatric* or paediatric* fetal or foetal or perinatal or pregnan* or childbirth or birth or labor or labour) near (mortality or death*)):ti,ab,kw
#26	((confidential enquir* or confidential inquir*) and (stillbirth* or "sudden infant death*" or sids or "cot death*" or "crib death*")):ti,ab,kw
#27	(cemach or cmace or cemd or cmde):ti,ab,kw
#28	("saving mothers lives" or "making pregnancy safer" or "making childbirth safer"):ti,ab,kw

(Continued)

#29	((("verbal autops*" or "social autops*") near (maternal or mother* or maternity or child* or infan* or pediatric* or paediatric* fetal or foetal or perinatal or pregnan* or childbirth or birth or labor or labour)):ti,ab,kw
#30	((("near miss*" or "significant event*" or "critical event*" or "critical incident*") and (maternal or mother* or maternity or child* or infan* or pediatric* or paediatric* fetal or foetal or perinatal or pregnan* or childbirth or birth or labor or labour)):ti,ab,kw
#31	#22 or #23 or #24 or #25 or #26 or #27 or #28 or #29 or #30

CINAHL, EBSCOHost

#	Query
S27	S26 OR S22 OR S19
S26	S17 AND S25
S25	S23 NOT S24
S24	S18 OR S20
S23	TI (economic* OR cost* OR pharmacoeconomic* OR pharmaco-economic* OR "value of life" OR "quality adjusted life year*" OR qaly* OR qald* OR qale* OR "disability adjusted life year*" OR daly* OR sfg OR "short form 6" OR shortform6 OR "quality of life" OR euroqol oR "euro quality" OR eq5d OR eq-5d OR hye OR "health year equivalent*" OR "healthy year equivalent*" OR "willingness to pay" OR "standard gamble" OR "time trade off" OR "time tradeoff" OR tto OR vas OR "visual analog" OR "resource use" OR "resource utilisation" OR "decision tree*" OR "decision model*" OR "decision analysis") OR AB (economic* OR cost* OR pharmacoeconomic* OR pharmaco-economic* OR "value of life" OR "quality adjusted life year*" OR qaly* OR qald* OR qale* OR "disability adjusted life year*" OR daly* OR sfg OR "short form 6" OR shortform6 OR "quality of life" OR euroqol oR "euro quality" OR eq5d OR eq-5d OR hye OR "health year equivalent*" OR "healthy year equivalent*" OR "willingness to pay" OR "standard gamble" OR "time trade off" OR "time tradeoff" OR tto OR vas OR "visual analog" OR "resource use" OR "resource utilisation" OR "decision tree*" OR "decision model*" OR "decision analysis")
S22	S17 AND S21
S21	S20 NOT S18
S20	TI (intervention* or effect or effects or impact or impacts or controlled or "control group" or "control groups" or (before N5 after) or (pre N5 post) or ((pretest or "pre test") and (posttest or "post test")) or quasiexperiment* or "quasi experiment*" or evaluat* or "time series" or "time point*" or "repeated measur*") OR ((intervention* or effect or effects or impact or impacts or controlled or "control group" or "control groups" or (before N5 after) or (pre N5 post) or ((pretest or "pre test") and (posttest or "post test")) or quasiexperiment* or "quasi experiment*" or evaluat* or "time series" or "time point*" or "repeated measur*"))
S19	S17 AND S18
S18	TI (random* or blind* or allocat* or assign* or trial* or placebo* or crossover* or cross-over*) OR AB (random* or blind* or allocat* or assign* or trial* or placebo* or crossover* or cross-over*)

(Continued)

S17	S1 OR S2 OR S3 OR S4 OR S5 OR S6 OR S7 OR S8 OR S9 OR S10 OR S11 OR S12 OR S13 OR S14 OR S15 OR S16
S16	TI (("near miss*" or "significant event*" or "critical event*" or "critical incident?") and (maternal or mother* or maternity or child* or infant* or pediatric* or paediatric* fetal or foetal or perinatal or pregnan* or childbirth or birth or labor or labour) and (review* or audit* or meeting? or enquir* or inquir*)) or AB (("near miss*" or "significant event*" or "critical event*" or "critical incident?") and (maternal or mother* or maternity or child* or infant* or pediatric* or paediatric* fetal or foetal or perinatal or pregnan* or childbirth or birth or labor or labour) and (review* or audit* or meeting? or enquir* or inquir*))
S15	TX ((verbal autops* or social autops*) N5 (maternal or mother* or maternity or child* or infant* or pediatric* or paediatric* fetal or foetal or perinatal or pregnan* or childbirth or birth or labor or labour))
S14	TX saving mothers lives or making pregnancy safer or making childbirth safer
S13	TX cemach or cmace or cemd or cmde
S12	TX ((confidential enquir* or confidential inquir*) and (stillbirth* or "sudden infant death*" or sids or "cot death*" or "crib death*"))
S11	TX ((confidential enquir* or confidential inquir*) and ((maternal or mother* or maternity or child* or infant* or pediatric* or paediatric* fetal or foetal or perinatal or pregnan* or childbirth or birth or labor or labour) N3 (mortality or death*)))
S10	TX ((stillbirth* or "sudden infant death*" or sids or "cot death*" or "crib death*") N5 (review* or audit* or meeting* or enquir* or inquir*))
S9	TI (((death* or mortality) N3 (review* or audit* or meeting* or enquir* or inquir*)) and (maternal or mother* or maternity or child* or infant* or pediatric* or paediatric* fetal or foetal or perinatal or pregnan* or childbirth or birth or labor or labour)) or AB (((death* or mortality) N3 (review* or audit* or meeting* or enquir* or inquir*)) and (maternal or mother* or maternity or child* or infant* or pediatric* or paediatric* fetal or foetal or perinatal or pregnan* or childbirth or birth or labor or labour))
S8	(((MH "Pregnancy Complications+/MO") OR (MH "Perinatal Death") OR (MH "Sudden Infant Death"))) AND (TI ((review* or audit* or meeting? or enquir* or inquir*)) or AB ((review* or audit* or meeting? or enquir* or inquir*)))
S7	(((MH "Mortality") OR (MH "Cause of Death")) AND ((MH "Expectant Mothers") OR (MH "Child+"))) AND (TI ((review* or audit* or meeting? or enquir* or inquir*)) or AB ((review* or audit* or meeting? or enquir* or inquir*)))
S6	(((MH "Child Mortality") OR (MH "Infant Mortality") OR (MH "Maternal Mortality"))) AND (TI ((review* or audit* or meeting? or enquir* or inquir*)) or AB ((review* or audit* or meeting? or enquir* or inquir*)))
S5	(((stillbirth* or "sudden infant death*" or sids or "cot death*" or "crib death*")) AND (MH "Audit"))
S4	(((maternal or mother* or maternity or child* or infant* or pediatric* or paediatric* fetal or foetal or perinatal or pregnan* or childbirth or birth or labor or labour) N3 (mortality or death*))) AND (MH "Audit")
S3	(((MH "Pregnancy Complications+/MO") OR (MH "Perinatal Death") OR (MH "Sudden Infant Death"))) AND (MH "Audit")

(Continued)

S2	((MH "Mortality") OR (MH "Cause of Death")) AND ((MH "Expectant Mothers") OR (MH "Child+")) AND (MH "Audit")
S1	((MH "Child Mortality") OR (MH "Infant Mortality") OR (MH "Maternal Mortality")) AND (MH "Audit")

ClinicalTrials.gov

Other terms = (death review OR mortality review OR death reviews OR mortality reviews) AND Condition=maternal OR mother OR mothers OR maternity OR child OR children OR childhood OR infant OR infants OR pediatric OR paediatric OR fetal OR foetal OR perinatal OR pregnant OR pregnancy OR childbirth OR birth OR labor OR labour	205
Other terms=((death OR deaths OR mortality) AND (audit OR audits OR enquiry OR enquiries OR inquiry OR inquiries OR meeting OR meetings)) AND Conditions=(maternal OR mother OR mothers OR maternity OR child OR children OR childhood OR infant OR infants OR pediatric OR paediatric OR fetal OR foetal OR perinatal OR pregnant OR pregnancy OR childbirth OR birth OR labor OR labour)	144
(audit OR audits OR enquiry OR enquiries OR inquiry OR inquiries OR meeting OR meetings) stillbirth OR stillbirths OR sudden infant death OR sudden infant deaths OR sids OR cot death OR cot deaths OR crib death OR crib deaths	4
confidential enquiry OR confidential enquiries OR "confidential inquiry" OR "confidential inquiries" OR cemach OR cmace OR cemd OR cmde OR saving mothers lives OR making pregnancy safer OR making childbirth safer	39
verbal autopsy OR verbal autopsies OR social autopsy OR social autopsies maternal OR mother OR mothers OR maternity OR child OR children OR childhood OR infant OR infants OR pediatric OR paediatric OR fetal OR foetal OR perinatal OR pregnant OR pregnancy OR childbirth OR birth OR labor OR labour	20
(near miss OR significant events OR critical events OR critical incidents OR significant event OR critical event OR critical incident) AND (audit OR audits OR enquiry OR enquiries OR inquiry OR inquiries OR meeting OR meetings) (maternal OR mother OR mothers OR maternity OR child OR children OR childhood OR infant OR infants OR pediatric OR paediatric OR fetal OR foetal OR perinatal OR pregnant OR pregnancy OR childbirth OR birth OR labor OR labour)	44
	456

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Intervention=(death review OR mortality review OR death reviews OR mortality reviews) AND Condition=maternal OR mother OR mothers OR maternity OR child OR children OR childhood OR infant OR infants OR pediatric OR paediatric OR fetal OR foetal OR perinatal OR pregnant OR pregnancy OR childbirth OR birth OR labor OR labour	2
Title=(death OR deaths OR mortality) AND (audit OR audits OR enquiry OR enquiries OR inquiry OR inquiries OR meeting OR meetings)	45

Death audits and reviews for reducing maternal, perinatal and child mortality (Review)

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(Continued)

Title=(stillbirth OR stillbirths OR sudden infant death OR sudden infant deaths OR sids OR cot death OR cot deaths OR crib death OR crib deaths) AND Intervention=(audit OR audits OR enquiry OR enquiries OR inquiry OR inquiries OR meeting OR meetings)	4
Title=(confidential enquiry OR confidential enquiries OR “confidential inquiry” OR “confidential inquiries” OR cernach OR cernace OR cerned OR cernde OR saving mothers lives OR making pregnancy safer OR making childbirth safer)	3
Title=(maternal OR mother OR mothers OR maternity OR child OR children OR childhood OR infant OR infants OR pediatric OR paediatric OR fetal OR foetal OR perinatal OR pregnant OR pregnancy OR childbirth OR birth OR labor OR labour) AND Intervention=(verbal autopsy OR verbal autopsies OR social autopsy OR social autopsies)	2
Title=(near miss OR significant events OR critical events OR critical incidents OR significant event OR critical event OR critical incident)	13
	69

WHAT'S NEW

Date	Event	Description
30 March 2020	Amended	Added missing declaration of interest statement.

CONTRIBUTIONS OF AUTHORS

Conceiving the protocol: all authors.

Designing the protocol: MW, BN, JP, BS, AD, AH.

Co-ordinating the protocol: MW.

Designing search strategies: MW, NR.

Writing the protocol: MW, BN, JP, BS.

Searching and screening studies: JP, SS, MW.

Providing general advice on the protocol: AH, AD.

Performing previous work that was the foundation of the current study: AD, MW, BN, AH, HA, VM.

DECLARATIONS OF INTEREST

MW: none.

JP: none.

SS: none.

BN: none.

BS: none.

NR: none.

HA: none.

Death audits and reviews for reducing maternal, perinatal and child mortality (Review)

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VM: none.

AD is the lead author of one of the trials included in this review. He was not involved in assessing the quality of the trial he led.

AH: none.

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Internal sources

- No sources of support supplied

External sources

- NIHR, UK.

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- Department for International Development, UK.

Project number 300342-104

DIFFERENCES BETWEEN PROTOCOL AND REVIEW

We did not search the two grey literature collections, OpenGrey and the Grey Literature Report. However, we consulted widely, in particular with experts at the World Health Organization, to identify further studies.

We did not specify “Perinatal morbidity related to suboptimal care” as an important outcome in the protocol, but we consider this as an important outcome as it relates to quality of care, and have included it in the 'Summary of findings' table.