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Primary care strategies to improve childhood immunisation uptake in developed countries: systematic review

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Summary

Objectives To conduct a systematic review of strategies to optimize immunisation uptake within preschool children in developed countries.

Design Systematic review.

Setting Developed countries

Participants Preschool children who were due, or overdue, one or more of their routine primary immunisations.

Main outcome measures Increase in the proportion of the target population up to date with standard recommended universal vaccinations.

Results Forty-six studies were included for analysis, published between 1980 and 2009. Twenty-six studies were randomized controlled trials, 11 were before and after trials, and nine were controlled intervention trials. Parental reminders showed a statistically significant increase in immunisation rates in 34% of included intervention arms. These effects were reported with both generic and specific reminders and with all methods of reminders and recall. Strategies aimed at immunisation providers were also shown to improve immunisation rates with a median change in immunisation rates of 7% when reminders were used, 8% when educational programmes were used and 19% when feedback programmes were used.

Conclusion General practitioners are uniquely positioned to influence parental decisions on childhood immunisation. A variety of strategies studied in primary care settings have been shown to improve immunisation rates, including parental and healthcare provider reminders.

Introduction

Childhood vaccines currently save 3 million lives a year globally and are among the most successful and cost-effective public health interventions of the 20th century. Immunisation has been responsible for substantial falls in serious bacterial infectious diseases in children including tetanus, diphtheria, meningococcal serogroup C (MenC), measles and Haemophilus Influenzae B (HiB),
since their inclusion in the UK primary vaccination schedule. Immunising children not only protects individuals from infection but also contributes to population-based immunity by reducing the circulation of infectious agents leading to community-wide health gains.

To maximize the potential population-wide benefits of routine vaccination through herd immunity, the World Health Organization (WHO) has set national targets of 95% coverage annually for each antigen in the routine immunisation schedule by 2 years of age. Yet recent figures show that these targets are not being met in many countries. In England, in the quarter between January and March 2010, vaccination coverage for the measles, mumps and rubella vaccine (MMR), pneumococcal vaccine (PCV) booster and the Hib/MenC vaccines were 88%, 88% and 90%, respectively, at 24 months of age.

Coverage tends to be lowest in deprived, urban areas with mobile populations. For example, childhood immunisation rates for London are substantially below the national average with levels in one East London borough (Bexley) of just 63%, 71% and 74% for the PCV (Pneumococcal conjugate vaccine) booster, MMR and Hib/MenC, respectively. In areas of low MMR coverage, this has resulted in outbreaks, with a high proportion of those affected being under 1 year of age, who were most likely infected by unimmunised older contacts. One consequence of this is that measles and mumps have once again become endemic in the UK, with over 1100 cases of measles and over 7000 cases of mumps reported in the UK in 2009, 14 years after local transmission of measles was halted.

Barriers to immunisation can stem from parental concerns about risks, inadequate knowledge and provision by providers, and generalized systemic barriers involving the organization of the health system and access to services. Ninety-eight percent of infants born in the UK are registered with a UK general practitioner (GP) and their first contact with their GP is often at the primary vaccination. Hence, GPs and practice nurses are uniquely positioned to influence a parent’s decisions to have their child immunised.

Our objective was to conduct a systematic literature review aimed at providing GPs with up-to-date, evidence-based guidelines on how to improve uptake rates of primary immunisations for children registered under their care. Our research question was ‘How can primary care practitioners in developed countries improve preschool immunization uptake?’

**Methods**

**Search strategy and data sources**

We systematically searched electronic databases including MEDLINE, EMBASE, PsycInfo, Cochrane and OpenSIGL from inception to 1 June 2010 using MESH and Key Terms including immun*, vaccin*, innoc* and rates, uptake and coverage (Appendix 1) to identify studies reporting interventions to improve preschool immunisation uptake and to evaluate their effectiveness in children in developed countries.

We hand-searched the reference lists from retrieved studies and reviews to find additional studies and contacted experts in the field. We also identified relevant grey literature such as conference papers, dissertations and government guidelines.

**Study selection**

Two reviewers (NW and HW) independently screened titles and abstracts of all citations for eligibility and retrieved those that met the inclusion criteria. If insufficient information was available in the abstract to decide on eligibility the whole article was retrieved for review. Discrepancies were resolved by discussion and by involving a third reviewer (SS) when necessary.

We included experimental studies reporting original research including randomized controlled trials, controlled clinical trials, before and after studies and interrupted time-series studies, published in English. Our target population was children under the age of 5 years living in developed countries. We included studies reporting our main outcome measure; the increase in the proportion of the target population who were up to date with standard recommended universal vaccinations. Outcomes could be for either single vaccinations or combinations of vaccines due. We excluded studies for which the full article was not available, and studies that did not contain
any original data such as review articles, commentaries and correspondence.

Data extraction and quality assessment
The two reviewers independently extracted data from included studies on study setting, participant characteristics, clinical setting, interventions and outcomes measured. Included studies were graded for methodological quality using 26 points from the 27 point score devised by Downs and Black (Appendix 2).12

Results
A total of 32,624 studies were retrieved from the electronic searches of which 32,410 were excluded after reviewing their titles and abstracts. The remaining 214 papers were retrieved for full text review. Recommendations from experts in the field and reviews of reference lists identified a further 14 studies. A total of 46 papers met the inclusion criteria (Figure 1).

The commonest reasons for exclusion were failure to report interventions or outcomes of interest and poor study design.

The 46 included studies were published between 1980 and 2009. Twenty-six were randomized controlled trials, 11 were before and after trials and nine were controlled intervention trials. The studies originated mainly from the USA (36) and the remainder originated from the UK and Ireland (7), Australia (1), New Zealand (1) and Finland (1). The study participants ranged in age from 6 weeks to 11 years of age but the majority of studies focused on preschool children. Studies were conducted in different settings including paediatric outpatient clinics, family practices, primary care clinics, community health centres, managed care organizations and health maintenance organizations in the USA; and community clinics and general practices in the UK, New Zealand and Australia.

Studied interventions aimed to remind and/or recall parents of upcoming or overdue vaccinations; targeted providers to improve uptake through feedback, audit or chart prompts; provided simple education to parents within a general practice setting or consisted of multicomponent interventions.

Client-based interventions
Client-based interventions target the parent and child to increase the demand for immunisation services. Educating parents and communities on the benefits of vaccinating their children can empower parents to practice preventative healthcare and thereby improving immunisation uptake.

Reminder and recall
Reminders aim to advise parents of upcoming vaccinations that are due and remind parents of those children that are overdue. They vary in methodology from automated telephone calls and generic postcards to personalized letters and even home visits.

Twenty-two included papers reported on 41 intervention arms studying parental reminders and recalls.13–34 Details of the included studies are seen in Table 1. The average score for study quality using Down and Black’s quality scoring framework was 24.8 out of a potential 31 (range 21–29.5). Fourteen (34%) of the 41 intervention arms showed a statistically significant ($P < 0.05$) increase

Figure 1

Quorom diagram

- 32,624 potentially relevant citations identified and screened for retrieval
- 32,410 excluded after screening title and abstract
- 214 included for full text analysis
- 168 excluded after full text analysis including
  - 89 duplicates
  - 79 did not satisfy inclusion criteria
- 46 papers satisfied inclusion criteria


Primary care strategies to improve childhood immunisation uptake in developed countries
Table 1
Study characteristics of reminder and recall studies

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<tr>
<th>Paper</th>
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<th>Design</th>
<th>Quality</th>
<th>Intervention</th>
<th>Outcome</th>
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<tbody>
<tr>
<td>Abramson et al.</td>
<td>1993</td>
<td>Public health centre and children’s hospital continuity clinic, Forsyth County, North Carolina, USA; low socioeconomic status</td>
<td>Randomized controlled trial</td>
<td>23</td>
<td>1. Usual care (control group) vs. 2. Postcard reminders followed by telephone reminders</td>
<td>Age appropriate immunisations (DTP/OPV/Hib) at 7 months of age 1 vs. 2, net change = 19% (P &lt; 0.00001)</td>
</tr>
<tr>
<td>Alemi et al.</td>
<td>1993–1994</td>
<td>Paediatric outpatient clinic, Cleveland, USA; children under 6 months of age at recruitment; urban; predominantly ethnic minorities; low socioeconomic status</td>
<td>Controlled intervention trial</td>
<td>22.5</td>
<td>1. Usual care (control group) vs. 2. Computer-generated telephone reminders and recalls</td>
<td>On time immunisation with complete series (DTP/OPV/MMR/Hib): 1 vs. 2, net change = 24.4% (P = 0.0005)</td>
</tr>
<tr>
<td>Alto et al.</td>
<td>1991</td>
<td>Family practice clinic, Colorado, USA; children between 2 months and 7 years old; low socioeconomic status</td>
<td>Randomized controlled trial</td>
<td>26</td>
<td>1. Usual care (control group) vs. 2. Specific postcard reminders followed by telephone reminders</td>
<td>Up to date with DTP/OPV/MMR/Hib vaccinations: 1 vs. 2, net change = 8% (P &lt; 0.011)</td>
</tr>
<tr>
<td>Daley et al.</td>
<td>2000</td>
<td>Primary Care Clinic, Denver, Colorado, USA; Predominantly Medicaid and uninsured patient population; Children aged 6 weeks to 22 months</td>
<td>Randomized controlled trial</td>
<td>29.5</td>
<td>1. Usual care (control group) vs. 2. Reminder letter followed by telephone recall 10 days later</td>
<td>Immunisation with one or more doses of PCV7: 1 vs. 2, net change = 2.8% (95% confidence interval 1.8% to 7.4%)</td>
</tr>
<tr>
<td>Dini et al.</td>
<td>1993–1996</td>
<td>County health department, Denver, USA; children aged 60–90 days who had received the first dose of DTP and/or IPV</td>
<td>Randomized controlled trial</td>
<td>25</td>
<td>1. Usual care (control group) vs. 2. Telephone messages vs. 3. Letter reminders vs. 4. Telephone messages followed by letters</td>
<td>Completed immunisation series by 24 months of age 1 vs. any intervention net change = 8.3% (RR (rate ratio) = 1.21; CI (confidence interval) = 1.01-1.44). 1 vs. 2 net change = 8.4%, 1 vs. 3 net change = 7.3%, 1 vs. 4 net change = 9%</td>
</tr>
<tr>
<td>Hawe et al.</td>
<td>Not reported</td>
<td>Local government-operated public vaccination clinic, Ballarat, Australia, 259 children due for measles vaccination aged 12 months</td>
<td>Randomized controlled trial</td>
<td>26.5</td>
<td>1. Usual reminder card 2. Health belief model reminder card</td>
<td>Usual card group 67% vaccinated Health belief model card group 79% 12% difference (0.026)</td>
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<tr>
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<tbody>
<tr>
<td>Irigoyen et al. 19</td>
<td>1997</td>
<td>Paediatric clinic, New York, USA; children aged 4–18 months</td>
<td>Controlled intervention trial</td>
<td>26.5</td>
<td>1. Usual care (control group) vs. 2. Postcard reminder vs. 3. Telephone reminder vs. 4. Postcard plus telephone reminder</td>
<td>No significant difference in vaccination coverage by study group</td>
</tr>
<tr>
<td>Irigoyen et al. 20</td>
<td>2001</td>
<td>Five inner-city community paediatric practices, New York, USA; children aged 6 weeks to 15 months due or late for DTP</td>
<td>Randomized controlled trial</td>
<td>27</td>
<td>1. Usual care (control group) vs. 2. Continuous postcard reminders (as many as needed) vs. 3. Limited postcard reminders (up to 3)</td>
<td>Up to date 4:3:1:3 at 3 months post randomization 1 vs. 2 net change = 5.3% (P &lt; 0.01), 1 vs. 3 net change = 1.5% (not significant) Up to date 4:3:1:3 at 6 months post randomization 1 versus 2 net change = 4.9% (p &lt; 0.05), 1 vs. 3 net change = 2.8% (not significant) Multivariate analysis showed reminders had no independent effect on immunization outcomes UTD at 12 months 1 vs. 2 net change 12% (P = 0.07)</td>
</tr>
<tr>
<td>Kempe et al. 21</td>
<td>1999</td>
<td>Outpatient paediatric clinic, urban teaching hospital, Denver, Colorado, USA; children aged 5–17 months; low socioeconomic status; highly transient population</td>
<td>Randomized controlled trial</td>
<td>26.5</td>
<td>1. Usual care (control group) vs. 2. Postcard reminder plus telephone reminders</td>
<td></td>
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<tr>
<td>LeBaron et al. 22</td>
<td>1996–1998</td>
<td>Fulton County, Georgia, USA; children born between 1 July 1995 and 6 August 1995</td>
<td>Randomized controlled trial</td>
<td>25</td>
<td>1. Usual care (control group) vs. 2. Automated telephone or mail reminder recall vs. 3. In-person telephone, mail or home visit recall vs. 4. Auto-dialler with outreach backup</td>
<td>UTD with DTP/OPV/MMR/Hib (4:3:4:1) at 24 months of age 1 vs. 2 net change = 6% (P &lt; 0.05), 1 vs. 3 net change = 3% (not significant), 1 vs. 4 net change = 4% (not significant)</td>
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<tr>
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<tbody>
<tr>
<td>Lieu et al.</td>
<td>1994–1995</td>
<td>Managed care organization, Northern California, children aged 20–24 months old, ( n = 149 ) control group, ( n = 172 ) intervention group</td>
<td>Randomized controlled trial</td>
<td>23.5</td>
<td>1. Usual care (control group) vs. 2. Patient recalls via computer-generated personalized letter</td>
<td>MMR by age 24 months, 1 vs. 2 = 19% net change ((P &lt; 0.001))</td>
</tr>
<tr>
<td>Lieu et al.</td>
<td>1996–1997</td>
<td>Health Maintenance Organization, California, USA; under-immunised 20-month old children</td>
<td>Randomized controlled trial</td>
<td>23</td>
<td>1. Automated telephone reminder vs. 2. Letter reminder vs. 3. Automated telephone reminder followed by a reminder letter 1 week later vs. 4. Letter reminder followed by an automated telephone message 1 week later</td>
<td>Proportion of under-immunised children who received any needed vaccinations by age 24 months, 1 vs. 2 no net change; 1 vs. 4 net change 14% ((P = 0.02)); 2 vs. 4 14% ((P = 0.01)); 1 vs. 3 net change 9% ((P = 0.1)), 2 vs. 3 net change 9% ((P = 0.09))</td>
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<tr>
<td>Mason and Donnelly</td>
<td>1998–1999</td>
<td>Local health authority, Wales, UK; children aged 21 months who had not received MMR vaccine</td>
<td>Randomized controlled trial</td>
<td>24</td>
<td>1. Usual care vs. 2. Personal reminder letter and educational leaflet</td>
<td>Immunised with MMR at age 21–24 months of age, 1 vs. 2, net change = 1.1% ((95% \text{ CI } -3.3–5.5)); immunised with MMR at &gt;24 months of age, 1 vs. 2 = 1.1% ((95% \text{ CI } -3.6–5.7))</td>
</tr>
<tr>
<td>Morgan et al.</td>
<td>1996</td>
<td>South Glamorgan, Wales, UK; children aged between 9 months and 21 months</td>
<td>Randomized controlled trial</td>
<td>27.5</td>
<td>1. Usual care (control group) vs. 2. Non-directive telephone call to child’s health visitor vs. 3. Mailed reminder</td>
<td>Immunicated with MMR 1 vs. 2 net change = −7%, 1 vs. 3 net change = −11%</td>
</tr>
<tr>
<td>Rodewald et al.</td>
<td>1994–1995</td>
<td>Nine primary care practices, Rochester, New York, USA; 3015 infants</td>
<td>Randomized controlled trial (two by two factorial design)</td>
<td>25.5</td>
<td>1. Usual care (control group) vs. 2. Tracking/Outreach/prompting- lay outreach workers using postcards, telephone calls and home visits 3. Tracking/outreach 4. Prompting- during visits to primary care office, marking charts</td>
<td>UTD net change 1 vs. 2 = 21%; 1 vs. 3 21%; 1 vs. 4 = 2% (none showed significance); 1 vs. 2 plus 3 ( P &lt; 0.001 )</td>
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Table 1
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<tr>
<td>Stehr-Green et al.</td>
<td>1990</td>
<td>Public health clinics, Atlanta, Georgia, USA; Average age 8.7–9.2 months; n = 110 control group, n = 112 intervention group</td>
<td>Randomized controlled trial</td>
<td>24.5</td>
<td>1. Usual care (control group) vs. 2. Automated telephone reminder (max 9 attempts)</td>
<td>DTP immunisation, 1 vs. 2 = 3% net change (non-significant)</td>
</tr>
<tr>
<td>Szilagyi et al.</td>
<td>1993–1996</td>
<td>Three geographical regions of Monroe County, New York, USA; children aged up to 2 years</td>
<td>Randomized controlled trial</td>
<td>24.5</td>
<td>1. Usual care 1993 (control group) vs. 2. Staged intervention of increasing intensity letters, postcards or phone calls and outreach 1999</td>
<td>UTD with DTP/OPV/MMR/Hib 4:3:1:1 1 vs. 2 net gain = Monroe county 20%, suburbs 15%, inner city 29%, rest of city 17% (significance not stated)</td>
</tr>
<tr>
<td>Tollestrup and Hubbard</td>
<td>1987</td>
<td>County health department clinics, Washington, USA; children aged up to 5 years</td>
<td>Controlled intervention trial</td>
<td>23.5</td>
<td>1. Usual care (control group) vs. 2. Recall letter if 1 month overdue</td>
<td>DTP vaccination within 5 months, 1 vs. 2 net change = 18% (P &lt; 0.01)</td>
</tr>
<tr>
<td>Vivier et al.</td>
<td>1998</td>
<td>Primary care clinics, Rhode Island, USA; children enrolled in a managed care programme, up to the age of 6 years</td>
<td>Randomized controlled trial</td>
<td>24</td>
<td>1. Usual care (control group) vs. 2. Telephone reminder vs. 3. Postal reminder vs. 4. Sequential postal/ telephone reminders</td>
<td>Immunisations UTD, 1 vs. 2, net change = 11% (P &gt; 0.05); 1 vs. 3, net change = 12% (P &gt; 0.05); 1 vs. 4, net change = 14% (P &gt; 0.05); 1 vs. 2–4 combined P &lt; 0.05 One immunisation received during study period 1 vs. 2 net change = 18% (P &lt; 0.001); UTD with immunisations 1 vs. 2 net change = 1% (not significant)</td>
</tr>
<tr>
<td>Wilcox et al.</td>
<td>1997</td>
<td>Philadelphia, Pennsylvania, USA; 1752 children aged 6–10 months</td>
<td>Randomized controlled trial</td>
<td>24.5</td>
<td>1. Usual care (control group) vs. 2. Outreach</td>
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(Continued)
in immunisation rates (Szilagyi et al.\textsuperscript{29} did not report significance). Overall, these studies reported a median point change of 11\% (mean 10\%, range \(-11\%\) to 24\%). Studies comparing usual care with postal reminders alone\textsuperscript{17,19,20,23–26,30,33,34} reported an overall median point change of 10\% (mean 8.6\%, range \(-11\%\) to 19\%). Studies looking at telephone reminders alone\textsuperscript{14,17,28,31} reported an overall median point change of 9.5\% (mean 12\%; range 3–24\%). Those studies looking at the effect of the combination of postal and telephone reminders on vaccination rates\textsuperscript{13,15–17,21,31} reported an overall median point change of 10.5\% (mean 10.8\%, range 2.8–19\%). One study\textsuperscript{18} compared a traditional reminder card with a card designed with health belief model in mind. Hawe reported a 12\% increase in immunisation rates when the health belief model reminder card was used.

Parental reminders have been shown to have an overall positive effect on immunisation uptake. These effects have been reported with both generic and specific reminders and with all methods of reminders and recall.

### Parental education

In the context of this review, we considered only educational programmes that could feasibly be delivered within the setting of primary care and we excluded studies reporting national or regional education programmes.

Two of the included papers reported on two intervention arms studying the effect of simple parental education programmes on immunisation uptake (Table 2)\textsuperscript{35,36}. One study assessed the impact of a promotional teddy bear featuring the address of an information website for MMR and one studied the impact of an interactive graphic card and verbal explanation on immunisation uptake. The quality of these two studies averaged at 23.8 points from a possible 31. Neither study showed a significant effect on immunisation rates and the limited number of studies precludes us from reaching an evidenced-based conclusion on the effect of these strategies on parental behaviour.

### Patient-held records

Only one study\textsuperscript{37} assessing the impact of patient-held records effect on immunisation rates fulfilled

<table>
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<tr>
<th>Paper</th>
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<tr>
<td>Yokley and Grewick\textsuperscript{33}</td>
<td>Public health clinic, Akron, Ohio, USA; children aged 37 months; (n = 183–195) in each group</td>
<td>Not stated</td>
<td>Group randomized controlled trial by family</td>
<td>1. Usual care (control group) vs. 2. General postal patient reminder vs. 3. Specific postal patient reminder vs. 4. Specific postal patient reminder plus special out-of-hours clinics vs. 5. Specific postal patient reminder plus parent incentive lottery</td>
<td>23.5</td>
<td>Vaccinated with at least one antigen after 3 months: 1 vs. 2 = 3% net change (non-significant), 1 vs. 3 = 13% (significant), 1 vs. 4 = 16% (significant), 1 vs. 5 = 18% (significant)</td>
</tr>
<tr>
<td>Young et al.\textsuperscript{34}</td>
<td>Ohio, USA; 6-month old high-risk children</td>
<td>1978</td>
<td>Randomized controlled trial</td>
<td>1. Usual care (control group) vs. 2. Mailed reminder</td>
<td>21</td>
<td>One immunisation received during study period 1 vs. 2 = 16% ((P = 0.02); UTD with vaccinations 1 vs. 2 net change = 12% ((P = 0.06))</td>
</tr>
</tbody>
</table>
Inclusion criteria (Table 3); it scored 22 from a possible 31 points on study quality. This study did not demonstrate a significant difference between usual care and a home-based record booklet. Therefore, it is not possible to come to an evidence-based conclusion on the effect of this strategy on immunisation uptake.

## Provider-based interventions

Provider-based interventions aim to improve vaccination rates by reducing opportunities missed by the medical professional to immunise children. These include provider reminder/recall, assessment and feedback, provider education and a combination of some or all of these strategies.

### Provider reminder/recall

This strategy aims to investigate client’s immunisation status either by manual searching of notes or by automatic computer notifications. Providers are then notified either with paper or computer-based chart prompts that the vaccination is due or overdue.

Five of the included papers reported on six intervention arms studying provider reminder/recall strategies (Table 4). The studies averaged a quality score of 23.7 from a possible 31 (range 21–27). Overall, these studies reported a median point change of 7% (mean 10%, range −2% to 33%). Both manual searching and electronic reminders were shown to have a positive effect on immunisation rates.

### Provider education

Provider education strategies aim to enhance the knowledge of the immunisation provider through a variety of methods including peer support and the use of educational resources. Educational tools may be one-off sessions or part of continuing medical education.

Four of the included papers reported on four intervention arms studying the effect of educating the provider of vaccinations on immunisation rates (Table 3). The average quality score for included studies was 22.4 (range 20–28). Overall, these studies reported a median point change of 8% (mean 10%, range 1–25%).

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**Table 2**

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<thead>
<tr>
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<tbody>
<tr>
<td>Porter-Jones et al.</td>
<td>Not reported</td>
<td>Flintshire, Wales, UK; children eligible for their first dose of MMR</td>
<td>Randomized controlled trial</td>
<td>1. Usual care (control group) vs. 2. Usual care plus a promotional teddy bear displaying an MMR information website and telephone number</td>
<td>22</td>
<td>MMR1, 1 vs. 2 net change 0.7% (P = 0.143)</td>
</tr>
<tr>
<td>Stille et al.</td>
<td>1997–1998</td>
<td>Three paediatric primary care sites, Connecticut, USA; children under 28 days old; Inner city; low socioeconomic status</td>
<td>Controlled intervention trial</td>
<td>1. Usual care (control group) vs. 2. Interactive graphic card plus verbal explanation</td>
<td>25.5</td>
<td>Rate of appropriate immunisation aged 7 months 1 vs. 2 net change 0.4% (P = 0.38); DTP1 by 3 months of age 1 vs. 2 net change 2.8% (P = 0.47)</td>
</tr>
</tbody>
</table>
educational interventions varied from 1-hour peer education sessions to regular continuing medical education within the practice. Two studies looked at provider education alone and two studies looked at education in combination with other interventions such as patient reminders.

Feedback
This strategy retrospectively evaluates the performance of providers in childhood immunisations and feeds this information back to the medical practitioner. Feedback can also be combined with other strategies for example financial incentives or provider education.

Four of the included papers (Taylor, Sinn, Harper, Fairbrother) reported on six intervention arms studying the effect of provider feedback combined with other strategies on immunisation rates (Table 3). The average quality score for these papers was 24.1 (range 21–28). Overall, these studies reported a median point change of 19% (mean 17%, range 12–19%).

Multicomponent interventions
Multicomponent interventions encompass strategies that use a combination of techniques to improve immunisation uptake. These strategies include combining interventions aimed at both the client and the provider.

Eight of the included studies reported on eight intervention arms that combined a variety of interventions aimed at improving immunisation uptake (Table 5). The average score for study quality was 20.5 from a possible 31 (range 17.5–24). Overall, these studies reported a median point change of 15% (mean 19%, range −4% to 47%). Three (38%) of the eight intervention arms reported a statistically significant difference in immunisation rate. Four (50%) studies did not report the significance level for their intervention arms. It is not possible to distinguish which component of the intervention has had the greatest effect on immunisation rates.

Discussion
Numerous studies have reported interventions to improve primary immunisation uptake in children. Effective interventions include parental reminders, which can increase uptake by 11% in the intervention arms. These effects were reported with both generic and specific reminders and with all methods of reminders and recall. Strategies aimed at immunisation providers were also shown to improve immunisation rates with a median change in immunisation rates of 7% when reminders were studied, 8% when educational programmes were studied and 19% when feedback programmes were studied.

Providers who were educated by peers and who received feedback on their performance as vaccine providers were shown to have improved immunisation uptake within their practice. There was limited evidence for patient-held records and parental education alone as strategies for improving immunisation uptake.

Multicomponent strategy studies included interventions aimed at parents alone (e.g. reminder cards plus educational posters) as well as those that combined parental and healthcare...
<table>
<thead>
<tr>
<th>Paper</th>
<th>Study period</th>
<th>Setting and population</th>
<th>Design</th>
<th>Quality</th>
<th>Intervention</th>
<th>Outcome</th>
</tr>
</thead>
<tbody>
<tr>
<td>Burns et al.</td>
<td>1995–1996</td>
<td>Two family health centres, Pittsburgh, Pennsylvania, USA; children aged up to 6 years; 448 intervention group, 529 control group</td>
<td>Randomized controlled trial</td>
<td>25</td>
<td>1. Usual care (control group) vs. 2. Chart prompt</td>
<td>1 vs. 2: Immunisation with DTP4 net change 15% ($P = 0.03$); Immunisation with MMR 1 net change 16% ($P = 0.01$); Immunisation with OPV3 net change 14% ($P = 0.04$); Immunisation with DTP3, DTP5, HepB3 and OPV4, no statistically significant difference 1 vs. 2</td>
</tr>
<tr>
<td>Christy et al.</td>
<td>1990–1991</td>
<td>Two hospital-based primary care centres, Rochester, New York, USA; Children aged 2 to 60 months old; Urban</td>
<td>Controlled intervention trial</td>
<td>21</td>
<td>1. Usual care at site 2, 1 July 1991 – 6 December 1991 (concurrent control group) vs. 2. Nursing intervention guided by an algorithm form at site 1 during same period vs. 3. Usual care at site 1, 1 July 1990 and 30 November 1990 (retrospective control group)</td>
<td>Up to date with DTP/OPV/MMR/Hib aged 24–36 months net change 1 vs. 2 = 0.4%, 2 vs. 3 = 10% (significance not tested)</td>
</tr>
<tr>
<td>Fairbrother et al.</td>
<td>1995–1996</td>
<td>Family and paediatric practices in nine neighbourhoods, New York City, USA; low socioeconomic status</td>
<td>Randomized controlled trial</td>
<td>24.5</td>
<td>1. Usual care (control group) vs. 2. Physician bonus and feedback vs. 3. Enhanced fee for service and feedback 4. Feedback only</td>
<td>UTD with DTP/Hib/OPV/MMR 8 months after baseline, 1 vs. 2 net change = 19.2% ($P &lt; 0.01$), no significant difference between group 3 and control group 1 vs. 2: Up to date for 4DTP/3IPV/1MMR/3Hib/3HepB/1varicella In well child net change 33% (95% CI 32.2–34.6), in sick child 22% (95% CI 20.6–23.1)</td>
</tr>
<tr>
<td>Fiks et al.</td>
<td>2004–2005</td>
<td>Four urban primary care centres, Philadelphia, Pennsylvania, USA; children under 24 months old; &gt;80% ethnic minorities</td>
<td>Before and after study</td>
<td>27</td>
<td>1. Usual care (control group) vs. 2. Electronic Health Record-based clinical reminders</td>
<td>(Continued)</td>
</tr>
<tr>
<td>Paper</td>
<td>Study period</td>
<td>Setting and population</td>
<td>Design</td>
<td>Quality</td>
<td>Intervention</td>
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<tr>
<td>Franzini et al.43</td>
<td>2003–2005</td>
<td>Paediatric and family health practices, Greater Houston, Texas, USA; children aged 12–23 months of age</td>
<td>Before and after study</td>
<td>20.5</td>
<td>1. Usual care vs. 2. 1-hour peer-based educational lunch presentation</td>
<td>Immunisation rates 1 vs. 2 net change = 4% (not significant)</td>
</tr>
<tr>
<td>Harper et al.48</td>
<td>October 1993–October 1994</td>
<td>Family practice clinic (intervention), community health centre (control), St Paul, Minnesota, USA; children aged 24–35 months old; predominantly white, low socioeconomic group</td>
<td>Controlled intervention trial</td>
<td>23</td>
<td>1. Usual care (non-equivalent control group) vs. 2. Physician chart reminder plus feedback on performance plus patient education</td>
<td>4:3:1 DTP / OPV / MMR respectively, 1 vs. 2 = 12% net change (P &lt; 0.02)</td>
</tr>
<tr>
<td>Margolis et al.44</td>
<td>Not reported</td>
<td>44 primary care practices, North Carolina, USA; children aged 24–30 months</td>
<td>Randomized controlled trial</td>
<td>28</td>
<td>1. Usual care (control group) vs. 2. Practice-based continuing medical education</td>
<td>Immunisation rates at 30 months post randomization 1 vs. 2 net change 0.6% (significance not reported)</td>
</tr>
<tr>
<td>Sinn et al.49</td>
<td>Not reported</td>
<td>Ten paediatric group practices, Virginia Beach, Virginia, USA; children aged 9–30 months</td>
<td>Before and after trial</td>
<td>21</td>
<td>1. Usual care (control group) vs. 2. ‘The physician leadership model’ including peer influence and review and goal setting with feedback</td>
<td>Up to date with immunisations at 24 months 1 vs. 2 net change 18.8% (P &lt; 0.001)</td>
</tr>
<tr>
<td>Smith et al.45</td>
<td>1994</td>
<td>Children under 2 years of age; low income families</td>
<td>Before and after study</td>
<td>21</td>
<td>1. Usual care vs. 2. Physician education plus free vaccines and assistance with billing disputes</td>
<td>UTD for age, 1 vs. 2, net change = 25% (P &lt; 0.00001)</td>
</tr>
<tr>
<td>Soljak and Handford41</td>
<td>1985</td>
<td>Clinics and offices, Northland, New Zealand; all children born during study periods</td>
<td>Controlled intervention trial for patient reminders; before and after study for provider reminders</td>
<td>21</td>
<td>1. Retrospective control group vs. 2. GP reminder vs. 3. GP reminder plus postal reminder cards</td>
<td>Up to date ‘with all appropriate antigens’ at 5 months 2 vs. 3 net change 0.4% (p = 0.86); 1 vs. 2 and 3 net change 4% (P = 0.034)</td>
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<tr>
<th>Paper</th>
<th>Study period</th>
<th>Setting and population</th>
<th>Design</th>
<th>Quality</th>
<th>Intervention</th>
<th>Outcome</th>
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</table>
| Szilagyi et al. | 1991–1993    | Teaching hospital paediatric clinic and neighbourhood health centre, Rochester, New York, USA; children aged up to 2 years (mean ages 7–13 months); 1988 participants before randomization; urban; low socioeconomic status | Randomized controlled trial | 24.5    | 1. Usual care (control group)  
2. Provider chart reminders                                                                                                                                                                                   | UTD with DTP/OPV/MMR/Hib, 1 vs. 2 net change = 3% at clinic ($P = 0.3$) and −2% at health centre ($P = 0.5$) |
| Taylor et al.  | Not reported | Private paediatric and family clinics, King County, Washington, USA; children aged 3–19 months                                                                 | Randomized controlled trial | 28      | 1. Immunisation rate feedback and assessment of current immunisation procedures  
2. As above plus peer educational programme                                                                                                                                                           | Immunisation rate 1 vs. 2 net change = 2.2% ($P = 0.24$)                                                                                           |
| Waterman 46   | 1992–1994    | San Diego, California, USA; children aged 2–4 years; low socioeconomic status                                                                 | Controlled intervention trial | 20      | 1. Usual care (control group)  
2. Walk-in clinics plus provider education plus patient reminders plus education and health promotion                                                                                     | DTP/OPV/MMR (4:3:1) 1 vs. 2 net change = 12% (not significant)                                                                                   |
<table>
<thead>
<tr>
<th>Paper</th>
<th>Study period</th>
<th>Setting and population</th>
<th>Design</th>
<th>Quality</th>
<th>Intervention</th>
<th>Outcome</th>
</tr>
</thead>
<tbody>
<tr>
<td>Carter and Jones</td>
<td>1982</td>
<td>Fife, Scotland, UK; children aged 2 years</td>
<td>Before and after study</td>
<td>17.5</td>
<td>1. Usual care vs. 2. Health education programme including mass media campaign and distribution of measles immunisation guidelines and updates</td>
<td>Immunisation with measles vaccine 1 vs. 2 net change 13% (significance not reported)</td>
</tr>
<tr>
<td>Hicks et al.</td>
<td>2002</td>
<td>Community health centre, Northeast Colorado, USA; children aged 13–35 months old; 75% population Latino</td>
<td>Before and after study</td>
<td>21</td>
<td>1. Usual care vs. 2. Reminder cards plus posters in examination rooms</td>
<td>Completely immunised with DTP/IPV/Hib/HepB/MMR/Varicella 1 vs. 2 net change = 12.2% ( (P &lt; 0.05) )</td>
</tr>
<tr>
<td>Melinkovich et al.</td>
<td>1996 (pre) 2006 (post)</td>
<td>Community and school health centres, Denver, Colorado, USA; children aged 12–35 months</td>
<td>Before and after study</td>
<td>19</td>
<td>1. Usual care vs. 2. Postcard reminders plus standing orders and provider education</td>
<td>Immunization rates in 12–23-month-old cohort, 1 vs. 2 net change = 26%; in 24–35 month-old-cohort, 1 vs. 2 net change = 47% (significance not tested)</td>
</tr>
<tr>
<td>Mohr et al.</td>
<td>2000–2001</td>
<td>University of North Carolina paediatric clinic; children under 24 months old; ethnically diverse</td>
<td>Before and after study</td>
<td>19</td>
<td>1. Usual care (control group) vs. 2. Clinical improvement model including parent education and standardizing immunization records</td>
<td>Proportion of children UTD with DTP/Polio/MMR/Hib and HepB vaccines (4:3:1:3:3) at 24 months old; 1 vs. 2 net change 26% ( (P = 0.04) )</td>
</tr>
<tr>
<td>Murphy et al.</td>
<td>1994–1995</td>
<td>General Practice, Dublin, Ireland; children aged between 6 months and 5 years; inner city</td>
<td>Before and after study</td>
<td>22</td>
<td>1. Usual care (control group) vs. 2. Postal reminders plus opportunistic immunisation</td>
<td>Vaccine uptake 1 vs. 2 net change for DTP= 27% ( (P &lt; 0.0005) ), Hib = 43% ( (P &lt; 0.0005) ), MMR 22% ( (P &lt; 0.0005) )</td>
</tr>
<tr>
<td>Oeffinger et al.</td>
<td>Not reported</td>
<td>Family practice residency programme in community and hospital, Texas, USA; children aged under 12 months</td>
<td>Controlled intervention trial</td>
<td>24</td>
<td>1. Usual care (control group) vs. 2. Clinic patient education plus non-specific reminder letter at 2 months old</td>
<td>UTD with 3 doses of DTP/OPV at 12 months, 1 vs. 2 = –4% net change ( (0 = 0.41) )</td>
</tr>
<tr>
<td>Paunio et al.</td>
<td>1982–1986</td>
<td>Community-wide, Finland; children aged 0–11</td>
<td>Time series study 21.5</td>
<td>21.5</td>
<td>1. Usual care vs. 2. Registry plus mass-media reporting local on vaccination rates plus provider reminders plus parent reminders</td>
<td>MMR 1 vs. 2 = 8% net change (significance not tested)</td>
</tr>
</tbody>
</table>

(Continued)
provider strategies (e.g. parental reminders plus provider education). Melinkovich showed when studying community and school health centres in Denver, USA that combining parental and provider strategies led to the greatest point increase in immunisation coverage in the multicomponent strategy group of studies.\textsuperscript{53} The most successful strategies appear to be those that target both the healthcare provider and those that are to be immunised.

Our search methods yielded more studies than any similar previous review. We combined studies from a variety of clinical settings and a range of socioeconomic populations in developed countries, which makes the results generalisable in this setting. Included studies were heterogeneous in setting, service delivery, intervention delivery and quality which made meta-analysis difficult. We did not include studies from developing countries as the barriers to immunisation are different from those in developed countries and include financial barriers that are generally not relevant to parents and general practitioners in many developed countries that offer universal access to primary care services. Grey literature, conference papers and government documents were also included in the review as well as research published in peer-reviewed journals to ensure the broadest possible range of studies and to minimize publication bias. We excluded studies published in languages other than English, which is the main weakness of this study.

We did not examine the effect of GP financial incentives on immunisation uptake. There is a large literature of ‘pay for performance’ in healthcare and this has been a subject of previous systematic reviews, the inclusion of which is outside the scope of this paper.\textsuperscript{59,60}

Various sociodemographic factors have been shown to reduce the likelihood of a child being up to date with recommended vaccinations including being from a lone parent family\textsuperscript{61,62} from an ethnic minority\textsuperscript{63,64} and living in urban areas.\textsuperscript{65} The high level of mobility seen in these populations is thought to contribute to these differences.\textsuperscript{66,67} Interventions to increase vaccination rates have a greater effect on those who are most at risk of being under-immunised. Hence, it is important that vaccine coverage data are collected in a way that highlights differences in uptake rates between socioeconomic and

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**Table 5** Continued

<table>
<thead>
<tr>
<th>Paper</th>
<th>Study period</th>
<th>Setting and population</th>
<th>Design</th>
<th>Quality</th>
<th>Intervention</th>
<th>Outcome</th>
</tr>
</thead>
<tbody>
<tr>
<td>Rossdale et al. \textsuperscript{58}</td>
<td>1982–1985</td>
<td>Well-baby clinic, Bristol, UK; ethnically diverse, low socioeconomic status</td>
<td>Before and after study</td>
<td>20</td>
<td>1. Usual care prior to changes 2. Open access to clinic plus immunization reminder on notes plus 'well baby card' for parents to keep noting due immunizations and recording those received</td>
<td>1 vs. 2 immunised with DTP net change = 11%; Pertussis = 16%; Measles 1% (significance not performed)</td>
</tr>
</tbody>
</table>
ethnic groups. This would help in the implementation and evaluation of public health programmes to improve immunisation rates in the groups in the most need of intervention.

Childhood immunisation can continue to benefit the health of children. For example, the UK has seen a successful campaign to add pneumococcal vaccination to the primary schedule with a catch-up campaign for older children. This has resulted in a substantial reversal of previously increasing trends with falls in hospital admissions for bacterial pneumonia (20%) and empyema (22%) in the 2 years after implementation, linked with uptake rates of 80% and 98% after the first and second years of the campaign, respectively.68

Conclusions

Maintaining high vaccine uptake rates is an essential component of the success of any vaccination programme and in improving the health status of children. Health planners and professionals must engage actively with parents and the public and invest in process measures that ensure children receive primary prevention. Our review has highlighted a number of interventions that can help improve childhood immunisation rates in developed countries. These include reminding parents and providers of upcoming and overdue immunisations and educating and providing feedback to the vaccination providers. Some additional research is required to test the cost-effectiveness of these interventions and their impact in groups with poor immunisation rates or high risks of complications from vaccine preventable diseases.

References

2 Roush SW, Murphy TV. Historical comparisons of morbidity and mortality for vaccine-preventable diseases in the United States. JAMA 2007;298:2155–63
6 Baker D, Garrow A, Shields C. Inequalities in immunisation and breast feeding in an ethnically diverse urban area: cross-sectional study in Manchester, UK. J Epidemiol Community Health 2011;65:346–52
12 Downs S, Black N. The feasibility of creating a checklist for the assessment of the methodological quality of both randomised and non-randomised studies of health care intervention. J Epidemiol Community Health 1998;52:377–84
14 Alemi F, Alemagno S. Computer reminders improve on-time immunization rates. Med Care 1996;34 (Suppl. 10):OS45–51
Primary care strategies to improve childhood immunisation uptake in developed countries

34 Young S, Halpin T, Johnson DA, Irvin JJ, Marks JS. Effectiveness of a mailed reminder on the immunization levels of infants at high risk of failure to complete immunizations. *Am J Public Health* 1980;70:422–4
Appendix 1

<table>
<thead>
<tr>
<th>TERM</th>
<th>LINKING TERM</th>
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</thead>
<tbody>
<tr>
<td>Immunis*</td>
<td>OR</td>
</tr>
<tr>
<td>Vaccin*</td>
<td>OR</td>
</tr>
<tr>
<td>Inoculat*</td>
<td>OR</td>
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<tr>
<td>Tetanus</td>
<td>OR</td>
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<tr>
<td>Diptheria</td>
<td>OR</td>
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<tr>
<td>Polio*</td>
<td>OR</td>
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<tr>
<td>Measles</td>
<td>OR</td>
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<td>Mumps</td>
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<td>Rubella</td>
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<td>Pertussis</td>
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<td>Whooping</td>
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<td>Rates</td>
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<td>Coverage</td>
<td>OR</td>
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<tr>
<td>Uptake</td>
<td>AND</td>
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</table>

Appendix 2


Reporting

1. Is the hypothesis/aim/objective of the study clearly described?
   Yes = 1
   No = 0

2. Are the main outcomes to be measured clearly described in the Introduction or Methods section?
   If the main outcomes are first mentioned in the Results section, the question should be answered no.
3. Are the characteristics of the patients included in the study clearly described?
   Inclusion and/or exclusion criteria should be given.
   Yes = 1
   No = 0

4. Are the interventions of interest clearly described?
   Treatments and placebo (where relevant) that are to be compared should be clearly defined.
   Yes = 1
   No = 0

5. Are the distributions of principal confounders in each group of subjects to be compared clearly described?
   Yes = 2
   Partially = 1
   No = 0

6. Are the main findings of the study clearly described?
   Simple outcome data (including denominators and numerators) should be reported for all major analyses and conclusions. (This question does not cover statistical tests which are considered below).
   Yes = 1
   No = 0

7. Does this study provide estimates of the random variability in the data for the main outcomes?
   In non normally distributed data the interquartile range of results should be reported. In normally distributed data the standard error, standard deviation or confidence intervals should be reported. If the distribution of the data is not described, it must assume that the estimates used were appropriate and the question should be answered yes.
   Yes = 1
   No = 0

8. Have all important adverse events that may be a consequence of the intervention been reported?
   Not going to use this one.
   Yes = 1
   No = 0

9. Have the characteristics of patients lost to follow up been described?
   Yes = 1
   No = 0

10. Have actual probability values been reported (e.g. 0.035 rather than < 0.05) for the main outcomes except where the probability value is less than 0.001?
    Yes = 1
    No = 0

**External validity**

All the following criteria attempt to address the representativeness of the findings of the study and whether they may be generalised to the population from which the study subjects were derived.

11. Were the subjects asked to participate in the study representative of the entire population from which they were recruited?
    The study must identify the source population for patients and describe how the patients were selected.
    Yes = 1
    No = 0
    Unable to determine = 0

12. Were those subjects who were prepared to participate representative of the entire population from which they were recruited?
    The proportion of those asked who agreed should be stated. Validation that the sample was representative would include demonstrating that the distribution of the main confounding factors are the same in the study sample and the source population.
    Yes = 1
    No = 0
    Unable to determine = 0

13. Were the staff, places and facilities where the patients were treated, representative of the treatment the majority of patients receive?
    Yes = 1
    No = 0
    Unable to determine = 0

**Internal validity – bias**

14. Was an attempt made to blind study subjects to the intervention they have received?
    Yes = 1
    No = 0
    Unable to determine = 0

15. Was an attempt made to blind those measuring the main outcomes of the intervention?
Yes = 1
No = 0
Unable to determine = 0

16. If any of the results of the study were based on “data dredging”, was this made clear?
Any outcome that had not been planned at the outset of the study should be clearly indicated.
If no retrospective unplanned subgroup analyses were reported, then answer yes.
Yes = 1
No = 0
Unable to determine = 0

17. In trial and cohort studies, do the analyses adjust for different lengths of follow up of patients?
Where follow-up was the same for all study patients the answer should be yes.
Yes = 1
No = 0
Unable to determine = 0

18. Were the statistical tests used to assess the main outcomes appropriately?
Yes = 1
No = 0
Unable to determine = 0

19. Was compliance with the intervention/s reliable?
Where there was non compliance with the allocate treatment or where there was contamination of one group, the question should be answered no. For studies where the effect of any misclassification was likely to bias any association to the null, the question should be answered yes.
Yes = 1
No = 0
Unable to determine = 0

20. Were the main outcome measures used accurate (valid and reliable)?
For studies where the outcome measures are clearly described, the question should be answered yes. For studies which refer to other work or that demonstrates the outcome measures are accurate, the question should be answered as yes.
Yes = 1
No = 0
Unable to determine = 0

Internal validity – confounding (selection bias)

21. Were the patients in different intervention groups (trial and cohort studies) or were the cases and controls (case control) recruited from same population?
EG patients for all comparison groups should be selected from the same hospital. The question should be answered unable to determine for cohort and case control studies where there is no information concerning the source of patients included in the study.
Yes = 1
No = 0
Unable to determine = 0

22. Were study subjects in different intervention groups (trials and cohort studies) or were the cases and controls (case-control studies) recruited over the same period of time?
For a study which does not specify the time period over which patients were recruited, the question should be answered unable to determine.
Yes = 1
No = 0
Unable to determine = 0

23. Were study subjects randomised to intervention groups?
Studies which state that subjects were randomised should be answered yes except were method of randomisation would not ensure random allocation. For example alternate allocation would score no because it is predictable.
Yes = 1
No = 0
Unable to determine = 0

24. Was the randomised intervention assignment concealed from both patients and health care staff until recruitment was complete and irrevocable?
All non-randomised studies should be answered no.
Yes = 1
No = 0
Unable to determine = 0

25. Was there adequate adjustment for confounding in the analyses from which the main findings were drawn?
This question should be answered no for trials if: the main conclusion of the study were based on analyses of treatment rather than intention to treat; the distribution of known confounders in the different treatment groups was not described, or the distribution of known confounders differed between the treatment groups but was not taken into account in the analyses. In non randomised studies if the effect of the main confounders was not investigated or confounding was demonstrated but no adjustment was made in the final analyses the questions should be answered no.

Yes = 1
No = 0
Unable to determine = 0

26. Were losses of patient to follow up taken into account?
If the number of patient lost to follow up are not reported, the question should be answered as unable to determine. If the proportion lost to follow up was too small to affect the main findings, the question should be answered yes.

Yes = 1
No = 0
Unable to determine = 0

27. Did the study have sufficient power to detect a clinically important effect where the probability value for a difference being due to chance is less than 5%?
Sample sizes have been calculated to detect the difference of x% and y%.

<table>
<thead>
<tr>
<th>Size of smallest intervention group</th>
<th>A</th>
<th>B</th>
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<th>D</th>
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<td>N1-n2</td>
<td>N3-n4</td>
<td>N5-n6</td>
<td>N7-n8</td>
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<td>1</td>
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