

Interventions to Improve Influenza and Pneumococcal Vaccination Rates Among Community-Dwelling Adults: A Systematic Review and Meta-Analysis

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ABSTRACT

PURPOSE Influenza and pneumococcal vaccination rates remain below national targets. We systematically reviewed the effectiveness of quality improvement interventions for increasing the rates of influenza and pneumococcal vaccinations among community-dwelling adults.

METHODS We included randomized and nonrandomized studies with a concurrent control group. We estimated pooled odds ratios using random effects models, and used the Downs and Black tool to assess the quality of included studies.

RESULTS Most studies involved elderly primary care patients. Interventions were associated with improvements in the rates of any vaccination (111 comparisons in 77 studies, pooled odds ratio [OR] = 1.61, 95% CI, 1.49-1.75), and influenza (93 comparisons, 65 studies, OR = 1.46, 95% CI, 1.35-1.57) and pneumococcal (58 comparisons, 35 studies, OR = 2.01, 95% CI, 1.72-2.3) vaccinations. Interventions that appeared effective were patient financial incentives (influenza only), audit and feedback (influenza only), clinician reminders, clinician financial incentives (influenza only), team change, patient outreach, delivery site changes (influenza only), clinician education (pneumococcus only), and case management (pneumococcus only). Patient outreach was more effective if personal contact was involved. Team changes were more effective where nurses administered influenza vaccinations independently. Heterogeneity in some pooled odds ratios was high, however, and funnel plots showed signs of potential publication bias. Study quality varied but was not associated with outcomes.

CONCLUSIONS Quality improvement interventions, especially those that assign vaccination responsibilities to nonphysician personnel or that activate patients through personal contact, can modestly improve vaccination rates in community-dwelling adults. To meet national policy targets, more-potent interventions should be developed and evaluated.

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INTRODUCTION

Influenza and pneumococcal disease are vaccine-preventable causes of morbidity and mortality.¹⁻³ Clinical practice guidelines have recommended routine influenza and pneumococcal vaccinations for elderly and nonelderly high-risk patients.⁴⁻⁶ More recently, influenza vaccinations have been recommended for all individuals older than 6 months.⁷ Even so, vaccination rates remain low.^{8,9}

Studies of interventions for improving adult influenza and pneumococcal vaccination rates are numerous and have been synthesized in several systematic reviews. Jacobson and Szilagyi found that patient reminder and recall systems improved vaccination rates.¹⁰ The US Preventive Services

Task Force's (USPSTF) Community Guide to Preventive Services found supporting evidence for numerous interventions aimed at universally recommended vaccines¹¹ and for combinations of multiple interventions for vaccines targeted to high-risk groups.^{12,13} Stone et al found that interventions involving organizational changes and teamwork were most effective for improving influenza or pneumococcal vaccination rates.¹⁴ Most recently, Thomas et al found evidence of moderate quality that increasing community demand, vaccinating seniors during home visits, and deploying prevention facilitators working with health professionals improved influenza vaccination rates.¹⁵ Though important, these reviews have a variety of limitations. For example, Thomas et al included randomized controlled trials, most of which were graded low in quality. Consequently, the authors were able to recommend only that practitioners implement home visits (2 studies) and practice prevention facilitators (4 studies), to improve vaccination rates.¹⁵ The work of the USPSTF combined many vaccinations for different patient groups under targeted and universally recommended vaccinations.^{11,12} Stone et al, in their review of controlled clinical trials, examined the evidence more than a decade ago. We know of more than 50 additional studies that could be included, today.¹⁴

Because previous reviews may be of limited currency and breadth, we undertook a systematic review and meta-analysis of randomized and nonrandomized studies of the effectiveness of quality improvement interventions for improving adult influenza and pneumococcal vaccination rates in the community. Our review is intended to provide a comprehensive quantitative summary of the results achieved by previous quality improvement studies.

METHODS

Study Selection and Data Extraction

We searched medical literature databases, including MEDLINE, EMBASE, Cochrane Library, Web of Science, and 5 other databases, as well as the reference lists of previous reviews up to August 2010, for relevant studies (Supplemental Appendix A, Exhibit A.2, available at <http://annfamned.org/content/10/6/538/suppl/DC1>).

English language studies published in peer-reviewed journals were included if they involved elderly adults or adults with chronic diseases, involved a quality improvement intervention (see below), featured a parallel control group, and reported influenza or pneumococcal vaccination rates. We focused exclusively on the community setting to maximize relevance to primary care. Studies reporting sufficient data to estimate log odds ratios (ORs) and standard errors were eligible for meta-analysis.

Two reviewers (D.L. and J.H.) selected studies and

extracted data from each study in duplicate. Study quality was measured using the Downs and Black instrument, which assesses both randomized and non-randomized studies on the same items.¹⁶ Disagreements were resolved by consensus; remaining disagreements were resolved by the senior authors (J.A.J. and S.R.M.).

Data Synthesis

We synthesized results by performing random-effects meta-analyses of log odds ratios. We stratified analyses by vaccination type and intervention category. To categorize the interventions, we modified the taxonomy developed by Shojania et al used to classify quality improvement strategies (<http://dx.doi.org/10.1001/jama.296.4.427>).^{17,18} Comparisons were included in meta-analyses if the control group was usual care; a control intervention aimed at nonvaccination behaviors or a different intervention for improving vaccination rates if the intervention was provided to both study arms. When study arms contributed to more than 1 comparison in a meta-analysis, the vaccination rate numerator and denominator were divided among the comparisons to avoid counting patients more than once. We accounted for unit of analysis errors by adjusting standard errors for literature-based values of intraclass correlations.^{19,20} Although we reported all pooled odds ratios, we interpreted only those odds ratios comprised of 3 or more comparisons. Heterogeneity was characterized with I^2 statistics. We explored heterogeneity by substratifying interventions with clear grounds for delineating strata and sufficient studies to divide into strata of 3 or more comparisons. Clinician reminders were stratified according to whether the reminder system was immunization specific or targeted a range of preventive care behaviors, and whether reminders were generated from patients' medical histories. Patient outreach interventions were stratified by communication medium. Finally, team change interventions were stratified by type of personnel involved and whether they administered vaccine independently.

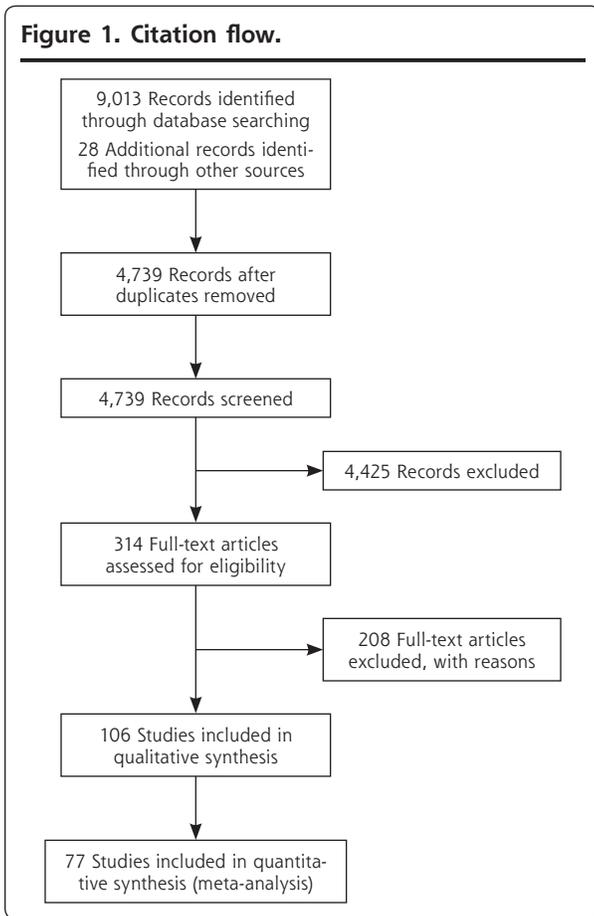
The effects of Downs and Black scores and randomization on pooled odds ratios were examined by meta-regression. Finally, we tested for publication bias by visual inspection of funnel plots and by using Harbord's test.²¹ Harbord's test is an alternative to Egger's test that mitigates false positives in meta-analyses of odds ratios. Analyses were performed using Stata 11 (StataCorp LP).²²

RESULTS

Overview of Studies

We included 106 and excluded 208 citations (Figure 1). Citations were most commonly excluded because

Figure 1. Citation flow.



they lacked a concurrent control group (n = 112, 54%) or took place in hospital or in a nursing home (n = 27, 13%) (Supplemental Appendix A, Exhibit A.3 and A.4). Interreviewer reliability for inclusion of electronic search citations was substantial (91% agreement, $\kappa = 0.8$). The included studies featured 470,175 patients (Table 1 and Supplemental Appendix A, Exhibit A.5). Studies took place primarily in the United States (82 studies), Canada (9 studies), and the United Kingdom (6 studies). A range of settings was represented, including academic primary care practices (41 studies), community practices (21 studies), managed care organizations (13 studies), Medicare-affiliated organizations (11 studies), and Veterans Affairs medical centers (8 studies). A few studies intervened at nonclinical sites, such as senior centers or workplaces. Most stud-

ies targeted the elderly for vaccination, either alone (54 studies), or in combination with high-risk nonelderly patients (27 studies).

Quality of Included Studies

Seventy-seven studies provided sufficient data for meta-analyses of odds ratios (Table 1 and Supplemental Appendix A, Exhibit A.6). Fifty-six studies (75%) were randomized or quasi-randomized controlled trials. The remaining studies were controlled before-and-after (7 studies) and observational (12 studies) designs. The median Downs and Black scores ranged from 14 to 26, with a median score of 21 points (the median Downs and Black scores ranged from 14 to 26, with a median score of 21 points (total possible score = 32, with greater scores indicating higher methodologic quality).

We examined individual items of the Downs and Black instrument (Supplemental Appendix B, <http://annfammed.org/content/10/6/538/suppl/DC1>). The most important weaknesses were unit-of-analyses errors and insufficient reporting and adjustment for potential confounders. We corrected for unit-of-analysis errors in 38 (51%) of studies. Potential confounders included previous vaccination status, health status, and demographic characteristics. The proportion of studies reporting and accounting for these confounders, whether by showing that randomization achieved a balanced distribution of covariates or by statistical adjustment, was 60%. Additional methodological weaknesses were lack of blinding of study subjects or assessors to intervention allocation and contamination, in which the intervention may have affected the treatment of nonintervention patients at



Table 1. Patients, Studies, and Comparisons by Quality Improvement Strategy

Quality Improvement Intervention	No. of Patients	No. of Studies	No. of Comparisons	No. of Comparisons Eligible for Meta-Analysis
Audit and feedback	103,577	13	15	5
Case management	2,924	6	6	4
Clinician education	20,806	18	20	10
Clinician reminders	48,614	40	48	36
Community engagement	23,879	3	3	3
Continuous quality improvement (or similar)	20,097	9	9	3
Delivery site change	35,163	9	12	7
Financial incentive (clinicians)	87,260	4	5	3
Financial incentive (patients)	16,395	4	5	5
Patient outreach	371,218	72	102	71
Team change	155,726	26	28	23
Visit structure change	321	1	1	1
Overall	470,175	106	151	111

the same site. Contamination is prevented by allocation at the physician, practice, or region level, which occurred in only 31 studies (40%).

Main Meta-Analyses

There were 111 comparisons from 77 studies that contributed to the overall meta-analysis (Table 1). The pooled odds ratio expressing the effectiveness of all quality improvement interventions for either vaccination was 1.61 (95% CI, 1.49-1.75; *P* <.001; *I*² = 85%).

Influenza Vaccination

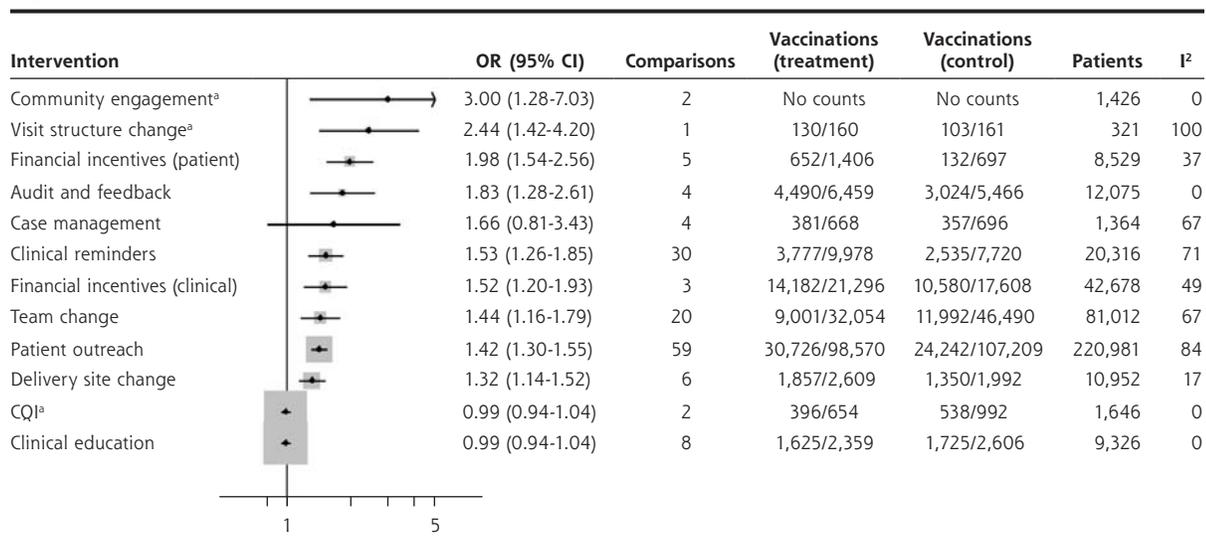
There were 93 comparisons from 65 studies included in meta-analyses for influenza vaccinations. The median treatment and control group vaccination rates were 0.45 ([interquartile range [IQR] = 0.27-0.66), and 0.31 (IQR = 0.20-0.52), respectively. The odds ratio for influenza vaccination, pooled across all interventions, was 1.46 (95% CI, 1.35-1.57; *I*² = 81%). Fewer than 3 comparisons were available for each of community engagement, visit structure change, and continuous quality improvement–like interventions. Excluding these interventions, most components were associated with statistically significant improvements in vaccination rates (Figure 2. See also Supplemental Appendix B, Exhibit B.1 for forest plots featuring individual studies). Interventions featuring patient financial incentives (OR = 1.98, 95% CI, 1.54-2.56; *I*² = 37%) and audit and feedback (OR = 1.83, 95% CI, 1.28-2.61; *I*² = 0%) were effective.

Patient incentives that eliminated out-of-pocket costs in a patient-pay environment^{23,24} appeared to be more effective than those providing a small reward in addition to preexisting third-party vaccination coverage.^{25,26} Insufficient studies were available to test this hypothesis statistically, however. Audit and feedback findings were driven largely by results from the single study by Buffington et al, who were able to improve vaccination rates with regularly updated posters in physician offices tracking vaccination progress.²⁷ Clinician reminders (OR = 1.53, 95% CI, 1.26-1.85; *I*² = 71%), clinician financial incentives (OR = 1.52, 95% CI, 1.20-1.93; *I*² = 49%), team change (OR = 1.44, 95% CI, 1.16-1.79; *I*² = 67%), patient outreach (OR = 1.42, 95% CI, 1.30-1.55; *I*² = 84%), and delivery site changes (OR = 1.32, 95% CI, 1.14-1.52; *I*² = 17%) were also associated with improvements in vaccination rates. Delivery site changes included workplace vaccination clinics²⁸ and clinics in public housing buildings.²⁹ These interventions were effective overall, but what elements of these were effective and where are difficult to discern, because a wide variety of intervention sites were implemented in a small number of studies. Case management and clinician education were ineffective.

Pneumococcal Vaccinations

There were 48 comparisons from 35 studies included in the meta-analyses. The median treatment and control group vaccination rates were 0.19 (IQR = 0.11-0.33), and 0.08 (IQR = 0.04-0.22), respectively. The

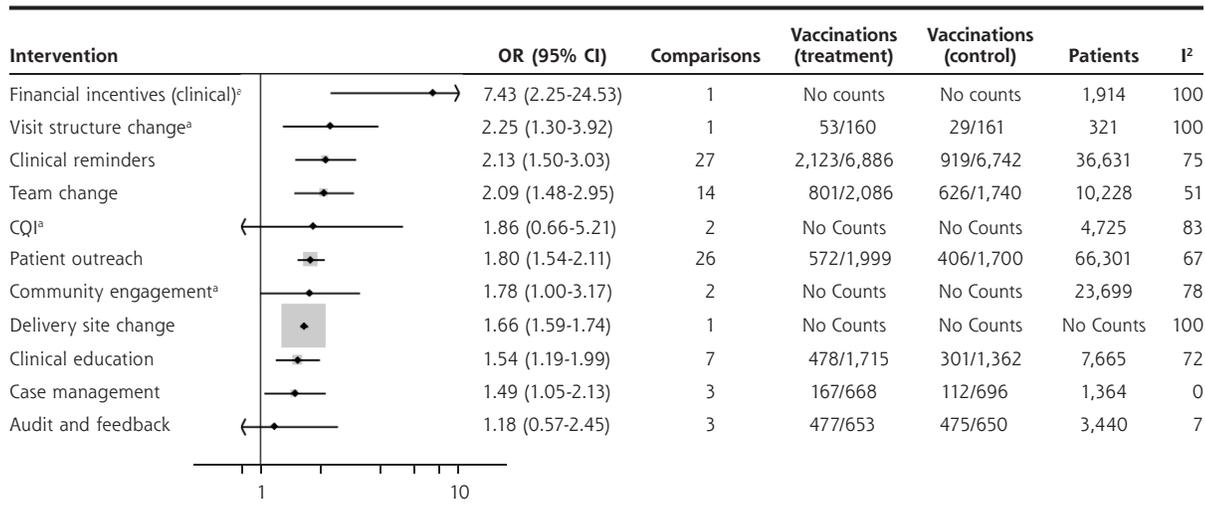
Figure 2. Effect of quality improvement interventions on influenza vaccination rates.



CQI = continuous quality improvement; OR = odds ratio.

Note: Forest plot showing pooled odds ratios from random effects meta-analyses. Vaccination rates provided are crude estimates generated by summing patients among studies. Many studies contributing odds ratios for meta-analysis did not provide crude counts.

^a Pooled odds ratios from fewer than 3 comparisons are reported but considered insufficient for interpretation.

Figure 3. Effect of quality improvement interventions on pneumococcal vaccination rates.

CQI = continuous quality improvement; OR = odds ratio.

Note: Forest plot showing pooled odds ratios from random effects meta-analyses. Vaccination rates provided are crude estimates generated by summing patients among studies. Many studies contributing odds ratios for meta-analysis did not provide crude counts.

^a Pooled odds ratios from fewer than 3 comparisons are reported, but considered insufficient for interpretation. No comparisons involving patient financial incentives were available for meta-analysis.

odds ratio for pneumococcal vaccinations, pooled across all interventions, was 2.01 (95% CI, 1.72-2.36; $I^2 = 72\%$). Three or more comparisons were available for clinician reminders, team change, patient outreach, clinician education, case management, and audit and feedback. Except for audit and feedback (OR = 1.18, 95% CI, 0.57-2.45; $I^2 = 7\%$), these interventions were associated with improvements in vaccination rates (Figure 3. See also Supplemental Appendix B, Exhibit B.2 for forest plots featuring individual studies). Interventions featuring clinician reminders (OR = 2.13, 95% CI, 1.50-3.03; $I^2 = 75\%$), team change (OR = 2.09, 95% CI, 1.48-2.95; $I^2 = 51\%$), and patient outreach (OR = 1.80, 95% CI, 1.54-2.11; $I^2 = 67\%$) had the highest odds ratios. Clinician education (OR = 1.54, 95% CI, 1.19-1.99; $I^2 = 72\%$) and case management (OR = 1.49, 95% CI, 1.05-2.13; $I^2 = 0\%$) were also associated with improvements in pneumococcal vaccination rates.

Meta-Analyses Within Intervention Substrata

Interventions featuring clinician reminders, team change, and patient outreach had moderate to high heterogeneity and sufficient comparisons for substratification. For clinician reminders, most heterogeneity was explained by declining odds ratios with time. For patient outreach and team change, the results of meta-analyses within intervention substrata are presented in Table 2 (forest plots are available in Supplemental Appendix C, at <http://annfamned.org/content/10/6/538/suppl/DC1>). Several findings require

qualification. Among patient outreach strategies for influenza, community media campaigns appeared most effective. This finding should be interpreted cautiously, however, because studies took place in settings with relatively captive audiences and a high prevalence of high-risk patients (eg, seniors' centers).^{29,30} For pneumococcal vaccination, the pooled odds ratio for waiting and examination room posters may also be misleadingly high, because there were few comparisons ($n = 5$), and the highest performing comparisons combined posters with other effective interventions.^{31,32} In 2 studies that considered them alone, waiting and examination room posters were not significantly associated with vaccination rates.^{31,32}

Generally, outreach methods involving personal contact with patients achieved higher pooled odds ratios. For influenza vaccinations, the most effective intervention, excepting community media campaigns, was telephone reminders delivered by clinic staff. For pneumococcal vaccinations, office brochures handed out to eligible patients by clinic staff before their appointments was most effective. Meta-regression detected significant differences between pneumococcal vaccination outreach strategies. Office brochures at the point of care were 3.87 times more effective than mailed reminders, whereas community media campaigns, patient-held preventive care checklists, and waiting or examination room posters were, respectively, 0.85, 0.77, and 0.75 times less effective than mailed reminders. Among team change interventions

Table 2. Results of Meta-Analyses Within Substrata of Patient Outreach and Team Change

Influenza Vaccination	Pooled OR (95% CI)	I ²	Pneumococcal Vaccination	Pooled OR (95% CI)	I ²
Patient outreach medium			Patient outreach medium		
Community media campaign ^a	3.16 (1.35-7.37)	0	Emergency medical technician outreach ^b	8.65 (0.02-4899.87)	100
Telephone reminders ^a	2.74 (1.23-6.12)	67	Brochures at office visit ^a	5.86 (3.29-10.44)	0
Waiting/examination room posters ^b	1.78 (0.53-6.01)	95	Telephone reminders ^b	2.86 (2.31-3.56)	0
Mailed print material ^a	1.45 (1.30-1.61)	89	Waiting/examination room posters ^a	1.92 (1.09-3.40)	57
Brochures at office visit ^b	1.38 (0.82-2.33)	0	Mailed print material ^a	1.66 (1.59-1.74)	0
Patient-held preventive care schedule	1.28 (0.82-1.99)	53	Home visit education ^b	1.52 (0.74-3.11)	100
Home visit education ^b	0.94 (0.64-1.40)	0	Community media campaign ^a	1.31 (1.28-1.55)	0
Emergency medical technician outreach ^b	0.67 (0.01-36.06)	100	Patient-held preventive care schedule ^a	1.29 (1.06-1.57)	0
Team change: type of additional personnel			Team change: type of additional personnel		
Multidisciplinary team ^b	2.44 (1.42-4.20)	100	Emergency medical technician	8.65 (0.02-4899.87)	100
Nurse, autonomous vaccinations ^a	1.63 (1.30-2.04)	7	Nurse, autonomous vaccinations ^b	7.03 (2.98-16.57)	0
Nurse, no autonomous vaccinations	1.14 (0.88-1.48)	60	Multidisciplinary team ^b	2.25 (1.30-3.92)	100
Pharmacist	1.11 (0.62-1.98)	0	Nurse, no autonomous vaccinations ^a	1.96 (1.28-3.03)	60
Emergency medical technician ^b	0.67 (0.01-36.06)	100	Pharmacist	1.03 (0.62-1.74)	0

OR = odds ratio.

Note: See forest plots in Supplemental Appendix C for more detail.

^a Pooled odds ratios significant at $P < .05$.^b No. <3. Although we avoided interpreting pooled odds ratios from fewer than 3 studies, they are presented here for completeness.

for influenza vaccinations, we found that having nurses assume responsibility for administering vaccinations was effective, whereas interventions in which nurses or pharmacists assessed patients and reminded physicians, but did not themselves administer vaccinations, were ineffective. We were unable to examine this relationship in studies of pneumococcal vaccinations because of insufficient comparisons.

Numbers Needed to Treat

Results for effective quality improvement strategies are summarized as numbers needed to treat (NNT), assuming baseline levels of vaccination similar to those reported in community studies of elderly adults⁸ (Table 3 and Supplemental Appendix C, Exhibit C.5).

Sensitivity Analyses and Publication Bias

Randomized study design was not significantly associated with study odds ratios within intervention strata. After excluding 2 clear outlier studies,^{33,34} quality score was also not significantly associated with study odds

ratios for any intervention. Funnel plots showed higher odds ratios in smaller studies. Harbord's test was positive for small study effects among studies of patient outreach for influenza and pneumococcal vaccinations and of team change for influenza vaccinations. These findings suggest potential publication bias.

DISCUSSION

We reviewed the evidence for effectiveness of quality improvement interventions for increasing influenza and pneumococcal vaccination rates. Most interventions were associated with modest improvements in vaccination rates.

Team change, patient outreach, and clinician reminders were effective for both influenza and pneumococcal vaccinations. We found that interventions involving team change were effective, especially where nurses had been assigned responsibilities for administering vaccine. Configuring additional personnel so that they are able relieve physicians of vaccinations

seems important to successful team change.¹⁴ Additionally, patient outreach may better increase vaccinations to the extent that direct personal contact is achieved. A previous review has similarly reported that reminders involving person-to-person telephone contact were most effective.¹⁰

Clinician reminders and education were associated with greater improvements for pneumococcal than for influenza vaccinations. Awareness and support may be less common for pneumococcal³⁵ than for influenza vaccinations,³⁶⁻³⁸ making pneumococcal vaccinations relatively low hanging fruit. Audit and feedback appeared effective for influenza, but not pneumococcal, vaccinations. Audit and feedback may have been effective for influenza vaccinations because of the prominent tracking posters used in Buffington et al.²⁷ The use of materials with high visual appeal and clarity has been previously associated with increased

vaccination rates.¹⁴ Clinician and patient financial incentives were both effective for influenza vaccinations, but they could not be evaluated for pneumococcal vaccinations. The 2 successful studies of patient financial incentives took place in out-of-pocket payment environments.^{23,24} Where demand for vaccinations is not pent up by inability to pay, the benefit of patient incentives may be smaller.^{25,26} Case management, surprisingly, was not very effective—possibly because case managers may have prioritized other disease-related process of care.

Several limitations of our review should be borne in mind. Our funnel plots and associated tests suggested publication bias, which may have led our pooled odds ratios to be overly optimistic. Our review also did not address the economic value of the interventions. Additionally, the included studies may not generalize well to nonelderly adults or adults not in a physician's care, for whom vaccination recommendations have recently been expanded.⁷

More importantly, we have taken a highly inclusive approach toward meta-analysis. There are 2 major limitations of this approach. First, our analysis of Downs and Black items identified a high prevalence of design or reporting flaws in the included studies. Lack of blinding may be relatively unimportant for quality improvement interventions designed to act, in part, by increasing awareness of vaccinations and for outcomes that can be measured relatively objectively by reviewing charts or billing data. Only 60% of studies reported and accounted adequately for potential confounders, however. This proportion was higher in randomized than in observational studies.

We have nonetheless reported odds ratios pooled from all studies. Neither randomization nor Downs and Black scores were associated with significant differences in odds ratios. The inclusion of a wide range of studies allowed us to produce quantitative summaries for many intervention categories. In particular, interventions requiring policy support or action on a community scale, such as audit and feedback and community media campaigns, are difficult to randomize—observational studies comprise an important source of insight.³⁹ Our study quality tables (Supplemental Appendix B) provide further detail on methodological issues for potential users.

Second, many of our pooled estimates contained residual heterogeneity. Our ability to explore heterogeneity was limited by lack of evidence.⁴⁰ For example, reasons for decreases in the effectiveness of clinician reminders in recent years are unknown. We have incorporated heterogeneity into our meta-analysis by using a random-effects approach. Users should interpret pooled odds ratios as estimates of the average

Table 3. Numbers Needed to Treat to Obtain an Additional Vaccination

Characteristics	NNT
Influenza vaccinations	
Baseline vaccination rate of 70% assumed	
Patient outreach (community media)	6
Patient outreach (telephone reminders)	6
Financial incentives, patient	8
Audit and feedback	9
Team change (nurse vaccine administration)	11
Clinician reminders	12
Financial incentives, clinician	13
Patient outreach (mailed print materials)	14
Team change (overall)	14
Patient outreach (overall)	15
Delivery site change	18
Pneumococcal vaccinations	
Baseline vaccination rate of 60% assumed	
Patient outreach (brochures handed out before appointments)	3
Clinician reminders	6
Team change (overall)	6
Team change (nurses without vaccine administration responsibilities)	7
Patient outreach (waiting/examination room posters)	7
Patient outreach (overall)	8
Clinician education	9
Patient outreach (mailed print materials)	9
Case management	11
Patient outreach (community media)	13
Patient outreach (preventive care checklists)	17

NNT = number needed to treat.

Note: Interventions included in this table had summary odds ratios statistically greater than 1.0 ($P < .05$) based on 3 or more studies. Numbers needed to treat are provided assuming other baseline vaccination rates in Supplemental Appendix C, Exhibit C.5.

intervention effect, as opposed to a single, true effect. Our 95% confidence limits may provide bounds on the expected performance of the intervention under most circumstances. In any event, a single true effect would not likely be useful, because most users can identify mitigating or potentiating factors unique to their circumstances. Our estimates provide a preliminary basis for selecting interventions; potential users should examine our summaries of individual studies (Supplemental Appendix A, Exhibit A.5) and intervention-specific forest plots (Supplemental Appendix B) in light of their own circumstances and a theoretical understanding of behavior change.^{41,42}

Building on previous reviews, we have produced a comprehensive, quantitative summary of the effectiveness of interventions to improve influenza and pneumococcal vaccination rates. Our results suggest that (1) shifting vaccine administration from physicians to members of the primary care team with clear responsibilities for chronic and preventive care and (2) activating patients through personal outreach may stand the best chance of improving vaccination rates in community dwelling adults. Nonetheless, practitioners and policy makers should temper their expectations of quality improvement interventions. In few treatment arms had vaccination rates improved sufficiently to meet national policy targets.^{43,44} Further research is required to develop and evaluate more potent approaches and to better understand how and why they work.

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