Delivery of patient adherence support: a systematic review of the role of pharmacists and doctors

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Abstract: We conducted a systematic review of adherence support programs involving doctors and pharmacists. We searched MEDLINE®, Embase, International Pharmaceutical Abstracts, PsycINFO®, and CINAHL using the keywords “pharmacist” or “doctor” and “adhere*” or “compli*” and “randomized controlled trials”. We found 89 studies involving pharmacists; in contrast, only 14 studies involved doctors. The roles of pharmacists and doctors ranged from providing education and counseling to adjusting treatment. Most interventions that specified a patient group were carried out with patients with chronic conditions (n=79) and only six included short-term treatments. The majority of interventions improved adherence and clinical outcomes to some extent, although the size of effect size was sometimes small. Resource utilization (eg, hospitalization rates, visits to doctors) did not change in the majority of studies that reported it. Few studies included cost analyses. All but one study had high risk of performance bias due to the nature of the interventions, which made it impossible to blind the participants. The majority of studies did not report tailoring the interventions to patient needs and the vast majority of papers did not report taking a concordant patient-centered approach or considering patients’ own views and experiences when providing adherence support. In addition, the majority of studies did not describe training for the health care professionals involved in providing adherence support. Providing training for doctors and pharmacists to take a more patient-centered concordant approach would be expected to increase the effectiveness of adherence support further.

Keywords: interventions, patient views, patient experiences, patient-centered approach, resource utilization

Introduction to patient adherence support and programs

One of the biggest challenges in health care worldwide is to ensure that patients are both willing and able to take their treatments as prescribed, and that they persist in doing so over the designated amount of time. When they do not do this, it is most often referred to as “treatment nonadherence” or “treatment noncompliance”, and the consequences can be severe. Nonadherence to appropriately prescribed treatments may lead to reduced clinical benefit and increased risk of morbidity and mortality.1 For example, it has been estimated that nonadherence is responsible for 48% of asthma deaths, an 80% increased risk of death in diabetes and a 3.8-fold increased risk of death in the year following a heart attack.2 The related economic burden on health care systems is significant. In the UK, the cost of unused and unwanted medicines has been estimated to reach £300 million per year,3 and, in the USA, nonadherence has been estimated to cost the US health care system US$310 billion (note US billion =109) annually.4
Nonadherence is prevalent in all disease categories and similar rates of nonadherence are found in chronic conditions such as diabetes (33%) and skin disorders (23%) as in life-threatening conditions such as cancer (20%) and end-stage renal disease (30%). However, rates may vary between patients as well as within individual patients across different treatments and over time. The rate of nonadherence that is clinically significant also varies between diseases. Typically, lifestyle recommendations such as diet (average percentage of patients that were nonadherent across studies 41%) and exercise (average percentage of patients nonadherent 28%) display more adherence challenges than pharmaceutical medicines (average percentage of patients that were nonadherent 21%). Asymptomatic patients have been found to be less adherent than symptomatic patients in, for example, HIV. It is important to point out that differing adherence definitions and measurements used across studies also influence the adherence rates captured.

The reasons for nonadherence are numerous and multifaceted, and the patients’ relationship and communication with their health care providers are key factors that influence both their motivation and their ability to adhere. The patient may decide not to take their treatment as prescribed (“intentional nonadherence”) for several different reasons. For example, the patient may experience side-effects that they are not prepared to tolerate or may have concerns over the long-term effects of taking the medication, while other patients may not fully appreciate the necessity or effectiveness of the medication to manage their condition. Other patients may not be able to take their treatment as prescribed, even though they intend to do so (“unintentional nonadherence”). Forgetting is the most common reason patients’ give for missing doses unintentionally, but problems such as difficulties with dexterity and swallowing, reduced access to medication, and high prescription co-payments can also hinder treatment adherence. It is worth bearing in mind that forgetting is an example of unintentional nonadherence that might be influenced by intentional factors, such as perceived need for treatment and concerns about treatment.

Both international and national policy drivers, such as the World Health Organization and National Institute for Health and Care Excellence in the UK, have called for increased patient involvement in treatment decisions to improve adherence and health care outcomes. The term “concordance” can be used to emphasize the shared decision-making process between patient and health care provider that should ideally be achieved when prescribing treatments. A scoping review of concordance, adherence, and compliance in medicine taking noted that concordance initially focused on the consultation process, in which doctor and patient agree therapeutic decisions that incorporate their respective views and moved to a wider concept that stretches from prescribing communication to patient support in medicine taking. In the concordance model, a decision not to take a medicine is an acceptable and good outcome.

In addition, numerous adherence support programs have been developed over the years. Key criteria for success are that interventions are appropriate for implementation in practice and that they are cost-effective. To ensure robust development, implementation, and evaluation of interventions, it has been recommended that development is theory driven and that the behavioral-change techniques (BCTs) that are used is clearly defined.

A number of large-scale reviews have evaluated the evidence around adherence support programs. For example, Haynes et al reviewed randomized controlled trials that measured both adherence and clinical outcome. Haynes et al found 44% and Kripanali et al found 54% of interventions successful at increasing adherence. However, Haynes et al noted that “even with the most effective methods for long-term treatments, improvements in drug use or health were not large”. Kripanali et al and Haynes et al concluded that complex interventions were more likely to be effective, for example, by including combinations of more convenient care, information, reminders, self-monitoring, reinforcement, counseling, family therapy, psychological therapy, crisis intervention, manual telephone follow-up, and supportive care (although they could not determine the relative importance of the individual components). There was also no consensus regarding whether motivational, behavioral, or combined approaches were preferable.

The literature has not been able to provide guidance regarding what specific theoretical models and BCTs are needed to improve the effectiveness of adherence interventions, but there are some indications of factors that may increase effectiveness. Kripanali et al concluded that behavioral interventions that reduced the dosing demands of therapies consistently improved adherence with a large effect size. Haynes et al argued that interventions delivered by allied health care professionals (HCPs) such as nurses and pharmacists were worthy of future research. In addition, others have found that tailoring interventions to individual patients’ needs, rather than using a “one-size-fits-all” approach, is likely to be more effective. Interventions that are tailored to individual needs can, for example, distinguish...
between a patient’s lack of motivation to take medication and practical barriers to taking medication, as well as address an individual’s unique mix of factors in order to improve adherence. Lastly, a recent meta-analysis found that cognitive-based BCTs are effective at improving adherence, can be effectively delivered by routine HCPs, and can have effects that go beyond those achieved by educational or behavioral interventions.15

Some more recent reviews have focused on particular modes of delivery. For example, Rubio-Valera et al17 and Morgado et al18 reviewed pharmacist-led adherence support programs for patients prescribed antidepressants and antihypertensives, respectively, and these interventions were deemed promising and worthy of further study. Pharmacists may be considered particularly suited to deliver adherence interventions given their responsibility to monitor and optimize patients’ pharmacological treatment as part of providing pharmaceutical care.19 Cutrona et al20 reviewed interventions for cardiovascular medication in which the patient’s doctor was involved, but the interventions appeared to be less effective when a doctor was involved than when other HCPs were involved.20

Due to the promising findings regarding pharmacist-led interventions from these disease-specific reviews, one of the aims of the current review was to expand the scope and review the evidence regarding pharmacist-led adherence support programs across diseases categories. Second, despite the negative impact of physician involvement in adherence support programs for cardiovascular medications found by Cutrona et al,20 it was deemed important to include a wider search for evidence from such interventions in other populations. The role of nurses was beyond the scope of this project.

Methods
Search strategy
We carried out a systematic review to evaluate delivery of patient adherence support involving pharmacists and/or doctors. We included evidence related to all aspects of patient adherence (eg, adherence to medication, diet, lifestyle changes, screening procedures).

SG and LE developed a search strategy and performed an electronic search of the following databases: MEDLINE® (1946 to September 6, 2013), Embase (1980 to August 2013), International Pharmaceutical Abstracts (1970 to August 2013), PsychINFO® (1806 to the first week of September 2013), and CINAHL (September 9, 2013). We used the keywords “pharmacist” or “doctor” or “physician” and “adhere*” or “compli*” and “randomized controlled trials”. MA also hand searched the bibliographies of included papers and obtained the full text of any original studies that potentially met the inclusion criteria.

Inclusion and exclusion criteria
Inclusion criterion
The inclusion criterion was RCTs of interventions to improve adherence in which pharmacists and/or doctors had a defined role in terms of delivery. All countries and settings were included.

Exclusion criteria
The exclusion criteria were: study designs other than RCTs, interventions without a defined role for pharmacists and/or doctors, papers not published in English, conference abstracts, and protocols only.

Screening and data extraction
All database search results were combined into a Reference Manager® (v 11; Thomson Reuters, New York, NY, USA) database. An electronic duplicate search was conducted using Reference Manager followed by a manual duplicate search. All duplicate papers were removed. MA then screened each title and abstract to determine whether the full research paper should be retrieved or whether it was evident it did not meet the inclusion criterion at that stage. SG independently screened a random 10% sample of abstracts to check the reliability of the screening process (agreement level 90%). All discrepancies were resolved through discussion. MA then reviewed all retrieved and screened full manuscripts to determine whether each article met the inclusion criterion and SG independently reviewed a further random 10% sample of full papers to check reliability (agreement level 96%). MA then extracted data from the included articles regarding pharmacists’ and/or doctors’ delivery of support on patient adherence. SG then independently reviewed a random 10% of the data extraction table to check reliability (agreement level 90%).

The following data were extracted directly into electronic tables: study author and year, the country in which the research was carried out, sample size, clinical diseases of interest, the nature of the intervention (including whether it was tailored to each patient or fixed for all participants), the setting and the HCP involved, training for HCP, the HCP’s involvement in the intervention, measured outcomes, and the effectiveness of the intervention.

To assess the quality and the risk of biases of the included papers, the Cochrane Collaboration’s tool for assessing risk of
bias in randomized trials was used.\textsuperscript{21} MA performed a quality assessment of the included papers, and SG conducted a 10% reliability check (agreement level 94%). All discrepancies were resolved through discussion.

**Results**

**Overview**

A total of 1,031 abstracts were screened and 219 full-text articles were obtained. Of these, 103 met the inclusion criterion\textsuperscript{22–124} (Figure 1). Details of the included studies are shown in Supplementary Table S1 and Table S2.

**Quality assessment**

The full quality-assessment results are found in Supplementary Table S2. Of the 103 studies, 13 were found to have low risk of bias in all domains (selection bias, detection bias, attrition bias, selective reporting bias, and other bias) except for the risk of performance bias. Only three studies had high risk of bias in at least four of the six domains.\textsuperscript{25,78,83}

**Performance bias**

All but one study had high risk of performance bias. The high risk of performance bias was due to the nature of the interventions, which made it impossible to blind the participants. The only study that did not have high risk of performance bias had an unclear risk, because the patients were blinded but the physicians were not.\textsuperscript{29}

**Selection bias**

The risk of bias when generating the random sequence was low in 50 studies and high in 14 studies. In 39 papers, the method of generating a random sequence was not described, making the risk of selection bias unclear. The risk of selection bias when concealing the allocation was low in 44 studies, high in 47 studies, and unclear in 12 studies.

**Detection bias**

In 26 studies, the risk of detection bias was low. In 69 studies, information regarding blinding of outcome measurements was insufficient, making the risk of detection bias unclear, and, in eight of the studies, the risk was high.

**Attrition bias**

The majority of studies had a low risk of attrition bias. Conversely, 17 studies were found to have a high risk of attrition bias, and, in 14 studies, the information provided was insufficient to decide the risk of attrition bias.

**Selective reporting bias**

In 99 studies, no selective reporting could be identified, indicating low risk of reporting bias. However, in one study, some results were only compared within the groups and not between them.\textsuperscript{109} In three studies (3%), data were incomplete.\textsuperscript{42,58,103}

![Flow chart of papers identified, screened and evaluated.](https://www.dovepress.com/)

**Figure 1 Flow chart of papers identified, screened and evaluated.**

**Abbreviation:** RCT, randomized controlled trial.
Other risks of bias occurred when there were differences between the intervention group and control group at baseline. This type of bias occurred in 29 studies. Examples were difference in age of patients declining to participate and those who consented,\(^5\) dropout rate,\(^6\) adherence,\(^7\) blood pressure,\(^8\) and comorbidity.\(^9\)

**Key findings**

**The HCP delivering the intervention**

Only 14 of the interventions were delivered by doctors.\(^{29,35,38,53,61,64,72,87,95,96,98,110}\) The rest of the interventions were delivered by different types of pharmacists (clinical, hospital, community, research, senior, or unspecified).

**Settings**

In 52 papers, the setting was described as special clinics or practices, and, in 24 papers, the studies were carried out at hospitals. Eighteen studies were carried out at community pharmacies.

A total of 44 studies were carried out in the USA, with the remaining studies having been conducted in a range of international countries, including those in Europe, Africa, Asia, Canada, and Australia.

**Clinical diseases**

The most common diseases targeted in the interventions were hypertension (16 studies), type II diabetes (eleven studies) and heart failure (seven studies). In eleven studies, there was no specific disease group studied. Most interventions were carried out with patients with chronic conditions and only a few included short-term treatments—for instance, *Helicobacter pylori*-infected ulcers or other infections.\(^{24,34,57,78,108}\)

**Training received by the HCP**

In the majority of studies, no additional training for the HCP was described (63%). However, when additional training was reported (37% of the studies), it included educational workshops on the specific disease, training on the intervention protocol or relevant guidelines, or interview training. In 5% of the studies, the authors described providing specific training in patient-centered approaches, respecting patients’ wishes, or helping patients set goals for their treatment.\(^{30,65,82,90,96}\) In one study, the authors explicitly stated that the HCP did not receive additional training.\(^{42}\)

**The role of the HCP in the intervention**

When pharmacists were involved in the delivery of the intervention, in 93% of cases, they carried out the full intervention. In contrast, it was more common for doctors to deliver only part of the intervention (64% of the studies in which doctors were involved), when compared with interventions delivered by pharmacists. The pharmacists often had a role as a patient educator, providing either tailored or fixed patient education, as well as making treatment recommendations. In the studies in which the doctors delivered the intervention, they gave advice and adjusted patients’ treatments, and also provided patient education. In four studies, the doctor delivered the intervention in collaboration with other HCPs.\(^{53,64,77,87}\)

**Tailored versus fixed interventions**

Fifteen of the interventions were fixed. These included structured education,\(^{37,38}\) written information,\(^{96}\) mailed letters,\(^{77,95,111}\) videotapes, or telephone calls.\(^{51}\) In six studies, several methods were used.\(^{25,29,31,35,45}\) In one study, the patients were given a blood-pressure measuring device to take home and instructions on how to use it.\(^{44}\) Another study included a demonstration dose of sublingual nitroglycerin, for which the physician was present.\(^{72}\)

Forty percent of the interventions were tailored. These included tailored patient counseling,\(^{22,58}\) education,\(^{40,75,87,92,111,118}\) assessment and instructions,\(^{35,48,57}\) or a combination of different approaches.\(^{67,78,109}\) In the studies in which the interventions were tailored, the HCP tailored the interventions based on patients’ needs, giving tailored information and advice and prioritizing the recommendations based on the patients’ medical records.\(^{62,107}\) Hederos et al\(^{64}\) used the concept of “concordance” (defined as reaching an agreement with parents of pediatric patients on how to look upon asthma and its management). They used this approach with the hypothesis that the parents would understand the advice and education better. Personalized medication adherence plans, individual goal values, and identification of adherence barriers were evaluated and used in five studies.\(^{28,48,52,61,82}\) In total, 14 studies explicitly reported using some method of tailoring the intervention by taking the patients’ own views into account rather than simply the HCP’s viewpoint.\(^{28,30,36,47,48,50,52,61,64,65,82,90,96,120}\)

In 46% of the studies, it was unclear to what extent the interventions were tailored, as the interventions were not described in sufficient detail.

A greater proportion of studies carried out in the last 15 years than older studies reported being tailored to patients’ needs (46% versus [vs] 6%). All the studies reporting the use of a more patient-centered concordant approach were carried out in the last 15 years. However, in the last 15 years, there was still a higher proportion of studies that did not describe
tailoring the intervention to individual patients’ needs than those that did.

Effectiveness of the patient adherence support provided by pharmacists and doctors

Short-term versus long-term treatments
The effectiveness of interventions for studies focusing on short- and long-term treatments were similar. For short-term treatments, interventions showed benefits in five studies, and one paper failed to show any benefits other than patient satisfaction with pharmacy service. For research focusing on long-term treatments, interventions showed benefits in 64 out of 79 (81%) studies. Consequently, 15 studies failed to show any improvement.

A range of different outcomes was used in measuring the effectiveness of the adherence support provided. These included medication adherence, attending screening appointments, clinical outcomes, quality of life (QoL), medication knowledge, patient satisfaction, resource utilization, and cost analyses. The effects of the adherence intervention on these outcomes are summarized in Table 1.

Medication adherence
Medication adherence was measured in several different ways, including pill counts, serum concentrations, Medication Event Monitoring System, self-report, questionnaires, and pharmacists’ refill records. The interventions statistically significantly enhanced adherence in 68 studies. However, 35 interventions failed to show statistically significant improvement in adherence. Some interventions did not result in a statistically significant change in adherence but did result in a statistically significant improvement in clinical outcomes – for example, blood pressure and fasting blood glucose. When differences in adherence were significant, a range of effect sizes was seen.

Where adherence was measured dichotomously and patients classified as either adhering or not adhering to treatment, in 16 of 41 (40%) studies, the number of patients in the intervention group adhering was double or more than double the number in the control group, with the remaining studies (25/41, 60%) showing more modest effects. When adherence was measured as an adherence rate, the mean adherence rates were between 3% and 20% greater in the intervention group than in the control group.

Inhaler technique was measured in six out of eleven studies regarding asthma, with improvement being demonstrated in five of them. Inhaler technique was not measured in any of the studies regarding chronic obstructive pulmonary disease.

Screening appointments
Seven studies concerned adherence to cancer screenings. The authors recorded patients’ adherence to screening appointments to evaluate the effectiveness of the interventions. One study measured attitude toward the screening tests and did not find any statistically significant difference between the intervention group and control group. Of the six remaining, five showed a statistically significant improvement in screening rates, and only one paper failed to show any improvement.

Clinical outcomes
A range of clinical outcomes was assessed including: blood pressure, lipid levels, HbA1c levels, severity of symptoms, and adverse drug events (ADEs).

Blood pressure was an outcome measure in 21 papers. Blood pressure was statistically significantly improved in 16 (76%) of them. Inhaler technique was measured in six out of eleven studies regarding asthma, with improvement being demonstrated in five of them. Inhaler technique was not measured in any of the studies regarding chronic obstructive pulmonary disease.

Table 1 Effectiveness of the interventions

<table>
<thead>
<tr>
<th>Outcome</th>
<th>Papers measuring the outcome (n=103)</th>
<th>Papers showing statistically significant benefits/improvements, n (%)</th>
<th>Papers failing to show statistically significant benefits/improvements, n (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Medication adherence</td>
<td>103</td>
<td>68 (66)</td>
<td>35 (34)</td>
</tr>
<tr>
<td>Cancer screening rates</td>
<td>6</td>
<td>5 (83)</td>
<td>1 (17)</td>
</tr>
<tr>
<td>Quality of life</td>
<td>23</td>
<td>16 (70)</td>
<td>7 (30)</td>
</tr>
<tr>
<td>Blood pressure</td>
<td>21</td>
<td>16 (76)</td>
<td>5 (24)</td>
</tr>
<tr>
<td>Lipid values</td>
<td>13</td>
<td>7 (54)</td>
<td>6 (46)</td>
</tr>
<tr>
<td>HbA1c</td>
<td>9</td>
<td>8 (89)</td>
<td>1 (11)</td>
</tr>
<tr>
<td>Severity of symptoms</td>
<td>15</td>
<td>2 (13)</td>
<td>13 (87)</td>
</tr>
<tr>
<td>Adverse drug reactions</td>
<td>8</td>
<td>1 (13)</td>
<td>7 (87)</td>
</tr>
<tr>
<td>Medication knowledge and understanding</td>
<td>27</td>
<td>20 (74)</td>
<td>7 (26)</td>
</tr>
<tr>
<td>Resource utilization</td>
<td>19</td>
<td>3 (16)</td>
<td>16 (84)</td>
</tr>
<tr>
<td>Cost analyses</td>
<td>5</td>
<td>4 (80)</td>
<td>1 (20)</td>
</tr>
</tbody>
</table>

Abbreviation: HbA1c, hemoglobin A1c.
Thirteen papers measured patients’ lipid levels, but the majority (seven) failed to show any significant difference.\textsuperscript{33,51,52,65,82,93,106}

Nine papers measured HbA\textsubscript{1c} levels.\textsuperscript{42,44,65,68,70,83,94,106} All but one of them\textsuperscript{89} demonstrated a statistically significant improvement in the intervention group.

Severity of symptoms was measured in 15 papers, and only two studies (13\%) showed statistically significant changes in this outcome.\textsuperscript{83,89}

Eight interventions used ADEs as an outcome measure. Seven (87\%) of them failed to show any statistically significant benefit, and the other one detected a statistically significant improvement in one arm of the study.\textsuperscript{85} Murray et al\textsuperscript{85} investigated the effect of adherence support on patients with complicated and uncomplicated heart failure and/or hypertension, and found a statistically significant reduction in ADEs in the complicated stratum but not in the uncomplicated stratum.

QoL
QoL was used as an outcome measure in 24 of the studies. Improvement was noted in nine (38\%) of them\textsuperscript{27,67,68,83,99,100,104,109,112} and 15 (62\%) failed to show any improvement.\textsuperscript{30,37,47,62,66,70,72,84,86,87,90,113,114,117,118}

Medication knowledge and understanding
Medication knowledge and medication understanding were used as measures in 27 studies. Medication knowledge was statistically significantly improved in 20 (74\%) of the studies that used it as an outcome measure.\textsuperscript{22,25–27,30,31,57,62,68,70,74,80,81,83,86,88,91,97,99,102,103,109,112,119} and seven (26\%) failed to show statistically significant improvement.\textsuperscript{26,30,62,86,103,109,119}

Patient satisfaction
Sixteen papers measured patient satisfaction. In nine of the studies (56\%), satisfaction increased as a result of the intervention.\textsuperscript{25,26,36,52,75,90,108,112,118} and, in seven studies (44\%), patient satisfaction did not statistically significantly change.\textsuperscript{34,40,58,62,67,91,113}

Resource utilization
A number of outcome measures were related to resource utilization, including hospitalization rates, visits to doctors, and visits to emergency rooms. Nineteen papers investigated patients’ resource utilization; a decrease was found in three (16\%) studies,\textsuperscript{22,112,121} with the remaining 16 (84\%) studies not finding any change.

Cost analysis
Six studies also did some form of cost analysis as a part of evaluating the effectiveness and feasibility of the interventions. One study\textsuperscript{46} was cost-effective and another showed a gain of 42.2 working days.\textsuperscript{84} Beaucage et al calculated that each intervention would cost US$3.74.\textsuperscript{34} Another study aimed to calculate total direct health care costs, but the variation in costs was too large for a comparison to be possible.\textsuperscript{90} In a fifth study, an economic evaluation of the intervention was conducted, determining that the intervention group had a statistically significantly lower cost to the National Health Service (NHS) than the control.\textsuperscript{90} Zhang et al did not find any difference in cost of drugs or hospitalization between the intervention and control group.\textsuperscript{124}

Discussion
Evaluation: effectiveness of the role played by pharmacists and doctors in providing patient adherence support
Involving pharmacists and doctors in adherence support appears to improve outcomes to some extent, although the size of effect is sometimes small. Of the interventions involving pharmacists and doctors, 66\% led to some improvements in medication adherence, while 73\% led to some improvements in clinical outcomes. These findings need to be viewed in the context of a high possibility of performance bias, because the nature of the interventions made it impossible to blind the participants. In addition, there is a possibility of reporting bias and of a higher proportion of interventions showing an effect being published than those not doing so.

The percentage of adherence interventions showing improvements in medication adherence and clinical outcomes in this review was higher than in a Cochrane review of medication adherence interventions\textsuperscript{13} in which 44\% were found to lead to improvements in medication adherence and 31\% to improvements in clinical outcomes. In addition, improvement in adherence was higher than that found in another large-scale review.\textsuperscript{14} Our review focused on interventions involving doctors and pharmacists, while the other two reviews\textsuperscript{13,14} included all adherence interventions. This indicates that involving pharmacists and doctors in adherence interventions may make them more effective. However, we are cautious with the interpretation of this finding, because the inclusion and exclusion criteria were not identical in the reviews. In addition, the studies covered by the reviews used many different types of interventions and different methods of measuring adherence and clinical outcomes, making direct comparisons impossible.
It is of interest that in both our review and that of Morgado et al.,\textsuperscript{18} who reviewed pharmacy interventions to enhance blood-pressure control and adherence to antihypertensive therapy, more studies showed improvement in clinical outcomes than showed improvement in adherence. This contrasts with reviews of general adherence support programs\textsuperscript{13,14} (not necessarily involving a pharmacist), in which adherence was found to improve more frequently than clinical outcomes. The difference may be due to the multifaceted nature of interventions and the fact that pharmacists may make clinical recommendations as well as provide adherence support. The difference may also be due to the nonlinear relationship between adherence and clinical outcome in some conditions, and because some clinical outcomes, such as high blood pressure, are affected by factors that may not have been part of the intervention (such as exercise or diet).

**Recommendations: ways to improve the delivery of patient adherence support**

**Recommendations for practice**

Our review demonstrates that while the majority of adherence support provided by pharmacists and doctors leads to some improvements in outcomes, there is room to improve the provided support further. The successful features of adherence interventions that have been identified by previous research were not reported as being present in the majority of adherence interventions provided by pharmacists and doctors.

Previous research has suggested that adherence support involving pharmacists that is tailored to individual needs, rather than being fixed, is more likely to be effective.\textsuperscript{15} In our review, while there was a trend toward more tailored approaches being used in the last 15 years, less than 50% of adherence support programs in the last 15 years were tailored to individual needs, with the remainder either being fixed or not reported in sufficient detail to determine if they had been tailored. We would therefore recommend increasing the level of tailoring of interventions to improve the delivery of patient adherence support.

In addition, taking a more concordant approach, where the aim is to formulate an agreed treatment plan between the HCP and patient rather than expecting the patient to follow the HCP’s instructions, has been shown to improve treatment outcomes.\textsuperscript{1} In our review, out of the 41 studies in which tailoring to patient need was reported, only 14 described a concordant approach whereby patients’ individual goal values and identification of adherence barriers were evaluated and agreement reached. In the rest of the studies, the tailoring appeared to be a result of the HCP’s own view of the patient’s needs. We therefore recommend using a patient-centered approach and involving patients in decisions when tailoring adherence support.

Finally, the majority of studies did not describe any training for the HCPs involved. When training was reported, it was sometimes more focused on the specific clinical diseases involved rather than on how to give adherence support. We would recommend that HCPs receive specific training on how to develop a concordant approach with patients and formulate a joint treatment plan that meets patients’ own identified needs and that therefore may be more likely for patients to adhere to.

**Recommendations for research**

The majority of interventions were delivered by pharmacists rather than doctors. Evaluation of more adherence support programs involving doctors would inform this area of research, particularly as a previous review of adherence support in patients with cardiovascular problems and diabetes suggested doctors were less effective than other HCPs in providing effective adherence support in these disease groups.\textsuperscript{20}

A cost analysis economic evaluation was provided by very few of the studies. Cost analyses are important to evaluate the cost-effectiveness of interventions and would be helpful to incorporate into evaluations. The outcomes used in the studies included adherence to medication and adherence to attending screening appointments. However, use of these outcome measures makes the assumption that increased adherence is always a positive outcome. Nonadherence may be an appropriate decision by the patient. None of the studies included a measure of concordance or patient involvement in decision making. It would be appropriate to include such outcome measures – for example, the Leeds Attitude to Concordance scale\textsuperscript{125} – in future studies.

**Limitations of the review**

Due to the large volume of literature on adherence support strategies, our review only focused on the role of pharmacists and doctors in providing adherence support. Further reviews are needed on the role of other HCPs, especially nurses, in providing adherence support.

The search strategy for this review was broad and included adherence support for all treatments and preventive...
health care. This meant that we did not include search terms for specific treatments and preventive strategies such as “cancer screening” and may have missed some relevant studies in these areas. In addition, we excluded studies not written in English and did not use alternative names for “pharmacist”, such as “chemist”.

As with all reviews of adherence interventions, the definition of what is acceptable adherence is problematic. It is defined differently by different researchers and will vary from condition to condition. Relatedly, the differentiation between people who miss occasional doses and those who take drug “holidays” is often not made. Again, as for all such reviews, the measurement of adherence in studies presents difficulties, as there is no gold standard for such measurement.

Conclusion
Adherence support programs involving pharmacists and doctors appear to improve outcomes to some extent, although the size of effect is not always large. Their effectiveness may be increased by working with patients to tailor adherence strategies according to individual patient needs. A much greater volume of studies have evaluated the role of pharmacists than the role of doctors; therefore, more research is needed to evaluate the role of doctors in providing adherence support.

Acknowledgments
This study had no specific source of funding. However, the Centre for Medication Safety and Service Quality is affiliated with the Centre for Patient Safety and Service Quality at Imperial College Healthcare NHS Trust, which is funded by the National Institute of Health Research (NIHR). This paper represents independent research supported by the Imperial NIHR Patient Safety Translational Research Centre. The views expressed are those of the author(s) and not necessarily those of the NHS, the NIHR, or the Department of Health. The work contributed by LE and CJ was funded by Atlantis Healthcare.

Disclosure
LE and CJ work for Atlantis Healthcare, which provides consulting services to pharmaceutical companies. This research was not funded by a pharmaceutical company. DKR is co-founder and academic advisor to Luto Research (http://www.luto.co.uk), which develops, refines, and tests consumer health information. The authors declare no other conflicts of interest in this work.

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