Task-sharing interventions for improving control of diabetes (1977) (1974) in low-income and middle-income countries: a systematic review and meta-analysis





Joseph Linju Maria*, T N Anand*, Boban Dona, Jose Prinu, Dorairaj Prabhakaran, Panniyammakal Jeemon

Summary

Background Task-sharing interventions using non-physician health-care workers might be a potential diabetes management strategy in health systems that are constrained by physician shortages, such as those in low-income and middle-income countries (LMICs).

Methods We did a systematic review and meta-analysis of task-sharing intervention strategies for managing type 2 diabetes in LMICs. We searched PubMed, Embase, and CINAHL from database inception to Sept 25, 2019, for studies that were randomised control trials or cluster randomised trials with task-shifted or task-shared interventions delivered to adults (≥18 years) by non-physician health workers versus usual care, done in LMICs with glycated haemoglobin (HbA₁) or fasting blood sugar (FBS) as outcome measures. The methodological quality of included studies was assessed using the Cochrane risk of bias tool. Random-effects model meta-analysis was used to estimate the population average pooled mean difference for HbA₁, and FBS with 95% CIs. Our study protocol was registered in the PROSPERO database (CRD42018081015).

Findings We found 4213 studies from the literature search, of which 46 (1.1%) were eligible for the narrative synthesis, including a total of 16973 participants. 16 of these studies were excluded from the meta-analysis due to high risk of bias. 24 studies with a total of 5345 participants were included in the meta-analysis of HbA₁, and 18 studies with a total of 3287 participants for FBS. Interventions led to an average reduction in HbA₁, when tasks were delivered by nurses (averaged pooled mean difference -0.54% [95% CI -0.89 to -0.18]; P=80%) and pharmacists (-0.91% [-1.15 to -0.68]; P=58%), but not when they were delivered by dietitians (-0.50% [-1.10 to 0.09]; P=54%) or community health workers (0.05% [0.03 to 0.07]; P=0%). A reduction in average FBS was also observed when interventions were delivered by pharmacists (average pooled mean difference -36 · 26 mg/dL [-52 · 60 to -19 · 92]; P=78%) but not nurses (-7.46 mg/dL [-18.44 to 3.52]; P=79%) or community health workers (-5.41 [-12.74 to 1.92]; P=71%). Only one study reported on FBS when tasks were delivered by dietitians, with a mean difference of -35.00 mg/dL (-65.96 to -4.04).

Interpretation Task sharing interventions with non-physician healthcare workers show moderate effectiveness in diabetes management in LMIC settings. Although relatively high heterogeneity limits the interpretation of the overall findings, interventions led by pharmacists and nurses in LMICs with relatively high physician density are effective strategies in the management of diabetes.

Funding Wellcome Trust-Department of Biotechnology India Alliance.

Copyright © 2020 The Author(s). Published by Elsevier Ltd. This is an Open Access article under the CC BY 4.0 license.

Introduction

Non-communicable diseases (NCDs) continue to be a dominant cause of global deaths. For example, the Global Burden of Disease study¹ estimates that NCDs contributed up to three-quarters of total annual deaths in 2019. Furthermore, low-income and middle-income countries (LMICs) bear a disproportionately higher burden of NCDs than high-income countries. Diabetes is a major contributor to NCDs, affecting about 463 million people globally. Consequent to poor detection rates and awareness of diabetes, four of five adults with undiagnosed diabetes live in LMICs. Additionally, the treatment and control rates of diabetes are abysmally poor in LMICs.2 It has been estimated that the total unmet need for diabetes care in terms of poor detection, treatment, and control is about 77% in LMIC settings.^{2,3}

The management of diabetes is complex, necessitating continuous effort for achieving better control and evidence-based targets.4 The American Medical Association recommends team-based, patient-centred care, integrated long-term treatment approaches to diabetes and comorbidities, and ongoing collaborative communication among providers for improving diabetes care and population health as per the standards of medical care in diabetes. 5.6 Further, the management of diabetes becomes more multifaceted with other concurrent chronic conditions. Data suggest that as many as 85% of people with diabetes have at least one other chronic condition.⁷

Lancet Glob Health 2020

Published Online November 27, 2020 https://doi.org/10.1016/ S2214-109X(20)30449-6

*Joint first authors contributed

Centre for Chronic Disease Control, New Delhi, India (LL Maria MSc. T N Anand MPH B Dona MPH. Prof D Prabhakaran DM); Public Health Foundation of India, New Delhi, India (J Prinu MPH, Prof D Prabhakaran): and Achutha Menon Centre for Health Science Studies, Sree Chitra Tirunal Institute for Medical Sciences and Technology, Trivandrum, India (P Jeemon PhD) Correspondence to:

Dr Panniyammakal Jeemon, Achutha Menon Centre for Health Science Studies, Sree Chitra Tirunal Institute for Medical Sciences and Technology, Trivandrum 695011, pjeemon@gmail.com

For statistics in the Diabetes Atlas of the International Diabetes Federation see https://www.diabetesatlas.org/

Research in context

Evidence before this study

Task-sharing has been proposed as a strategy to deal with the shortage of physicians in low-income and middle-income countries (LMICs). We searched PubMed, Embase, and CINAHL for task-sharing interventions in managing people with diabetes. Two previous reviews show that task sharing for managing non-communicable diseases is a potentially viable strategy, and another review reported reductions in population average blood pressure with task-sharing strategies. However, no systematic reviews or meta-analyses of task-sharing interventions for managing diabetes have been published.

Added value of this study

In this systematic review and meta-analysis, we determined the effectiveness of task-sharing interventions for managing diabetes in LMICs. We showed that task-sharing interventions led by nurses and pharmacists are effective in achieving meaningful

population average reductions in glycated haemoglobin and fasting blood sugar. However, task-sharing interventions with community health workers did not show meaningful reductions. Additionally, the task-sharing strategy was more effective in LMICs with higher doctor–population ratios.

Implications of all the available evidence

Our data provide evidence to support task-sharing interventions that involved higher cadres of health-care providers, such as nurses and pharmacists, in managing diabetes in LMIC settings. The magnitude of the population-average reduction in glycated haemoglobin and fasting blood sugar are similar to the reductions associated with oral hypoglycaemic drugs in clinical trials. The supervision and levels of training needed for the health-care workforce should be determined and tailored according to the capacity of the health system.

Complications of poor glycaemic control, such as macrovascular diseases, blindness, and kidney failure, are often devastating for the individual and family, and incur catastrophic health spending in LMIC settings.^{8,9} There is a clear shortage of well trained health-care providers in LMICs, despite the increased demand for integrated care and the emphasis on the availability of trained personnel at decentralised levels of health care to provide integrated care for diabetes.^{10,11} The average physician consultation time is often low in LMICs, which leads to ineffective communication, clinical handover, and lifestyle counselling for patients with diabetes.¹²

Reorganising health service delivery using non-physician health-care workers has been successful in improving outcomes for maternal and child health care, and a similar reorganising approach might improve outcomes of diabetes care. Task shifting, 13 an approach for redistributing human resources rationally, could help in improving health-care delivery.14 In general, task-shifting describes a situation in which a task that is normally performed by a physician is transferred to a health professional with a different or lower level of education and training. This approach can also include lower cadres of health-care professionals being trained in a particular task. 15,16 The term task sharing has been used recently to describe this concept, as it better describes the process of care in teambased provision of integrated care for patients with chronic conditions.17 Previous systematic reviews highlight task sharing as a potential strategy for managing NCDs. 13,18 We examined the nature and effectiveness of task sharing interventions for managing diabetes in LMIC settings.

Methods

Search strategy and selection criteria

For this systematic review and meta-analysis, we did a systematic search to summarise task-sharing interventions for managing diabetes. We developed a search strategy in PubMed using previous reviews, ¹⁸ which we modified for use in other databases (Embase and CINAHL) for locating articles from inception to Dec 28, 2018. We then did an updated search for articles published before Sept 25, 2019. No language restrictions were used. A range of search terms was used, relating to diabetes, task sharing, task shifting, and a list of LMICs based on the 2018 World Bank database¹⁹ (appendix pp 41–45). Hand searching was done using citations and reference lists of the studies included.

We included randomised control trials or cluster randomised trials with task-shifted or task-shared interventions versus usual care that were done in LMICs. To be included, trials had to be of patients aged 18 years or older with type 2 diabetes and had to involve measurement and reporting of glycaemic outcomes as the change in glycated haemoglobin or fasting blood glucose. Task shared interventions had to be designed to improve glycaemic control and to be delivered by community health workers, nurses, pharmacists, or allied health professionals (eg, dietitians). The tasks shared included non-pharmacological measures (eg, patient education for lifestyle modification) and pharmacological measures (eg, initiation or refill of prescription medications and titrating the dose of medications). We excluded studies of children, mothers with gestational diabetes, or patients with type 1 diabetes, and studies that did not involve tasksharing interventions for diabetes or cardiovascular risk management. Interventions led by peer educators or home carers were also excluded, because they are not health-care professionals, and studies of task-sharing activities that are exclusive to traditional healers or alternative therapies (eg, acupuncture and homoeopathic medicine), or that promoted only self-care or informal caregiver health education. Additionally, we excluded

See Online for appendix

reviews, pre-post studies, cross-sectional studies, casecontrol studies, case series studies, and drug efficacy studies, as well as studies that had a duration of less than 3 months and studies that measured knowledge and attitude or practice outcomes without reported glycaemic measures.

Two authors (JLM and TNA) did the literature search independently, using the search strategy developed in consultation with the other authors. Two authors (JLM and BD) independently removed the duplicates manually. Further, they reviewed all the titles independently, and any conflicts in article selection were resolved after mutual discussion (between JLM and BD). The remaining abstracts were assessed for potential eligibility by the same reviewers (JLM and BD). Finally, three authors (JLM, BD, and JP) independently reviewed the full texts of the included articles, and any disagreements were resolved through discussion with a fourth reviewer (PJ). Our study protocol was registered in the PROSPERO database (CRD42018081015).

Data analysis

Studies that met the inclusion criteria after the full-text review were assessed for quality by two authors (JLM and BD) using the Cochrane risk of bias tool.20 Double data extraction was done by two authors (BD and IP) from the eligible full articles. Any discrepancies were discussed and clarified with two other authors (JLM and TNA). The arbitrator (PJ) reviewed any apparent discrepancies and made the final recommendation. We extracted the details of patients, the delivered interventions, components of the intervention, and relevant results of the studies. We extracted the outcome measurements for glycated haemoglobin (HbA_{1c}) and fasting blood glucose (FBS) that were taken before and after the intervention, for both the intervention and control groups. Information regarding the country where the study was conducted and study population type and size were also noted.

We did a qualitative synthesis of types of intervention and sample characteristics of the included trials. For quantitative synthesis, we included eligible randomised controlled trials and clustered randomised trials with at least 30 participants in each treatment arm for our metanalysis. For cluster randomised trials, we estimated the effective study sample size using the reported design effect, or we calculated the design effect on the basis of the intra-cluster correlation coefficient and average cluster size. A change in glycaemic levels was estimated using the difference between the mean HbA_{lc} or FBS in the intervention arm (ie, task sharing) and the control group (ie, usual care).

We adopted a random effect model for meta-analysis because we assumed a greater study-level variability due to differences in task-sharing groups, types of interventions, and study populations. Appropriate weights were assigned for individual studies included in the meta-analysis on the basis of the inverse variance

method. The Der Simonian and Laird method was used for assessing between-study variance.²³ We estimated the pooled mean difference for HbA_{1c} and FBS along with their 95% CIs. We generated prediction intervals to assess the uncertainty of the summary estimate across different study settings.^{21,24} We conducted independent heterogeneity assessments for HbA_{1c} and FBS analyses. Heterogeneity was assessed using the *I*² statistic, and the statistical significance of heterogeneity was tested with Q statistics.²⁵ The heterogeneity contribution from each study was assessed by omitting each study and recording the change in overall heterogeneity.

An exploratory subgroup analysis was done with the covariates sample size, duration of intervention, country physician density, study population characteristics, and geographical regions as classified by WHO. We did univariate and multivariate meta-regression analyses to identify the effect of covariates on the effect size. Let univariate be publication bias was graphically assessed with funnel plots and contour-enhanced funnel plots. We also used Egger's regression test for the statistical significance of publication bias. We analysed data using the meta package of R version 3.5.1. The quality of evidence was evaluated using GRADE. Ethics approval was not required for this study.

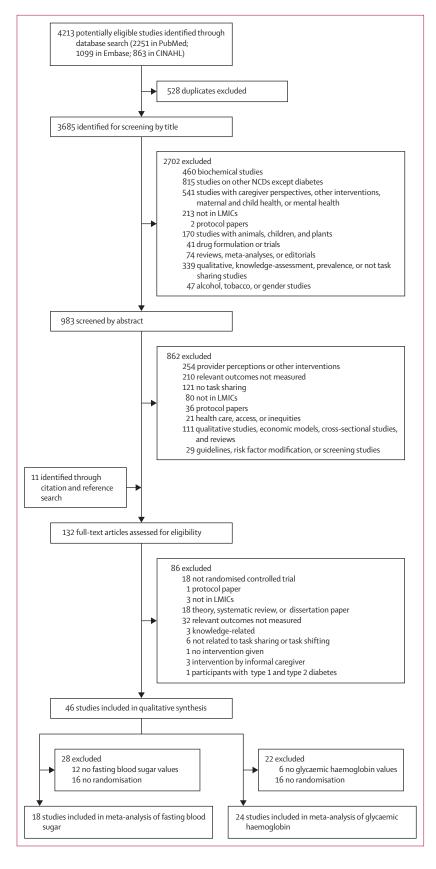
Role of the funding source

The funder had no role in the study design, data collection, data analysis, data interpretation, or the writing of the report. All authors had full access to all the data in the study and had final responsibility for the decision to submit for publication.

Results

We identified 4213 references from all the databases (figure 1), 528 (12 \cdot 5%) of which were duplicates. We screened 3685 (87 \cdot 5%) of the titles and 983 (23 \cdot 3%) abstracts for eligibility, and promoted 121 (2 \cdot 9%) articles for full-text review. Additionally, 11 studies were obtained after hand searching and were also included in the full-text review. Of the 132 articles selected for full-text review, 86 (65 \cdot 2%) were excluded, most commonly because the relevant outcomes were not measured, and 46 (34 \cdot 8%) were included in the narrative synthesis, including a total of 16 973 participants (figure 1).

The characteristics of the included studies are summarised in the appendix (pp 2–16). Among the 46 included studies, there were eight (17%) cluster randomised trials ^{31–38} and 38 (83%) randomised trials. ^{39–76} There were nine trials each from Brazil and China. The studies were done in primary health centres (n=10 [22%]), ^{31,33,35,50,63,65,66,70,72,74} diabetes clinics or community health centres (n=12 [26%]), ^{32,34,36,45,47–49,51,67,68,71,75} hospitals (n=21 [44%]), ^{39,40–44,46,52,53,55–61,62,64,69,73,76} and community settings (n=3 [9%]). ^{37,38,54} The sample size ranged from 53 participants⁶⁵ to 4393 participants. ³⁴ 34 (74%) of 46 studies were with patients with diabetes.



The other studies were with older patients (n=1 [2%]),⁵¹ patients with high risk of diabetes (n=2 [4%]),^{48,74} or patients with cardiovascular disease (n=1 [2%]),³⁷ hypertension (n=1 [2%]),³⁶ hypertension and diabetes (n=2 [4%]),^{38,66} metabolic syndrome (n=1 [2%]),⁴⁶ coronary artery disease (n=2 [4%]),^{46,49} myocardial infarction (n=1 [2%]),⁴³ or NCD (n=1 [2%]).³³ The participant follow-up ranged from 2 months^{42,43} to 36 months.^{66,71} 21 (46%) of 46 trials followed up participants for 12 months or more.

On the basis of the strategies used as the predominant component, interventions led by non-physician healthcare workers were broadly categorised as pharmacological or non-pharmacological. However, there were multiple elements in the interventions of the studies included. Pharmacological interventions involved the generation of a medication prescription following an algorithm or guidelines, dose adjustment or titration of medications, and drug modification. Non-pharmacological interventions included assessment, monitoring, lifestyle education, and counselling for the management of diabetes. The interventions were delivered by pharmacists (n=16 [35%]), 41,42,44,45,52-56,61,62,66,68-70,73 nurses [41%]), 31,33–36,39,40,43,46,49,57,59,64,65,67,71,72,75,77 dietitians or nutritionists (n=4 [9%]), 47,50,58,74 community health workers $(n=5 [11\%])^{32,37,38,48,63}$ or a combination of community health centre staff, nurses, or pharmacists (n=2 [4%]).51,60

Of the 46 trials, 37 (80%) delivered a non-pharmacological intervention, eight (17%) had both non-pharmacological and pharmacological components, 38,40,42,44,52,61,66,69 and one (2%)33 described a pharmacological intervention exclusively. Pharmacological components in the trials ranged from drug initiation (n=3 [7%])33,44,66 to dose adjustment (n=8 [17%])^{38,40,42,44,52,61,66,69} and the addition of new drugs (n=2 [4%]).33,69 The pharmacological interventions were task-shared with pharmacists (n=6 [13%]), 42,44,52,61,66,69 nurses (n=2 [4%]),33,40 and community health workers (n=1 [2%]).38 The non-pharmacological interventions described in the trials ranged from providing education regarding lifestyle modification (n=45 [98%]) to telephone follow-up (n=12 [26%]) $^{39-42,46,52,54,55,57,60,71,72}$ and home visits (n=5 [11%]).49,63,64,67,75 The contents of lifestyle education were focused on dietary changes (n=39 [85%]), physical activity (n=38 [83%]), and medication adherence (n=37 [80%]).

19 (41%) of the interventions were organised as individual sessions, 14 (30%) had group sessions, and seven (15%) had a mixture of both group and individual sessions. The format of intervention sessions was not described in six (13%) studies. The frequency of sessions with the participants in the intervention group

Figure 1: Study selection

 $\label{lower} NCDs = non-communicable\ diseases.\ LMICs = low-income\ and\ middle-income\ countries.$

varied. Some studies had weekly sessions and monthly follow-up classes, whereas others had one class per month for up to 6 months.

The detailed results of the quality assessment based on Cochrane risk of bias tool are presented in the appendix (pp 19–20). Of the 46 trials, 16 (35%) studies^{31,37,41,49,53,55,58,60-62,64,67-69,73,74} did not sufficiently describe randomisation, or the method of randomisation was unclear. Consequently, those trials were considered to have a high risk of bias and were not included in the meta-analysis. Allocation concealment was reported only in 14 (30%) studies,^{31,32,36,38,39,43,46,47,51,54,63,65,71,72} whereas 19 (41%) studies^{32,34,36-41,43-47,57,63,68-70,72} reported masking of outcome assessors.

In terms of the availability of outcome variables, of the 30 trials considered for meta-analyses, six did not report before and after values for HbA_{lc} . Therefore, 24 studies were included in the meta-analysis of the HbA_{lc} outcome. Similarly, 12 studies did not have before and after values for FBS, and therefore 18 studies were included in the meta-analysis of the FBS outcome.

Regarding the effect of interventions on HbA_{tc} , we included 24 studies (19 randomised controlled trials and five cluster randomised trials) in the analysis after assessing risk of bias. Overall, the population average pooled mean difference in HbA_{tc} was -0.58% (95% CI -0.86 to -0.30; $I^2=95\%$) with a prediction interval ranging from -1.87 to 0.71 (figure 2). Simple funnel plots for publication bias showed asymmetry (appendix p 27), and the Eggers regression test also reported significant bias (t=-3.41; df=22; p=0.0024). To assess the reason for the asymmetric funnel plot, we examined contour-enhanced funnel plots. Most of the missing studies were in the areas of high statistical significance and unlikely to be due to publication bias (appendix p 27).

In the pooled effects on HbA_{1c}, studies involving task sharing with community health workers showed an averaged pooled mean difference of 0.05% (95% CI 0.03 to 0.07), whereas for dietitians it was -0.50% (-1.10 to 0.09) and for nurses it was -0.54% (-0.89 to -0.18; figure 2). In the subgroup analysis, for nurses implementing non-pharmacological interventions the averaged pooled mean difference was -0.54% (-0.91 to -0.18), compared with -0.59% (-2.21 to 1.02) for them implementing pharmacological interventions (appendix p 28). Task sharing with pharmacists showed an average pooled mean difference of -0.91% (-1.15 to -0.68), and pharmacologists implementing both pharmacological and non-pharmacological interventions showed similar effects on HbA_{1c} (figure 2; appendix p 28).

In subgroup analysis of the effects on HbA_{1c} , task-sharing interventions in the diabetes population showed an average pooled mean difference of -0.68% (95% CI -0.92 to -0.44; appendix p 29), and restriction to sample sizes of more than 200 showed an average pooled mean difference of -0.17% (-0.34 to 0.00; appendix p 30). Subgroup analysis with different intervention time

periods showed an average pooled mean difference in HbA_{tc} of -0.45% (-0.78 to -0.12) for studies of 6-15 months and -0.57% (-0.86 to -0.28) for studies of more than 15 months (appendix p 31). The average pooled mean difference in study settings with up to ten doctors per 10000 population was -0.55 (-0.89 to 0.21), and in settings with more than 15 doctors per 10 000 population it was -0.71% (-1.01 to -0.41; appendix p 32). When studies were grouped by WHO geographical regions, the average pooled mean difference in HbA₁₆ were -0.24% (-0.57 to 0.09) for the western Pacific region, -0.63% (-0.86 to -0.41) for the Americas, 0.08%(-0.23 to 0.39) in the African region, -0.64%(-1.16 to -0.13) for the southeast Asia region, and -1.23% (-1.56 to -0.90) for the eastern Mediterranean region (appendix p 33).

We assessed the contribution to heterogeneity by doing sensitivity analysis by excluding each study (appendix p 21). The exclusion of two studies 48.56 with a high heterogeneity contribution from the meta-analysis reduced the overall heterogeneity moderately, whereas the pooled estimate remained similar (-0.57 [95% CI -0.80 to -0.35]; data not shown). We identified WHO regions, study population, duration of the task-sharing intervention, and trial sample size as major predictors of effect size based on multivariate meta-regression analysis, and these covariates together accounted for 93.4% of the heterogeneity (appendix p 22).

Concerning the effect of interventions on FBS, we included 18 individual and cluster randomised controlled trials in the analysis with 3635 participants. The overall population average pooled mean difference of FBS was -16.74 mg/dL (95% CI -24.20 to -9.29) with a prediction interval ranging from -46.18 mg/dL to 12.70 mg/d (figure 3). Funnel plots for publication bias showed asymmetry (appendix p 34) and the Eggers regression test reported significant bias (t=-3.84, df=16, p=0.0014). Visual examination of the contour-enhanced funnel plot also suggested bias, with few studies in the high significance area and missing studies in the non-significance area (appendix p 34).

We assessed mean difference in FBS when different groups implement task sharing, and it was statistically different across the groups (figure 3). Studies with community health workers showed an average pooled mean difference of –5·41 mg/dL (95% CI –12·74 to 1·92; figure 3). Task sharing with dietitians was reported in only one study⁵⁰ with a mean difference of –35·00 mg/dL (65·96 to –4·04; figure 3). Task sharing with nurses gave an average pooled mean difference of –7·46 mg/dL (–18·44 to 3·52), and for pharmacists it was –36·26 mg/dL (–52·60 to –19·92; figure 3).

For the subgroup analysis of FBS, the effects of task sharing with pharmacological and non-pharmacological interventions are in the appendix (p 35). Task-sharing interventions in the diabetes population showed an average pooled mean difference of -23·94 mg/dL

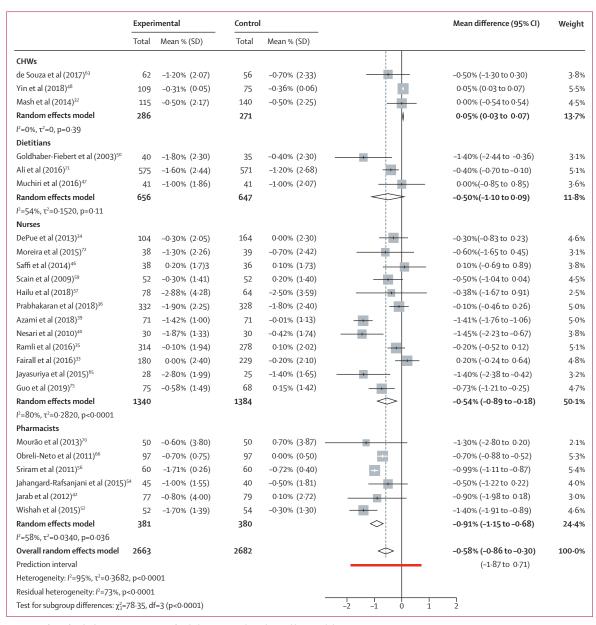


Figure 2: Effect of task-sharing interventions for diabetes control on glycated haemoglobin CHWs=community health workers.

(95% CI –38·51 to –9·37) for people with type 2 diabetes (appendix p 36), and for sample sizes with more than 200 participants the average pooled mean difference was –1·59 mg/dL (–17·40 to 14·21; appendix p 37). Subgroup analysis with different intervention time periods showed an average pooled mean difference in FBS of –35·00 mg/dL (–66·96 to –4·04) for up to 5 months, –14.86 mg/dL (–22·69 to –7·69) for 6–15 months, and –21·48 mg/dL (–35·27 to –7·69) for more than 15 months (appendix p 38). The average pooled mean difference in study settings with up to ten doctors per 10 000 population was –15·82 mg/dL

(–55·60 to 24·97), and for more than 15 doctors per 10 000 population it was $-25\cdot60$ mg/dL ($-39\cdot76$ to $-11\cdot43$; appendix p 39). The average pooled mean difference in FBS for the different WHO regions were $-9\cdot72$ mg/dL ($-17\cdot22$ to $-2\cdot21$) for the western Pacific region, $-24\cdot34$ mg/dL ($-37\cdot11$ to $-11\cdot56$) for the Americas, and $-32\cdot85$ mg/dL ($-74\cdot13$ to $8\cdot43$) for the eastern Mediterranean region (appendix p 40). We assessed the contribution to heterogeneity by doing sensitivity analysis by excluding each study (appendix p 23). In the multivariate meta-regression, the task-sharing group and sample size were significant

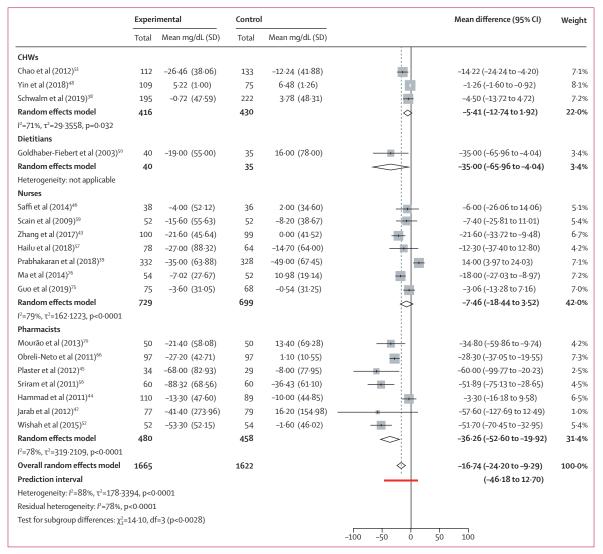


Figure 3: Effect of task-sharing interventions for diabetes control on fasting blood sugar CHWs=community health workers.

predictors of FBS change, and these covariates together accounted for $36\cdot8\%$ of heterogeneity (appendix pp 23–24).

Based on the GRADE criteria, the effect of nurse-led interventions on population average reduction in HbA_{1c} was graded as moderate (appendix p 25) and was downgraded for inconsistency. Due to indirectness and inconsistency in the relationship, the level of evidence from pharmacist-led interventions were also downgraded to moderate. The evidence for lowering HbA_{1c} and FBS was rated as low for interventions delivered by community health workers.

Discussion

We did a comprehensive review of available literature on task-sharing interventions and quantitatively synthesised the population-average pooled mean difference for HbA₁. and FBS levels. Our findings support the use of tasksharing interventions for the management of diabetes, with a moderate to good effect on the reduction in HbA₁ and FBS. However, the analyses with prediction intervals suggests that the effect of interventions on glycaemic outcomes might vary across study settings. In the subgroup analysis, the available evidence supports the engagement of higher cadres of non-physician health-care workers, such as nurses and pharmacists, for task-sharing activities related to management of diabetes. For example, non-pharmacological interventions by nurses and both non-pharmacological and pharmacological interventions by pharmacists resulted in clinically meaningful reductions in HbA1c. Additionally, nonpharmacological interventions delivered by pharmacists resulted in a large pooled average mean difference in FBS.

The moderate reduction in HbA $_{1c}$ of 0.58%, as observed in our overall pooled estimate for task-sharing interventions, might lead to significant public health effects at the population level in LMIC settings. For example, a 1% reduction in HbA $_{1c}$ was associated with a 37% reduction in microvascular complications in the UKPDS study. Similar glycaemic efficiency is reported when using dipeptidyl peptidase-4 inhibitors or gliptins, with a reduction of 0.5-1% when used as monotherapy and 0.6-1.1% when used in combination with metformin, depending on the drug, dose of therapy, and starting HbA $_{1c}$. A systematic review of general diabetes disease management programmes also show HbA $_{1c}$ reduction at a similar magnitude to that of our pooled analysis.

Task-shared interventions for diabetes management mainly comprise non-pharmacological lifestyle interventions and pharmacological components. When delivered by higher cadres of health workers, both the non-pharmacological and pharmacological interventions resulted in meaningful reductions in HbA_{1c}. In general, diabetes self-management education and support have been proven to be effective in glycaemic control.80,81 Additionally, diabetes self-management education programmes have been organised in high-income settings using health-care workers such as nurses, pharmacists, and certified diabetes educators.6 Health systems in LMICs have traditionally been organised around vertical disease management programmes and hence the care processes have mostly centred on physicians. This places a high work burden on physicians and poorly defines the roles of other health-care workers in LMICs.82 Therefore, it is necessary to restructure and organise roles for nonphysician health-care workers and to provide adequate training and supervision for effectively managing diabetes and other chronic non-communicable disease conditions in an integrated team-based care model.

The non-pharmacological and pharmacological interventions delivered by pharmacists in the studies included emphasised medication adherence as a key strategy that resulted in better glycaemic outcomes. Similar effects of glycaemic control were shown in studies that emphasised improving adherence to glycaemic medications.^{83,84} Hence, adherence improvement should be considered as a major component of task sharing intervention studies in management of diabetes outcomes in LMICs.

The wide prediction intervals of the effect estimates observed in our study could be attributable to the variation in glycaemic outcomes of task-sharing strategies across different WHO regions. The WHO regions differ in terms of health-care system characteristics and doctor-population ratios. Further, they employ different cadres of health workers for task-sharing interventions. Therefore, careful selection of the cadre of health workers and the capacity for supervision by physicians are important parameters in the scale-up of task-sharing interventions to improve glycaemic outcomes in LMICs.

The implications of our study for future research and practice are that public health interventions in diabetes management that are effective and useful in selected study settings need to be scaled up and implemented more widely to create wider health impacts. However, for effective task-sharing implementation, it is necessary to strengthen health systems and have health-care regulations in terms of policies supporting non-physician health-care workers. Task-sharing of interventions in diabetes management is an ideal candidate for scaling up as a general strategy for prevention and control of NCDs in LMICs. However, as recommended in the 2019 standards of medical care in diabetes,5 it is important to evaluate quality improvement strategies by incorporating reliable data metrics, ongoing data collection, and evaluation with the larger aim of improving processes of care and outcomes within the available resources. A recent systematic review⁸⁵ identifies several health system factors that support and impair the ability of non-physician health-care workers to manage NCDs. Adequate and standardised training sessions, proper guidance, reliable systems to track patient data, reasonable compensation or performance incentives for the work done, and logistical support are recognised as key facilitators to implement task-sharing interventions.85 Given the relative advantage of technology-assisted diabetes prevention and management interventions, more such options should be explored to expand the scope and coverage of task-sharing interventions.

The strengths of this review include a registered protocol and a comprehensive search strategy in multiple databases. This study also has limitations. First, the weak description of the intervention strategy in most studies included did not allow analysis of the effect of different types of interventions on glycaemic control. Second, significant heterogeneity was observed across the studies. However, to some extent, heterogeneity is inevitable in reviews of findings from many countries and public health interventions. This does not negate the applicability of task-shared interventions because the original populations, settings, and interventions could be quite diverse, increasing the likelihood that the evidence can be applied broadly.

In conclusion, clinicians and policy makers, when considering the organisation of care for patients with diabetes in LMICs, should consider task-sharing interventions with non-physician health-care workers. To achieve the optimal control of diabetes, non-pharmacological interventions for medication and lifestyle adherence are also needed. Prioritising potential interventions on the basis of health-care workers' availability and skills could improve glycaemic control in individuals with diabetes.

Contributor

DP and PJ conceived of the study. JLM, TNA, PJ, and DP developed the protocol. JLM and TNA did the literature search. JLM, BD, and JP appraised study quality, and extracted and analysed the data. JLM, TNA, DP, and PJ interpreted the data. JLM and TNA wrote the first draft of the article. JLM, TNA, BD, JP, DP, and PJ reviewed and critically evaluated the draft paper.

Declaration of interests

We declare no competing interests.

Data sharing

All datasets generated and analysed are available in the article and appendix.

Acknowledgments

This work was supported by a Wellcome Trust–Department of Biotechnology India Alliance Clinical and Public Health Intermediate Fellowship (grant IA/CPHI/14/1/501497) awarded to PJ. PJ also received research grants from the US National Heart Lung and Blood Institute (5R01HL125442-05), the Australian National Health and Medical Research Council (1160283 and 1169766), the World Diabetes Federation (WDF-15-959), and the Indian Council of Medical Research.

References

- 1 GBD 2019 Diseases and Injuries Collaborators. Global burden of 369 diseases and injuries in 204 countries and territories, 1990–2019: a systematic analysis for the Global Burden of Disease Study 2019. Lancet 2020; 396: 1204–22.
- Manne-Goehler J, Geldsetzer P, Agoudavi K, et al. Health system performance for people with diabetes in 28 low- and middle-income countries: a cross-sectional study of nationally representative surveys. PLoS Med 2019; 16: e1002751.
- 3 Abegunde DO, Shengelia B, Luyten A, et al. Can non-physician health-care workers assess and manage cardiovascular risk in primary care? Bull World Health Organ 2007; 85: 432–40.
- 4 Inzucchi SE, Bergenstal RM, Buse JB, et al. Management of hyperglycemia in type 2 diabetes: a patient-centered approach: position statement of the American Diabetes Association (ADA) and the European Association for the Study of Diabetes (EASD). Diabetes Care 2012; 35: 1364–79.
- 5 American Diabetes Association. Improving care and promoting health in populations: Standards of Medical Care in Diabetes—2020. *Diabetes Care* 2020; 43 (suppl 1): S7–13.
- 6 Powers MA, Bardsley J, Cypress M, et al. Diabetes self-management education and support in type 2 diabetes: a joint position statement of the American Diabetes Association, the American Association of Diabetes Educators, and the Academy of Nutrition and Dietetics. Clin Diabetes 2016; 34: 70–80.
- 7 Anjana RM, Pradeepa R, Deepa M, et al. Prevalence of diabetes and prediabetes (impaired fasting glucose and/or impaired glucose tolerance) in urban and rural India: phase I results of the Indian Council of Medical Research-INdia DIABetes (ICMR-INDIAB) study. Diabetologia 2011; 54: 3022–27.
- Forbes JM, Cooper ME. Mechanisms of diabetic complications. Physiol Rev 2013; 93: 137–88.
- Seuring T, Archangelidi O, Suhrcke M. The economic costs of type 2 diabetes: a global systematic review. *Pharmacoeconomics* 2015; 33: 811–31.
- 10 Kruk ME, Porignon D, Rockers PC, Van Lerberghe W. The contribution of primary care to health and health systems in low- and middle-income countries: a critical review of major primary care initiatives. Soc Sci Med 2010; 70: 904–11.
- 11 Lall D, Engel N, Devadasan N, Horstman K, Criel B. Models of care for chronic conditions in low/middle-income countries: a 'best fit' framework synthesis. BMJ Glob Health 2018; 3: e001077.
- 12 Irving G, Neves AL, Dambha-Miller H, et al. International variations in primary care physician consultation time: a systematic review of 67 countries. BMJ Open 2017; 7: e017902.
- 13 Joshi R, Alim M, Kengne AP, et al. Task shifting for non-communicable disease management in low and middle income countries—a systematic review. PLoS One 2014; 9: e103754.
- 14 WHO, President's Emergency Plan For AIDS Relief, UNAIDS. Task shifting: rational redistribution of tasks among health workforce teams: global recommendations and guidelines. Geneva: World Health Organization, 2007.
- 15 Joshi R, Thrift AG, Smith C, et al. Task-shifting for cardiovascular risk factor management: lessons from the Global Alliance for Chronic Diseases. BMJ Glob Health 2018; 3 (suppl 3): e001092.
- 16 Dawson AJ, Buchan J, Duffield C, Homer CSE, Wijewardena K. Task shifting and sharing in maternal and reproductive health in low-income countries: a narrative synthesis of current evidence. Health Policy Plan 2014; 29: 396–408.

- 17 Anand TN, Joseph LM, Geetha AV, Prabhakaran D, Jeemon P. Task sharing with non-physician health-care workers for management of blood pressure in low-income and middle-income countries: a systematic review and meta-analysis. *Lancet Glob Health* 2019; 7: e761–71.
- 18 Anand TN, Joseph LM, Geetha AV, Chowdhury J, Prabhakaran D, Jeemon P. Task-sharing interventions for cardiovascular risk reduction and lipid outcomes in low- and middle-income countries: a systematic review and meta-analysis. J Clin Lipidol 2018; 12: 626–42.
- 19 World Bank. World bank country and lending groups. 2018. https://datahelpdesk.worldbank.org/knowledgebase/articles/906519-world-bank-country-and-lending-groups (accessed Dec 10, 2018).
- 20 Higgins JPT, Altman DG, Gøtzsche PC, et al. The Cochrane Collaboration's tool for assessing risk of bias in randomised trials. BMJ (Clinical Research Ed) 2011; 343: d5928.
- 21 Riley RD, Higgins JPT, Deeks JJ. Interpretation of random effects meta-analyses. BMJ 2011; 342: d549.
- 22 Borenstein M, Hedges LV, Higgins JP, Rothstein HR. A basic introduction to fixed-effect and random-effects models for meta-analysis. Res Synth Methods 2010; 1: 97–111.
- 23 DerSimonian R, Laird N. Meta-analysis in clinical trials. Control Clin Trials 1986; 7: 177–88.
- 24 Borenstein MHL, Higgins JPT, Rothstein HR. Introduction to meta-analysis. West Sussex, UK: John Wiley & Sons, 2009.
- 25 Higgins JP, Thompson SG. Quantifying heterogeneity in a meta-analysis. Stat Med 2002; 21: 1539–58.
- 26 Thompson SG, Higgins JP. How should meta-regression analyses be undertaken and interpreted? Stat Med 2002; 21: 1559–73.
- 27 Peters JL, Sutton AJ, Jones DR, Abrams KR, Rushton L. Contour-enhanced meta-analysis funnel plots help distinguish publication bias from other causes of asymmetry. J Clin Epidemiol 2008; 61: 991–96.
- 28 Egger M, Davey Smith G, Schneider M, Minder C. Bias in meta-analysis detected by a simple, graphical test. BMJ 1997; 315: 629–34.
- 29 Lall D, Engel N, Devadasan N, Horstman K, Criel B. Models of care for chronic conditions in low/middle-income countries: a 'best fit' framework synthesis. BMJ Global Health 2018; 3: e001077.
- 30 Guyatt G, Oxman AD, Akl EA, et al. GRADE guidelines: 1. Introduction-GRADE evidence profiles and summary of findings tables. J Clin Epidemiol 2011; 64: 383–94.
- 31 Labhardt ND, Balo J-R, Ndam M, Manga E, Stoll B. Improved retention rates with low-cost interventions in hypertension and diabetes management in a rural African environment of nurse-led care: a cluster-randomised trial. Trop Med Int Health 2011; 16: 1276–84.
- 32 Mash RJ, Rhode H, Zwarenstein M, et al. Effectiveness of a group diabetes education programme in under-served communities in South Africa: a pragmatic cluster randomized controlled trial. Diabet Med 2014; 31: 987–93.
- 33 Fairall LR, Folb N, Timmerman V, et al. Educational outreach with an integrated clinical tool for nurse-led non-communicable chronic disease management in primary care in South Africa: a pragmatic cluster randomised controlled trial. PLoS Med 2016; 13: e1002178.
- 34 DePue JD, Rosen RK, Seiden A, et al. Implementation of a culturally tailored diabetes intervention with community health workers in American Samoa. *Diabetes Educ* 2013; 39: 761–71.
- 35 Ramli AS, Selvarajah S, Daud MH, et al. Effectiveness of the EMPOWER-PAR intervention in improving clinical outcomes of type 2 diabetes mellitus in primary care: a pragmatic cluster randomised controlled trial. BMC Fam Pract 2016; 17: 157.
- 36 Prabhakaran D, Jha D, Prieto-Merino D, et al. Effectiveness of an mHealth-based electronic decision support system for integrated management of chronic conditions in primary care: the mWellcare cluster-randomized controlled trial. Circulation 2018; published online Nov 10. https://doi.org/10.1161/ CIRCULATIONAHA.118.038192.
- 37 Khetan A, Zullo M, Rani A, et al. Effect of a community health worker-based approach to integrated cardiovascular risk factor control in India: a cluster randomized controlled trial. Glob Heart 2019; 14: 355–65.
- 38 Schwalm JD, McCready T, Lopez-Jaramillo P, et al. A community-based comprehensive intervention to reduce cardiovascular risk in hypertension (HOPE 4): a cluster-randomised controlled trial. *Lancet* 2019; 394: 1231–42.

- 39 Azami G, Soh KL, Sazlina SG, et al. Effect of a nurse-led diabetes self-management education program on glycosylated hemoglobin among adults with type 2 diabetes. J Diabetes Res 2018; 2018: 4930157.
- 40 Nesari M, Zakerimoghadam M, Rajab A, Bassampour S, Faghihzadeh S. Effect of telephone follow-up on adherence to a diabetes therapeutic regimen. Jpn J Nurs Sci 2010; 7: 121–28.
- 41 Farsaei S, Sabzghabaee AM, Zargarzadeh AH, Amini M. Effect of pharmacist-led patient education on glycemic control of type 2 diabetics: a randomized controlled trial. J Res Med Sci 2011; 16: 43–49
- 42 Jarab AS, Alqudah SG, Mukattash TL, Shattat G, Al-Qirim T. Randomized controlled trial of clinical pharmacy management of patients with type 2 diabetes in an outpatient diabetes clinic in Jordan. J Manag Care Pharm 2012; 18: 516–26.
- 43 Zhang P, Hu YD, Xing F-M, Li C-Z, Lan W-F, Zhang X-L. Effects of a nurse-led transitional care program on clinical outcomes, healthrelated knowledge, physical and mental health status among Chinese patients with coronary artery disease: a randomized controlled trial. *Int J Nurs Stud* 2017; 74: 34–43.
- 44 Hammad EA, Yasein N, Tahaineh L, Albsoul-Younes AM. A randomized controlled trial to assess pharmacist–physician collaborative practice in the management of metabolic syndrome in a university medical clinic in Jordan. J Manag Care Pharm 2011; 17: 295–303.
- 45 Plaster CP, Melo DT, Boldt V, et al. Reduction of cardiovascular risk in patients with metabolic syndrome in a community health center after a pharmaceutical care program of pharmacotherapy follow-up. Braz J Pharm Sci 2012; 48: 435–46.
- 46 Saffi MAL, Polanczyk CA, Rabelo-Silva ER. Lifestyle interventions reduce cardiovascular risk in patients with coronary artery disease: a randomized clinical trial. Eur J Cardiovasc Nurs 2014; 13: 436–43.
- 47 Muchiri JW, Gericke GJ, Rheeder P. Effect of a nutrition education programme on clinical status and dietary behaviours of adults with type 2 diabetes in a resource-limited setting in South Africa: a randomised controlled trial. *Public Health Nutr* 2016; 19: 142–55.
- 48 Yin Z, Perry J, Duan X, et al. Cultural adaptation of an evidence-based lifestyle intervention for diabetes prevention in Chinese women at risk for diabetes: results of a randomized trial. Int Health 2018; 10: 391–400.
- 49 Huang YJ, Parry M, Zeng Y, Luo Y, Yang J, He GP. Examination of a nurse-led community-based education and coaching intervention for coronary heart disease high-risk individuals in China. Asian Nurs Res (Korean Soc Nurs Sci) 2017; 11: 187–93.
- 50 Goldhaber-Fiebert JD, Goldhaber-Fiebert SN, Tristán ML, Nathan DM. Randomized controlled community-based nutrition and exercise intervention improves glycemia and cardiovascular risk factors in type 2 diabetic patients in rural Costa Rica. Diabetes Care 2003; 26: 24–29.
- 51 Chao J, Wang Y, Xu H, et al. The effect of community-based health management on the health of the elderly: a randomized controlled trial from China. BMC Health Serv Res 2012; 12: 449.
- 52 Wishah RA, Al-Khawaldeh OA, Albsoul AM. Impact of pharmaceutical care interventions on glycemic control and other health-related clinical outcomes in patients with type 2 diabetes: randomized controlled trial. *Diabetes Metab Syndr* 2015; 9: 271–76.
- 53 Samtia AMRM, Ranjha NM, Usman F, Javed I. A multifactorial intervention to enhance adherence to medications and disease-related knowledge in type 2 diabetic patients in southern Punjab, Pakistan. Trop J Pharm Res 2013; 12: 851–56.
- 54 Jahangard-Rafsanjani Z, Sarayani A, Nosrati M, et al. Effect of a community pharmacist-delivered diabetes support program for patients receiving specialty medical care: a randomized controlled trial. *Diabetes Educ* 2015; 41: 127–35.
- 55 Chung WW, Chua SS, Lai PSM, Chan SP. Effects of a pharmaceutical care model on medication adherence and glycemic control of people with type 2 diabetes. *Patient Prefer Adherence* 2014; 8: 1185–94.
- 56 Sriram S, Chack LE, Ramasamy R, Ghasemi A, Ravi TK, Sabzghabaee AM. Impact of pharmaceutical care on quality of life in patients with type 2 diabetes mellitus. *J Res Med Sci* 2011; 16 (suppl 1): S412–18.
- 57 Hailu FB, Hjortdahl P, Moen A. Nurse-led diabetes self-management education improves clinical parameters in Ethiopia. Front Public Health 2018; 6: 302.

- 58 Sun J, Wang Y, Chen X, et al. An integrated intervention program to control diabetes in overweight Chinese women and men with type 2 diabetes. Asia Pac J Clin Nutr 2008; 17: 514–24.
- 59 Scain SF, Friedman R, Gross JL. A structured educational program improves metabolic control in patients with type 2 diabetes: a randomized controlled trial. *Diabetes Educ* 2009; 35: 603–11.
- 60 Liang R, Dai X, Zuojie L, Zhou A, Meijuan C. Two-year foot care program for minority patients with type 2 diabetes mellitus of Zhuang Tribe in Guangxi, China. Can J Diabetes 2012; 36: 15–18.
- 61 Cani CG, Lopes LS, Queiroz M, Nery M. Improvement in medication adherence and self-management of diabetes with a clinical pharmacy program: a randomized controlled trial in patients with type 2 diabetes undergoing insulin therapy at a teaching hospital. Clinics (São Paulo) 2015; 70: 102–06.
- 62 Tawa O, Mahwi KAO. Role of the pharmaceutical care in the management of patients with type 2 diabetes mellitus. *IJPSR* 2013; 4: 1363–69.
- 63 de Souza CF, Dalzochio MB, Zucatti ATN, et al. Efficacy of an education course delivered to community health workers in diabetes control: a randomized clinical trial. *Endocrine* 2017; 57: 280-86
- 64 Mollaoğlu M, Beyazit E. Influence of diabetic education on patient metabolic control. Appl Nurs Res 2009; 22: 183–90.
- 65 Jayasuriya R, Pinidiyapathirage MJ, Jayawardena R, et al. Translational research for diabetes self-management in Sri Lanka: a randomized controlled trial. Prim Care Diabetes 2015; 9: 338-45.
- 66 Obreli-Neto PR, Guidoni CM, de Oliveira Baldoni A, et al. Effect of a 36-month pharmaceutical care program on pharmacotherapy adherence in elderly diabetic and hypertensive patients. Int J Clin Pharm 2011; 33: 642–49.
- 67 Wattana C, Srisuphan W, Pothiban L, Upchurch SL. Effects of a diabetes self-management program on glycemic control, coronary heart disease risk, and quality of life among Thai patients with type 2 diabetes. Nurs Health Sci 2007; 9: 135–41.
- 68 Phumipamorn S, Pongwecharak J, Soorapan S, Pattharachayakul S. Effects of the pharmacist's input on glycaemic control and cardiovascular risks in Muslim diabetes. *Prim Care Diabetes* 2008; 2: 31–37.
- 69 Lim PC, Lim K. Evaluation of a pharmacist-managed diabetes medication therapy adherence clinic. *Pharm Pract (Granada)* 2010; 8: 250–54.
- 70 Mourão AO, Ferreira WR, Martins MA, et al. Pharmaceutical care program for type 2 diabetes patients in Brazil: a randomised controlled trial. Int J Clin Pharm 2013; 35: 79–86.
- 71 Ali MK, Singh K, Kondal D, et al. Effectiveness of a multicomponent quality improvement strategy to improve achievement of diabetes care goals: a randomized, controlled trial. Ann Intern Med 2016; 165: 399–408.
- 72 Moreira RC, Mantovani MF, Soriano JV. Nursing case management and glycemic control among Brazilians with type 2 diabetes: pragmatic clinical trial. Nurs Res 2015; 64: 272–81.
- 73 Shao H, Chen G, Zhu C, et al. Effect of pharmaceutical care on clinical outcomes of outpatients with type 2 diabetes mellitus. Patient Prefer Adherence 2017; 11: 897–903.
- 74 Sartorelli DS, Sciarra EC, Franco LJ, Cardoso MA. Beneficial effects of short-term nutritional counselling at the primary health-care level among Brazilian adults. *Public Health Nutr* 2005; 8: 820–25.
- 75 Guo Z, Liu J, Zeng H, He G, Ren X, Guo J. Feasibility and efficacy of nurse-led team management intervention for improving the self-management of type 2 diabetes patients in a Chinese community: a randomized controlled trial. *Patient Prefer Adherence* 2019: 13: 1353–62.
- 76 Ma C, Zhou Y, Zhou W, Huang C. Evaluation of the effect of motivational interviewing counselling on hypertension care. *Patient Educ Couns* 2014; 95: 231–37.
- 77 Stratton IM, Adler AI, Neil HAW, et al. Association of glycaemia with macrovascular and microvascular complications of type 2 diabetes (UKPDS 35): prospective observational study. BMJ 2000; 321: 405–12.
- 78 Makrilakis K. The role of DPP-4 inhibitors in the treatment algorithm of type 2 diabetes mellitus: when to select, what to expect. Int J Environ Res Public Health 2019; 16: 2720.

10

- 79 Pimouguet C, Le Goff M, Thiébaut R, Dartigues JF, Helmer C. Effectiveness of disease-management programs for improving diabetes care: a meta-analysis. CMAJ 2011; 183: e115–27.
- 80 Steinsbekk A, Rygg LØ, Lisulo M, Rise MB, Fretheim A. Group based diabetes self-management education compared to routine treatment for people with type 2 diabetes mellitus. A systematic review with meta-analysis. BMC Health Serv Res 2012; 12: 213.
- 81 Ellis SE, Speroff T, Dittus RS, Brown A, Pichert JW, Elasy TA. Diabetes patient education: a meta-analysis and meta-regression. Patient Educ Couns 2004; 52: 97–105.
- 82 Lall D, Engel N, Devadasan N, Horstman K, Criel B. Challenges in primary care for diabetes and hypertension: an observational study of the Kolar district in rural India. BMC Health Serv Res 2019; 19: 44.
- 83 Cramer JA. A systematic review of adherence with medications for diabetes. *Diabetes Care* 2004; 27: 1218–24.
- 84 Krass I, Schieback P, Dhippayom T. Adherence to diabetes medication: a systematic review. *Diabet Med* 2015; 32: 725–37.
- 85 Heller DJ, Kumar A, Kishore SP, Horowitz CR, Joshi R, Vedanthan R. Assessment of barriers and facilitators to the delivery of care for noncommunicable diseases by nonphysician health workers in low- and middle-income countries: a systematic review and qualitative analysis. JAMA Network Open 2019; 2: e1916545