A Motivational Interviewing Intervention by Pharmacy Students to Improve Medication Adherence

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ABSTRACT

BACKGROUND: Diabetes mellitus (DM) patients with comorbid hypertension (HTN) are at a higher risk of developing microvascular and macrovascular DM complications. Through guideline-driven recommendations, angiotensin-converting enzyme inhibitors (ACEIs) and angiotensin II receptor blockers (ARBs) are highly recommended for these patients. Unfortunately, medication adherence to these products, though crucial to achieving therapeutic benefit, is frequently suboptimal. Motivational interviewing (MI) is a patient-centered collaborative communication style that is used to strengthen internal motivation for change that may prove effective in enhancing adherence.

OBJECTIVE: To examine the effect of an MI telephone intervention conducted by pharmacy students in improving adherence to ACEIs/ARBs among Medicare Advantage Plan (MAP) patients with both DM and HTN.

METHODS: A prospective study was conducted among patients enrolled in a Texas MAP. Medical claims data were used to identify patients with DM and HTN, and pharmacy claims were observed to recognize those who filled either an ACEI or an ARB during June 2014. Patients with a 6-month proportion of days covered (PDC) < 0.80 in the previous 6 months were determined nonadherent, and 75% of those were randomly selected to serve as potential subjects for the intervention, while 25% were randomly selected to serve as potential subjects for the control group. The intervention was a telephone call by a pharmacy student on rotation at the health plan, and 5 monthly follow-up calls. Before implementing calls, participating students attended a 3-day MI training course, where their proficiency for MI skills was evaluated. Refill data during the 6-month postintervention were evaluated to examine the intervention effect measured on 3 outcomes: PDC; PDC≥0.80 versus < 0.80; and discontinuation versus continuation. Multivariate linear and logistic regression models were constructed to adjust for any imbalances in baseline characteristics, including age, gender, number of other medications, regimen complexity, health low-income subsidy status, prescriber specialty, comorbidities, 6-month previous hospitalization, baseline 6-month PDC, and Centers for Medicare & Medicaid Services risk score.

RESULTS: A total of 11 students participated in the intervention implementation. Patients receiving calls were randomly selected from those potential subjects for the intervention arm until a target of 250 was reached; 500 controls were randomly selected from the potential subjects for the control arm. The final cohort included in multivariate models consisted of 743 patients. Patients completing the initial call and at least 2 follow-ups were less likely to discontinue (OR = 0.29; 95% Cl = 0.15-0.54; P < 0.001) and more likely to be adherent in the linear regression model ($\beta = 0.0604$, P < 0.001) and the logistic regression model (OR = 1.53; 95% Cl = 1.02-2.28; P = 0.009). Other factors significantly associated with better adherence included higher baseline PDC and number of medications. Depression status was significantly associated with lower adherence.

CONCLUSIONS: Patients receiving 2 or more calls had significantly better adherence and less discontinuation during the 6 months following initial calls compared with those who did not receive calls. This finding indicates that an MI-based telephone intervention by pharmacy students may be a promising intervention to improve adherence. Future research should examine the sustainability of the intervention effect for longer time periods and its influence on associated clinical outcomes.

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What is already known about this subject

- Motivational interviewing (MI) is a patient-centered communication style to strengthen motivation and commitment for change in a supportive, collaborative, empathetic manner and has been shown to promote behavioral change.
- Telephone interventions by different health care professionals have been reported to improve patient care among diabetes mellitus (DM) patients.
- Pharmacist interventions have been influential in promoting adherence among patients with DM and hypertension and among older adults, but they may have cost and time constraints.

What this study adds

- Patients who received the initial MI telephone intervention and completed 2 or more follow-up calls had significantly fewer discontinuation rates and better adherence during the 6-month postintervention period.
- This study demonstrated that an MI phone intervention had a promising effect on medication adherence.
- The feasibility of training pharmacy students in MI and implementing an intervention by students was demonstrated in this study.

A dherence to medication is a crucial step to achieving the full benefit of a prescribed treatment regimen and improving health outcomes. Approximately 20%-50% of patients with chronic illnesses are reported to be nonadherent to their medications, a situation that compromises treatment effectiveness.¹⁻³ Among elderly patients, even higher nonadherence rates, ranging from 40% to 86%, are reported,^{1.3,4} partially potentiated by an increased number of illnesses and a higher medication burden coupled with the cognitive decline of aging.^{2,5} The economic burden of nonadherence is considerable, costing \$100 billion annually in the United States. In addition, 33%-69% of medication-related hospital admissions result from poor adherence.^{3,6}

Diabetes mellitus (DM) and hypertension (HTN) both remain major public health issues as well as a growing economic burden, with notable increases in elderly and obese individuals.⁷ In 2010, approximately 6% of the U.S. population as a whole, and 27% of Americans aged 65 or older, had DM.^{8,9} It is projected that 50 million Americans will be living with DM and 100 million with HTN by 2050.¹⁰ DM and HTN independently increase a patient's risk for cardiovascular disease (CVD),^{7,11-14} a major global cause of morbidity and mortality.¹⁵ Approximately 70%-80% of patients with DM also have HTN,^{7,16} and studies suggest an additive adverse effect on CVD outcomes.⁹ There is also a significant increase in the risk of developing microvascular and macrovascular complications of DM among patients with comorbid HTN, commonly referred to as the deadly duet.¹⁴

Angiotensin-converting enzyme inhibitors (ACEIs) and angiotensin II receptor blockers (ARBs) are highly recommended for patients with both DM and HTN, according to American Diabetes Association guidelines,^{8,14,15} with a benefit beyond blood pressure (BP) control alone. Clinical trials have demonstrated that ACEIs/ARBs decrease the risk of developing microvascular and macrovascular adverse events in this patient population.8,17 ACEIs/ARBs have been shown to reduce allcause mortality,^{14,18-20} along with cardiac morbidity and mortality.14,18-21 Poor adherence to these medications continues to contribute to suboptimal BP control among DM patients,^{11,13,22-24} and fewer than 1 in 8 patients reach goal BP values as recommended by the guidelines.^{11,13,25} Reported nonadherence rates for DM populations within managed care organizations are 20%-50%, 23,26-30 and only 50% of patients have been shown to take 80% or more of their DM or cardiovascular medication.^{26,30} Long-term persistence to antihypertensive therapy among DM patients remains low as well,6,27 with reported adherence rates between 77% and 80%, 13,23,27 and a reported medication possession ratio to ACEI/ARB therapy of 0.77.8 Interventions to improve adherence to ACEIs/ARBs among this high-risk population are greatly needed.

Motivational interviewing (MI) is a patient-centered communication style used to strengthen motivation and commitment for change³¹⁻³³ in a supportive, collaborative, empathetic way.³⁴ MI has demonstrated effectiveness in improving adherence³⁵⁻³⁷ and is recommended by the American Association of Diabetes Educators.³⁸ MI is a tactic employed by many health care professionals across the spectrum in an effort to improve medication adherence and overall health outcomes, and pharmacists are uniquely positioned to provide such services. Studies have shown that pharmacists are beneficial in improving adherence among patients with DM³⁹⁻⁴³ and HTN,⁴⁴⁻⁴⁸ and among older adults.^{40,49,50} To enhance adherence, pharmacists can counsel patients about their chronic disease and treatment, monitor medication use, and communicate with other health care professionals.^{5,40,51} Although pharmacy students have the knowledge and experience to provide similar services under pharmacist supervision, literature examining the potential influence of student interventions on adherence is lacking. The objective of this study was to examine the effect of a MI-based telephone intervention conducted by pharmacy students in improving adherence to ACEIs/ARBs among Medicare Advantage Plan (MAP) patients with DM and HTN.

Methods

Study Design

A prospective study was conducted using members enrolled in a Texas-based MAP. University of Houston (UH) College of Pharmacy fourth-year pharmacy students, who participated in 6-week advanced pharmacy practice experiential (APPE) rotations supervised by preceptors and pharmacy residents at the MAP, conducted/implemented the intervention.

Data Description

Several computerized de-identified data files, which included a membership file, a member summary file, an institutional claims file, a professional claims file, the MAP's care management system (CCMS), and a pharmacy claims file, were available for analysis. Membership and member summary files included demographic, Centers for Medicare & Medicaid Services (CMS) risk scores, and yearly cost data for beneficiaries. Institutional claims included inpatient information claims with diagnostic information such as International Classification of Diseases, Ninth Revision, Clinical Modification (ICD-9-CM) codes and procedure information such as Current Procedural Terminology (CPT) codes. Professional claims contained outpatient encounter information as ICD-9-CM codes and CPT codes. Pharmacy files contained Part D pharmacy data provided by the MAP's pharmacy benefits manager and included patient and drug identifying information, fill dates, days supply, quantity dispensed, and dosing information for each prescription filled.

Inclusion Criteria

Subjects were included in the study if they had (a) enrollment in the MAP's drug plan between June 2014 and December 2015; (b) a diagnosis for DM and HTN identified through CCMS ICD-9-CM codes 250 and 401 between June 2014 and December 2014; and (c) a prescription filled for an ACEI or ARB medication during June 2014. Exclusion criteria included disenrollment from the plan, death before study end, a dementia diagnosis, and an ACEI/ARB use contraindication such as angioedema, hyperkalemia, and renal artery stenosis^{52,53} (identified by ICD-9-CM codes). This exclusion criteria was applied because patients with dementia have cognitive impairment that leads to noticeable decline in memory and thinking skills; therefore, adherence interventions to the general population cannot be effectively applied to this population.⁵⁴ In addition, even though ACEIs and ARBs are the most frequently used classes of antihypertensive treatments, their use can also be associated with functional renal insufficiency and/or hyperkalemia. Therefore, ACEIs/ARBs should be discontinued or switched when the patients have these conditions.

Nonadherent patients were identified by evaluating 6-month refill data during June 2014 through December 2014. Patients were categorized as nonadherent if they had a proportion of days covered (PDC) < 0.80 for their ACEI/ARB. If they switched to another ACEI or ARB, they were still considered to be covered. PDC is commonly used as a measure of adherence, which is defined as the number of follow-up days covered with medication divided by the total number of days in the follow-up period,^{55,56} and 0.80, or 80%, is generally considered adequate to achieve BP control.^{6,23}

Protection of Human Subjects

The protocol was approved by the UH institutional review board (IRB). Patients consented to the study before implementation of the intervention in accordance with all IRB requirements.

We randomly selected 75% of those patients meeting eligibility criteria identified as nonadherent to serve as potential subjects for the intervention group, and 25% of patients to serve as potential subjects for the control group using a random number generator program. More subjects were randomly selected as potential intervention subjects than controls to ensure that we could reach a minimum of 250 consenting patients by phone as determined by the sample size determination below. Baseline characteristics were equally distributed in these 2 groups.

Sample Size Determination

Using the G*Power 3.1 statistical software,⁵⁷ and a primary endpoint of postintervention adherence rate, we determined a total sample of 459 subjects was adequate to provide 90% power to detect significance with a 0.15 effect size (relatively small effect size) for a two-tail analysis t-test at 0.05 α level. A total of 491 subjects were needed to provide 90% power to detect significance with a 0.17 effect size (relatively small effect size) for a chi-square analysis at a 0.05 α level. A total of 409 patients were needed for a two-tail logistic regression analysis at a 0.05 α level and 0.10 β level (90% power) and for a 1.5 odds ratio (OR), and approximately 230 patients were needed for a multiple linear regression analysis with an effect size of 0.10. As a result, a total of at least 500 patients (250/group) were considered an adequate sample size given the various statistical tests and to have enough power to detect differences in case of a 5%-7% patient disenrollment from the plan (the usual rate in previous years is approximately 3%-5% per year).

MI Training

MI training for the pharmacy students was conducted over 3 days (June 5-7, 2015) by an MI trainer who is a member of the Motivational Interviewing Network of Trainers. Students rotating at the MAP between June 2015 and April 2016 attended the training. In the first 2 days, training included lecture, discussion, demonstration, videos, and observed role playing of MI skills. On the last day, all students conducted 3 simulated telephone conversations with trained standardized patients (SPs). SPs portrayed problems with adherence, like forgetfulness or fear of side effects. These problems were identified in this patient population in our previous study.⁵⁸

The practice calls were recorded for evaluation and feedback. The trainer evaluated each student's proficiency using a 7-point Likert-type scale (1 = poor/never to 7 = excellent/always) for MI skills and MI spirit, such as expressing empathy, using reflective listening, and eliciting change talk. General and individual feedback was given to students after evaluating the recorded calls. Students were then ready to begin implementing the intervention.

A total of 11 students at the MAP between June 2015 and April 2016 participated in the MI training. Mean scores of the trainer's evaluation of the student performances indicated average use of MI techniques, mostly ranging from 3 to 4 on a 7-point scale, where 7 is maximum. Students received general and individual feedback from the trainer. Overall, the feasibility of the 3-day MI training session for pharmacy students was evaluated and students demonstrated beginning MI proficiency.

Intervention

The intervention consisted of 6 MI phone calls conducted by the specially trained pharmacy students on rotation at the MAP. All students were supervised by an MI-trained pharmacist or pharmacy resident. Call recipients were randomly selected from the list of identified nonadherent patients and randomly assigned to the intervention group until a target of 250 patients was reached. Follow-up monthly calls with nonadherent patients were conducted until 6 months after the initial call.

During the initial call, students first confirmed the diagnosis, medication name, and dosage with the patient and followed a general outline based on the Ask-Provide-Ask approach,³² a pharmacist adaptation for the Elicit-Provide-Elicit approach of MI.⁵⁹⁻⁶³ Using MI, the students helped patients identify barriers and helped guide the patient to develop a plan to deal with these barriers. After each call, students recorded the date and wrote a summary of the interaction with the patient, documenting the main barriers identified in the conversation and the plan made with the patient to achieve therapy goals. The summary information of the initial call was reviewed by students with a supervising pharmacist at the MAP before each monthly follow-up call. During the follow-up calls, students discussed the status of identified barriers, as well as plans and any additional problem solving. They provided encouragement and reinforced with the patient strategies discussed in the previous calls. Follow-up information was recorded and used for subsequent calls. For every patient who received the initial phone call, 2 patients were randomly selected from the list of patients identified as nonadherent and assigned to the control group (n = 500).

Outcome Measures

An index date was defined as the initial intervention call date. For controls, the date when initial calls started was considered the index date. Refill data during the 6 months following the index date was evaluated to calculate adherence rates for both the intervention and control groups as PDC. PDC was calculated as the number of days during the 6 months on which the ACEI/ARB was available to the participant (total days supply) divided by the total number of days in that period, which was 180 days in this study. The PDC was truncated at 1.00 by removing any days of therapy extending beyond the analysis period, with a resulting ratio between 0 and 1.

ACEIs or ARBs were considered together. In the event that a patient switched from an ACEI to an ARB or vice versa, the days supply for both the ACEI and ARB were summed. For a patient who had an ACEI or ARB added to his or her existing ARB or ACEI, the adherence measurement was considered for the initial drug.

Patients in both groups were also categorized as adherent and nonadherent during the 6-month follow-up period based on PDC \geq 0.80 to create a categorical outcome variable. PDC is used for measuring medication adherence in the Medicare Star Ratings program; 80% is the cutoff used to define medication adherence for the Medicare Star Ratings program to monitor and provide incentives to health plans with good scores on Medicare Star metrics.⁶⁴ Discontinuation was also assessed, defined as no record of an ACEI/ARB during the 6 months following the initial intervention call.

Other Measures and Variable Definition

Baseline characteristics that could influence adherence and their comparability between the intervention and control groups were evaluated based on the index date. These characteristics included age; gender; number of other medications used/day; regimen complexity (defined as number of prescribed doses/day multiplied by number of medications, as used by Odegard et al. [2013, 2012])^{42,43}; income status (low-income subsidy [LIS] vs. no subsidy); prescriber specialty (primary care practitioner vs. specialist); comorbidities during the 6-month baseline period, including heart failure (HF), history of myocardial infarction, coronary artery disease, stroke, end-stage renal disease, and depression (categorized as having comorbidity vs. not having comorbidity); previous hospitalization during the 6 months before the index date versus no hospitalization; baseline adherence PDC during the 6 months before the index date; and the CMS risk score. The CMS risk score accounts for disease severity and medication burden and is calculated based on data from a pool of beneficiaries to estimate the average predicted costs for each of the component factors (e.g., gender, age, low-income status, individual disease groups). It consists of 189 disease classifications for use in risk adjusting of clinical outcomes in Medicare populations.^{65,66}

Statistical Analyses

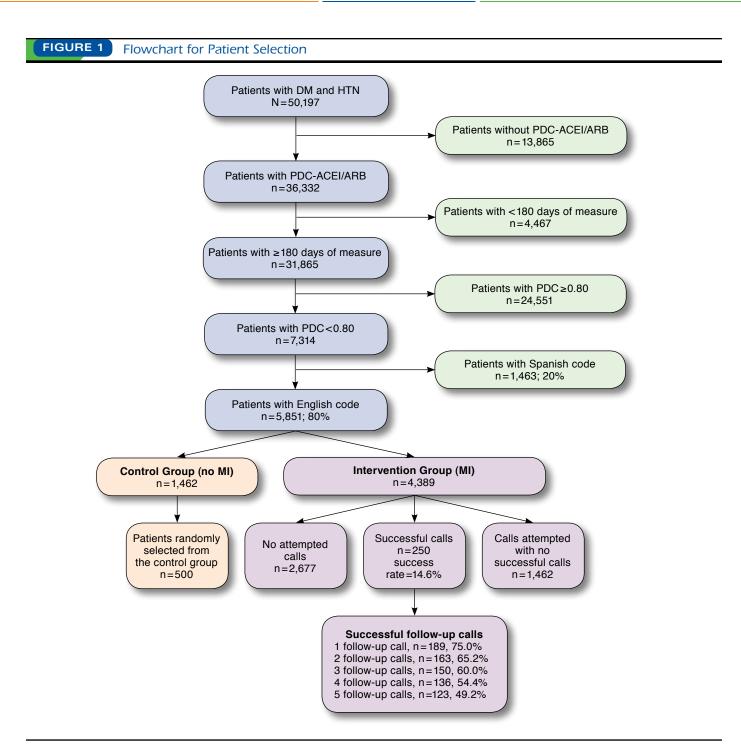
Baseline characteristics were compared between the 2 study groups (intervention vs. control) using Student t-tests for continuous variables and chi-square tests for categorical variables. PDCs for the invention and control groups during the 6 months following the intervention were compared using a t-test. Multiple linear regression with the 6-month PDC outcome was used to evaluate the intervention effect controlling for baseline characteristics not equally distributed between the 2 study groups. A logistic regression model with the categorical outcome based on a PDC≥0.80 cut-off was used to evaluate the intervention effect, adjusting for any imbalance in baseline characteristics. Logistic regression was also performed to assess the association between medication discontinuation with the outcome variable of discontinuation versus continuation, controlling for baseline characteristics. All statistical analyses were performed using SAS software version 9.3 (SAS Institute, Cary, NC) at a priori significance level of 0.05.

Results

Data Analysis Before Intervention

Data were obtained and analyzed between June 1 and June 21, 2015. A total of 50,197 patients were identified as having a diagnosis of DM and HTN. From this population, 36,332 were identified as having pharmacy claims for ACEIs or ARBs, with 31,865 having at least 180 days (6 months) of follow-up; 7,314 patients had a PDC < 0.80, with 5,851 (80%) coded as English-speaking.

Approximately 5,851 patients meeting eligibility criteria were randomly selected in a 3:1 ratio to serve as potential subjects for the MI intervention group and the control group, resulting in 4,389 patients being placed in the intervention group and 1,462 patients in the control group. Results are presented in Figure 1. Patient characteristics such as gender, age, and baseline PDC value were equally distributed between the intervention and control groups, and no significant differences were detected (P>0.05).



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ACEI = angiotensin-converting enzyme inhibitor; ARB = angiotensin II receptor blocker; DM = diabetes mellitus; HTN = hypertension; PDC = proportion of days covered; MI = motivational interviewing.

Intervention Calls

Initial calls by MI-trained pharmacy students, under pharmacist supervision, were completed between June 22 and October 29, 2015. More than 1,700 calls were attempted to reach the target

of 250 successful calls with consenting patients (success rate: 14.6%). On average, 18-20 successful calls were made per week by 2 students (averaging 9-10 per student). The average length of the initial call was approximately 12-13 minutes. Main

	Total N = 743 Frequency (%)	Intervention Group n=248 n (%)	Control Group n=495 n (%)	P Value
Age, years (±SD)	69.79 (±10.23)	72.35 (±10.01)	68.51 (±10.11)	<0.001ª
Gender				
Female	418 (56.26)	164 (39.23)	254 (60.77)	< 0.001ª
Male	325 (43.74)	84 (25.85)	241 (74.15)	
Specialty group				0.364
Specialist	374 (50.34)	119 (31.82)	255 (68.18)	
Primary care	369 (49.66)	129 (34.96)	240 (65.04)	
Regimen complexity	23.86 (21.04)	26.28 (20.81)	22.65 (21.07)	0.026ª
Alc lab value before intervention	6.52 (3.52)	6.59 (3.43)	6.47 (3.61)	0.784
Health plan				0.377
Low-income subsidy	451 (60.70)	145 (32.15)	306 (67.85)	
Other	292 (39.3)	103 (35.27)	189 (64.73)	
Heart failure	102 (13.73)	45 (44.12)	57 (55.88)	0.013ª
Depression	97 (13.06)	28 (28.87)	69 (71.13)	0.312
Stroke	33 (4.44)	33 (100.00)	0	<0.001ª
Myocardial infarction	14 (1.88)	3 (21.43)	11 (78.57)	0.338
Previous hospitalization				0.234
Yes	251 (33.78)	91 (36.25)	160 (63.75)	
No	492 (66.22)	157 (31.91)	335 (68.09)	
Number of medications at time of intervention (±SD)	10.93 (±6.75)	12.25 (±7.11)	10.27 (±6.47)	< 0.001ª
CMS risk score (±SD)	1.44 (±1.08)	1.75 (±1.12)	1.28 (±1.03)	<0.001ª
PDC before intervention (±SD)	0.6144 (±0.1505)	0.6348 (±0.1330)	0.6042(±0.1577)	0.005ª
PDC after intervention (±SD)	0.5969 (±0.3705)	0.6606 (±0.3376)	0.5650(±0.3822)	< 0.001ª

A1c=hemoglobin A1c; CMS=Centers for Medicare & Medicaid Services; PDC=proportion of days covered.

reasons for unsuccessful calls included call back requested (5%), declined (16%), left voicemail (22%), no answer (24%), disenrolled from the MAP (5%), and wrong/disconnected number (13.40%). The follow-up calls were ongoing until April 2016, with an average length of 5-7 minutes.

Approximately half of the patients (123, 49.2%) completed all 5 follow-up calls, while 136 (54.4%) completed 4, 150 (60%) completed 3, 163 (65.2%) completed 2, and 189 (75%) completed 1 follow-up call. The main barriers identified in calls included denial of nonadherence (32%) and forgetfulness (23.6%). Other reported barriers included high cost, modified (and often confusing) dosage instruction, high pill burden, side effects (perceived or experienced), transportation, and lack of disease state knowledge. The way each barrier was addressed is discussed in our previous study.⁵⁸

Multivariate Models Sample

The cohort included in the multivariate regression models consisted of 743 patients, with 248 in the intervention group and 495 in the control group. Seven patients were excluded because of missing values. The baseline sample characteristics are presented in Table 1.

Adherence

In the linear regression model, which examined factors associated with better adherence, the MI phone call intervention for patients who completed 2 or more follow-up calls (3 MI phone calls or more in total) was significantly associated with better adherence (β =0.0604, *P*<0.001). Higher medication adherence before intervention was also positively associated with better adherence after the intervention (β =0.2234, *P*<0.001). The number of medications taken by patients at the time of intervention was positively associated with medication adherence after intervention (β =0.0059, *P*=0.010). However, patients with HF (β =-0.1259, *P*=0.002) and depression (β =-0.0987, *P*=0.014) were less likely to adhere. These findings are summarized in Table 2.

In the logistic regression model, patients receiving 2 or more follow-up phone calls were more likely to become adherent after the intervention (OR: 1.53; 95% confidence interval [CI] = 1.02-2.28, P = 0.009). Patients with higher medication adherence before the intervention (prior PDC ≥ 0.50) were more likely to become adherent after the intervention (OR: 3.10; 95% CI = 1.98-4.86; P < 0.001). Patients taking a higher number of medications at the time of the intervention were more likely

TABLE 3

Variable	β Coefficient	P Value
Intercept	0.2567	0.017a
Baseline PDC	0.2234	<0.001a
Intervention	0.0604	<0.001ª
Number of medications at time of intervention	0.0059	0.010ª
CMS risk score	-0.0155	0.278
Regimen complexity	-0.00003075	0.325
Age	0.0020	0.135
Gender	0.0041	0.877
Heart failure	-0.1259	0.002ª
Depression	-0.0987	0.014ª
Previous hospitalization	-0.0265	0.385
Health plan	0.0031	0.908
Stroke	-0.0073	0.911
Myocardial infarction	0.0550	0.569
Specialty group	-0.0338	0.196

Result of Linear Regression Model to

to be adherent (OR: 1.03; 95% CI=1.002-1.060; P=0.033). However, patients with depression were less likely to be adherent (OR: 0.42; 95% CI=0.24-0.70; P=0.001). The results of the logistic regression model are presented in Table 3.

Results of the logistic regression for medication discontinuation showed that patients receiving 2 or more followup phone calls were less likely to discontinue (OR: 0.29; 95% CI=0.15-0.54; P<0.001) and patients with higher medication adherence before the intervention (prior PDC≥0.50) were less likely to discontinue (OR: 0.34; 95% CI=0.22-0.52; P<0.001). Additionally, patients with HF were more likely to discontinue (OR: 2.31; 95% CI=1.34-3.98; P=0.002). The results of the logistic regression model are presented in Table 4.

Discussion

TABLE 2

The MI-based telephone intervention by pharmacy students may be a promising intervention to improve adherence to ACEIs/ARBs among nonadherent patients enrolled in a MAP with comorbid DM and HTN. Patients who received the initial call and completed 2 or more follow-up calls had significantly less discontinuation and better adherence rates measured as PDC, as well as PDC>0.80, during the 6-month postintervention start.

There are more than 32 million Medicare patients receiving prescription drug benefits through MAPs or Medicare Part D.⁶⁷ Improving medication adherence among these patients can enhance CMS Star measures related to quality, which are linked to reimbursement,⁶⁸⁻⁷⁰ thus making adherence to ACEIs/ ARBs a priority for MAPs, given the health improvements and cost savings that can result.

Results of Logistic Regression Model to Examine Factors Associated with Better Adherence (N=743)

	1	
Variable	OR (95% CI)	P Value
Intervention		0.009ª
≥2 follow-up calls	1.53 (1.02-2.28)	
<2 follow-up calls	0.65 (0.38-1.10)	
No phone call	1	
Prior PDC		<0.001a
< 0.50	1	
0.50-0.80	3.10 (1.98-4.86)	
Age	1.01 (0.99-1.02)	0.315
Gender		0.830
Male	1.03 (0.75-1.42)	
Female	1	
Specialty group		0.180
Specialist	0.80 (0.59-1.10)	
Primary care	1	
Previous hospitalization		0.112
No	1	
Yes	0.74 (0.51-1.07)	
Regimen complexity	1.00 (0.990-1.001)	0.055
Number of medications at time of intervention	1.03 (1.002-1.060)	0.033a
CMS risk score	0.99 (0.83-1.18)	0.960
Depression	0.42 (0.24-0.70)	0.001ª
Health plan		0.967
Low-income subsidy	1	
Other	0.99 (0.71-1.38)	
Heart failure	0.62 (0.37-1.04)	0.073
Stroke	1.18 (0.53-2.62)	0.682
Myocardial infarction	1.55 (0.48-4.99)	0.461
aSignificance level a < 0.05		

^aSignificance level $\alpha < 0.05$.

CI=confidence interval; CMS=Centers for Medicare & Medicaid Services; OR=odds ratio; PDC=proportion of days covered.

Telephone-based interventions by different health care professionals have been reported to improve patient care among DM patients.⁷¹⁻⁷⁴ One study demonstrated the benefit of a telephone intervention by community pharmacists in improving adherence to antidiabetic medications and ACEIs.^{42,43} Pharmacists contacted patients who were 6 or more days late for refills, and in a brief amount of time, were able to successfully identify the patient's self-reported adherence barriers and provide assistance that improved overall adherence.⁴² Such interventions, however, cannot be widely implemented because of the high pharmacist cost and time constraints, as well as a pharmacy business model that is largely driven by the number of dispensed prescriptions.⁴²

However, pharmacy students, under the direct supervision of pharmacists, can be effective in providing similar services at a lower cost coupled with an enriched educational experience. Several studies have demonstrated the benefit of pharmacy student participation in a variety of clinical settings.⁷⁵⁻⁸³

to Examine Factors Associated with Medication Discontinuation (N=743)				
Variable	OR (95% CI)	P Value		
Intervention		< 0.001ª		
<2 follow-up calls	1.42 (0.81-2.47)			
≥2 follow-up calls	0.29 (0.15-0.54)			
No phone calls	1			
Prior PDC		<0.001ª		
< 0.50	1			
0.50-0.80	0.34 (0.22-0.52)			
Age	0.98 (0.96-1.01)	0.199		
Gender		0.798		
Male	0.95 (0.65-1.39)			
Female	1			
Specialty group		0.492		
Specialist	1.13 (0.78-1.65)			
Primary care	1			
Previous hospitalization		0.570		
No	1			
Yes	1.13 (0.73-1.75)			
Regimen complexity	1.00 (1.00-1.001)	0.837		
Number of medications at time of intervention	0.96 (0.93-1.001)			
CMS risk score	1.14 (0.93-1.38)	0.192		
Depression	1.41 (0.82-2.42)	0.208		
Health plan		0.715		
Low-income subsidy	1			
Other	0.93 (0.62-1.37)			
Heart failure	2.31 (1.34-3.98)	0.002a		
Stroke	0.82 (0.28-2.39)	0.727		
Myocardial infarction	0.99 (0.26-3.69)	0.990		
^a Significance level a < 0.05.				

Results of Logistic Regression Model

^aSignificance level α < 0.05.

TABLE 4

CI=confidence interval; CMS=Centers for Medicare & Medicaid Services;

OR = odds ratio; PDC = proportion of days covered.

An economic benefit was demonstrated in a previous study in which student interventions of patient education, order clarification, dosing adjustments, and lab monitoring yielded an overall savings of \$6,000-\$24,000 at a psychiatric hospital.⁸⁰ Pharmacy students conducting an interactive educational program on safe use of over-the-counter medications among older adults demonstrated positive intentions for behavior change among participants.⁸¹ One study examined the impact of a counseling program among HF patients led by pharmacy students and residents, and 89% of patients receiving the intervention reported better medication understanding. Many medication errors were also prevented, with a cost avoidance of \$4,241.82 A 5-minute consultation by a pharmacy student to improve knowledge about influenza and tetanus-diphtheriapertussis vaccines among 198 participants showed that a targeted consultation can increase vaccination rates and change public views.⁸³ Benefits described in these studies, however, were based primarily on self-reporting by students; our study examined refill data in a real-world environment to capture the potential improvement.

Not all pharmacist interventions, however, have led to improvements in adherence.^{84,85} A Medicare Part D telephone medication therapy management program did not document any medication adherence benefit despite reported cost savings.^{86,87} A brief pharmacist telephone intervention in a similar population of nonadherent patients with DM and HTN resulted in significantly better PDCs during the 6 months following the intervention, but the overall PDC rates in both the intervention and control groups were still lower than the recommended 80%.

Enhancing adherence to clinically meaningful values requires more than a brief pharmacist call. The style of communicating information to the patient can influence the intervention effectiveness. On the basis of our findings, incorporating MI strategies with follow-up calls to address barriers can be more influential in forming a sustainable behavioral change to improve adherence.

The intervention conducted in this study was based on the trans-theoretical model of behavioral change (TTM) in combination with MI techniques. The TTM describes behavior change as a progression through several stages,⁸⁸⁻⁹⁰ including the precontemplation, contemplation, preparation, action, and maintenance stages.⁸⁸⁻⁹⁰ TTM-based interventions have been beneficial among DM patients at different stages of readiness.⁹¹ MI strategies promote self-management behaviors such as medication adherence³⁴ by eliciting and strengthening the patient's internal motivation to change.^{92,93}

MI is well suited for delivery by phone^{34,75} and can be delivered by trained students.⁹⁴ MI promotes behavior change by contrasting the current behavior of nonadherence to desired goals of better outcomes and improved quality of life in a way that is supportive, collaborative, empathetic, and evocative. MI acknowledges autonomy; e.g., that the patient has every right to make no change, but uses a guided communication style that invites patients to consider their situation and find solutions to what they identify as problematic to facilitate a positive change while respecting patient autonomy.⁹³ Counseling can be based on patient readiness and stage of behavioral change to maximize engagement.³⁴

MI has been used to promote behavioral changes in alcohol abuse,⁹⁵ smoking cessation, weight loss, drug addiction,⁹⁶ selfcare for DM patients,^{93,97} and enhanced medication adherence with several chronic illnesses such as HTN,³⁵ asthma,³⁶ and HIV.^{37,98} Offering MI training during student introductory practice experiences and APPEs can help incorporate MI practices into the delivery of pharmaceutical care across various settings, with graduating pharmacy students entering the work force as well trained in these behavioral change techniques.

It is worth noting that the effectiveness of MI interventions can also be influenced by several other factors, such as the length and number of intervention sessions, MI delivery mode, and provider's credentials.⁹⁹ Menon et al. (2011) demonstrated that a single MI session may not be sufficient for a desired behavioral change.¹⁰⁰ Our study evaluated the feasibility and effectiveness of using 3 MI telephone intervention sessions to improve adherence.

Additionally, some comorbidities, including depression and HF, were negatively associated with better adherence in this study. The social isolation feelings accompanying depression may reduce cognitive functioning, which can affect patient willingness and motivation to follow treatment protocol, resulting in lower adherence, as shown in previous literature.^{2,101}

Nonadherence among patients with HF is common,^{102,103} possibly because of the increased regimen complexity and number of medications.¹⁰⁴ Previous research has been inconsistent regarding the influence of pill burden on adherence.¹⁰⁵⁻¹⁰⁷ In this study, higher pill burden was associated with improved adherence, which could be because patients with more medications may be sicker and therefore more attentive to taking their medications.¹⁰⁷ The baseline adherence before the intervention was also associated with better adherence, consistent with previous reports,¹⁰⁸⁻¹¹⁰ even though the baseline adherence of the patients in this study was below 80%.

Limitations

Several limitations to this study should be discussed. Prescription filling is not a guarantee that the patient actually took the medication, but refill rates are considered an acceptable measure of overall adherence.^{2,6} The study data did not capture prescriptions paid for by cash or verify the use of samples. While the initial random selection of the potential subjects for the intervention and control groups resulted in equal distribution of the major demographic variables between the groups, many patients randomly selected from the intervention group did not take the call or consent to the study, resulting in imbalances in some baseline characteristics. We later controlled for that in the multivariate analyses.

We also conducted analysis to test the difference in baseline sample characteristics among patients selected for the potential intervention group, across 3 groups: the group of patients with successful calls, the group of patients attempted to be reached but without successful calls, and the group of patients without attempted calls. There was no statistically significant difference in the baseline characteristics (age, gender, baseline PDC) across these different groups of patients. However, differences in other factors may exist between the patients who were willing to participate in the calls compared with those who were not willing or were not contacted. Such differences can only be addressed using a randomized design with an intentto-treat analysis. Additionally, most patients in the intervention group completed 2 follow-up calls, but approximately 50% of consenting patients finished all 5 follow-up calls. Planning an intervention with 2 to 3 follow-up calls may be more feasible for the future and still effective in improving adherence. Information on some potential confounders, like side effects, health literacy, and perceived health, was noted.

We also were unable to account for hospitalizations during a follow-up period. The patients in this study were all Texas patients enrolled in the drug plan, and generalizability of findings is limited to similar subpopulations.

Conclusions

Patients receiving 2 or more calls had significantly better adherence and less discontinuation during the 6 months following initial calls compared with those who did not receive calls. This finding indicates that the MI-based telephone intervention by pharmacy students may be a promising intervention to improve adherence. Future research should examine the sustainability of the intervention effect for longer time periods and its influence on associated clinical outcomes.

Pharmacy students are a valuable, cost-saving resource that can be used to improve medication adherence. Participating in MI counseling opportunities can enhance skills they can carry into a variety of future practice settings.

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