

Is there a nonadherent subtype of hypertensive patient? A latent class analysis approach

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Abstract: To determine subtypes of adherence, 636 hypertensive patients (48% White, 34% male) reported adherence to medications, diet, exercise, smoking, and home blood pressure monitoring. A latent class analysis approach was used to identify subgroups that adhere to these five self-management behaviors. Fit statistics suggested two latent classes. The first class (labeled “more adherent”) included patients with greater probability of adhering to recommendations compared with the second class (labeled “less adherent”) with regard to nonsmoking (97.7% versus 76.3%), medications (75.5% versus 49.5%), diet (70.7% versus 46.9%), exercise (63.4% versus 27.2%), and blood pressure monitoring (32% versus 3.4%). Logistic regression analyses used to characterize the two classes showed that “more adherent” participants were more likely to report full-time employment, adequate income, and better emotional and physical well-being. Results suggest the presence of a less adherent subtype of hypertensive patients. Behavioral interventions designed to improve adherence might best target these at-risk patients for greater treatment efficiency.

Keywords: adherence, hypertension, latent class analysis, self-management

Introduction

Hypertension is a major public health problem that affects 65 million Americans.¹ Several trials have led to established clinical guidelines, including following medication regimens, dietary and exercise recommendations, and smoking cessation.^{2–9} Adherence to these guidelines is considered necessary to optimize hypertension control. Unfortunately, hypertension control remains suboptimal, leading to an increase in the incidence of sequelae of hypertension, including heart failure and end-stage renal disease.^{10–14}

Key to improving hypertension control is identification of people at risk for nonadherence with the various components of a hypertension regimen. Several investigations attempting to identify such individuals^{15–21} share one major limitation, ie, they treat adherence to each recommendation as a discrete outcome. This method ignores the potential overlap between the different behaviors, in that unhealthy behaviors tend to coexist and may have a multiplicative adverse impact on health.^{10,22} Only a few studies have examined the collective impact of multiple unhealthy behaviors on health outcomes. Weir et al used cluster analysis to identify subtypes of hypertensive patients based on their medication use, lifestyle behaviors, and health beliefs.¹⁰ Their results indicated the presence of four subtypes. The first group effectively managed medication and lifestyle recommendations; the second group effectively managed medications, but not lifestyle recommendations; the third group reported nonadherence to medication, diet, and exercise, but had better adherence to smoking and alcohol recommendations; and the

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fourth group reported nonadherence to all recommendations. The former two groups were found to have better health outcomes than the two latter groups. The key strength of this study was its focus on identifying the subtypes of nonadherence for hypertensive patients. This would greatly improve the ability to manage hypertension clinically by allowing greater focus on patients at risk for poor self-management. However, cluster analysis has been criticized because of the arbitrary way in which groups are created.²³

Methodological limitations of cluster analysis may be avoided by using a latent class analysis (LCA) approach. LCA may be thought of as a categorical variable analog to factor analysis.²⁴ It assumes that a categorical latent variable, or “type”, causes a response on multiple observed variables. LCA has several advantages over traditional cluster analysis. LCA allows model fit criteria and rigorous statistical testing, whereas cluster analysis employs a relatively arbitrary method of clustering.^{23,25} An additional advantage is that LCA can be used with categorical data. Because many hypertension recommendations exist on a dichotomy (eg, stopping smoking), this can be an important methodology to employ with adherence behaviors. LCA may provide an important methodology for identifying subtypes of nonadherence, while circumventing limitations in the methodologies that have been employed in the literature.

The purpose of this study was to determine the presence of underlying adherence subtypes in hypertensive patients using an LCA approach. We were interested in identifying subtypes of hypertensive patients based on their self-management behaviors. Secondary aims were to characterize the subtypes based on a variety of demographic and psychosocial variables, ie, age, race, gender, financial situation, employment, quality of life, social support, stress, and attitudes towards hypertension. Characterization of the nonadherent subtype would potentially identify specific psychosocial factors worth targeting in adherence-improving interventions. Finally, we examined the association of adherence subtypes on blood pressure (BP) levels.

Methods

Participants

Six hundred thirty-six hypertensive patients were recruited from two Duke University Medical Center primary care clinics to participate in the Take Control of Your Blood Pressure study.²⁶ Briefly, this study tested two interventions (tailored behavioral intervention and BP self-monitoring) in a sample

of hypertensive patients.²⁷ The present study represents secondary, cross-sectional analyses of the baseline data from this trial. Participants were 66% female, and were evenly split between White and African American races (48.4% and 49%, respectively; 2.6% were classified as “other” for these analyses). All procedures were approved by the Institutional Review Board of Duke University Medical Center.

Inclusion and exclusion criteria

Patients were included if they had documented hypertension according to medical records (ICD-9 codes 401.9, 401.0, 401.1), if they had been enrolled in one of the two primary care clinics for at least a year, and if they were using a hypertensive medication, eg, an angiotensin-converting enzyme inhibitor or a beta-blocker. Patients were excluded if they were not on a BP medication; a family member was already enrolled in the study; they did not live in an eight-county catchment area; were receiving kidney dialysis; were pregnant or were planning to be pregnant; had an arm circumference greater than 17 inches and wrist circumference greater than 8.5 inches; had been hospitalized for a stroke, myocardial infarction, or coronary artery revascularization within the previous three months; had been diagnosed with metastatic cancer or dementia; resided in a nursing home or received home health care; did not speak or understand English; were enrolled in another hypertension study; were not receiving the majority of their healthcare through Duke University; had severely impaired hearing or speech; and/or had a history of organ transplantation. Using this method, 7646 potentially eligible patients were identified using medical records; 1325 were deemed eligible per the above criteria and were contacted. Six hundred and thirty-six patients (48% of the 1325) agreed to participate and were enrolled. For more details regarding the study design, refer to Bosworth et al.²⁶

Blood pressure measurement

Patients underwent BP screening involving two successive digitally-derived BP values taken by study staff. These were averaged to determine baseline systolic and diastolic BP (SBP and DBP, respectively). We were further interested in examining BP control. Using JNC 7 guidelines, SBP control was defined as $SBP \leq 140$ mmHg and DBP control as $DBP \leq 90$ mmHg.

Adherence measures

A variety of measures were used to assess adherence to medications, diet, exercise, smoking, and home BP monitoring. All adherence measures were dichotomized. Exercise,

medication adherence, and smoking recommendations were dichotomized at the point which best represented contemporary public health recommendations. Details regarding the cutoffs are provided below.

Medication adherence was assessed using the Self-reported Medication Taking Scale.²⁸ This four-item measure assesses medication-taking behavior. Respondents rate whether they forget to take medications, are careless about taking their medications, or stop taking their medications based on whether they feel better or worse. Items are rated on a four-point scale, from “strongly agree” to “strongly disagree”. In the current study, the internal consistency as measured by Cronbach’s alpha was 0.85. As in prior research, participants who endorsed at least one item by answering either “strongly agree” or “agree” were considered nonadherent to medications.

To define adherence to exercise recommendations, patients were asked: “On average, how much time per week do you spend on aerobic or body movement activities, such as brisk walking, jogging, or running, that elevates your heart rate for 20 minutes and makes you sweat/perspire?” Five answer options ranged from “never” to “more than four hours/week”. Participants were considered to be nonadherent to exercise if they reported exercising <2 hours/week, consistent with current recommendations.²⁹

Participants were asked: “On a scale of 1 to 10, with 1 being not at all hard and 10 being extremely hard, please rate how hard it is for you to follow recommendations to improve your blood pressure regarding diet”. Participants were considered nonadherent to dietary recommendations if they rated their difficulty with dietary recommendations ≥ 5 on the 10-point scale.

Smoking status was determined by a single yes/no item asking whether participants were current cigarette smokers. Participants were considered nonadherent to smoking recommendations if they reported that they currently smoked.

Patients were asked if they owned a home BP monitor and, if so, how often they used the monitor. Answers ranged from “never” to “frequently”. Patients were considered nonadherent if they answered “never”.

Psychosocial measures

Because we were also interested in characterizing the patients, we collected data on a variety of demographic variables and psychosocial constructs. A demographic questionnaire was administered to determine participants’ age, gender, ethnicity, marital status, education, employment status, and financial situation. Data were reduced for the following variables: race (White versus African American), marital status (married

versus unmarried), education (equal to or greater than high school versus less than high school), and employment status (employed versus unemployed). Participants were asked about the number of people residing with them because cohabitation has been shown to have a positive effect on adherence.³⁰

Two subscales of the MOS Short Form 12 (SF-12) were used to measure emotional well-being and physical well-being.^{31,32} Emotional well-being was measured using the Mental Component Summary Scale (MCS), a five-item subscale that measures general mental health and its impact on daily functioning. Physical well-being was measured using the Physical Component Summary Scale (PCS), a five-item subscale that measures general physical health and its impact on daily functioning. The mean of each subscale is 50, the standard deviation (SD) is 10, and the reliability ranges from 0.77–0.97.^{31,32} To allow for clinical interpretation, MCS and PCS scores were divided by 10, which is 1 SD. These scores were subsequently used in regression models.

Social support was measured in two ways. First, instrumental support was measured by asking participants: “If needed, is there someone who could help you with tasks such as taking you to the doctor, fixing lunch, or home repairs?” Second, emotional support was measured by asking participants: “Do you have someone you feel close to, someone you can trust and confide in?” The instrumental and emotional support items were answered either “yes” or “no”. Participants were further asked how much contact they had with the person in whom they could trust or confide. This was rated on a four-point Likert scale ranging from “no contact” to “a lot of contact”.

Participants were asked how often in the past month they had experienced stress. Answers were rated on a five-point Likert scale ranging from “never” to “very often”.

Attitudes towards hypertension were assessed by asking participants to what extent they believed hypertension was a serious condition. Answers were rated on a four-point Likert scale, from “very serious” to “not at all serious”. They were also asked how worried they were about hypertension. Answers were rated on a 10-point scale, from “definitely not worried” to “extremely worried”.

Data analysis

All analyses were conducted using Mplus 4.1 (Muthen and Muthen, Los Angeles, CA). The analytical strategy in LCA involves identifying the fewest number of classes that explain adherence across the five recommended self-management behaviors. The optimal number of classes was determined by progressively increasing the number of classes and testing each subsequent model. Because there remains debate on

the best way to determine number of classes, a combination of Bayesian information criterion (BIC) and parametric likelihood ratio tests with bootstrapped values (LRT) was used to determine model fit. The BIC aims to balance model fit with model parsimony such that lower scores represent better model fit.²⁵ The LRT compares progressive iterations of the more parsimonious models ($k-1$ classes) against models with greater number of classes (k classes). To establish the number of classes using LRT, the models with the fewest number of classes where $P < 0.05$ were accepted.³³ Using this combined strategy, we assigned each participant to a permanent class, based on their highest posterior class membership probability of being in each class.

After establishing class membership, logistic regression analyses were conducted to examine the odds of being a member of the classes given responses to the psychosocial and demographic questions, and SBP and DBP levels. The class less likely to be adherent was used as the referent. Significance for regression analyses was set at $P < 0.05$.

Results

The sample of 636 patients with hypertension (66% female, 49% African American) is described in Table 1. At baseline, the mean SBP was 125.0 mmHg and mean DBP was 71.3 mmHg. Approximately 84% of participants reported that they were nonsmokers, 58% reported adherence to medication, 55% reported adherence to dietary recommendations, 40% reported adherence to exercise recommendations, and 13% reported current home BP monitor use.

To determine the presence of latent classes for adherence behavior, 1, 2, and 3 class models were tested. As

Table 2 Fit statistics of the latent class analysis models

Classes	Log-likelihood	BIC	Bootstrapped p value from LRT for $k-1$ classes
1	-1823.97	3675.07	
2	-1796.09	3663.19	$P < 0.001$
3	-1791.82	3692.38	$P = 0.15$

Abbreviations: BIC, Bayesian information criterion; LRT, likelihood ratio test.

shown in Table 2, a 2-class model was superior to both 1- and 3-class models by virtue of lower BIC and $P < 0.05$ associated with the LRT 2 versus 3-class model. Class 1 consisted of 34.4% ($n = 219$) of the sample. These participants were consistently more likely to report adherence to medications, nonsmoking, diet, exercise, and home BP recommendations. Class 2 consisted of the remainder 65.6% ($n = 417$) of the sample. Compared with participants assigned to Class 1, Class 2 participants had a lower probability of reporting adherence to the five behaviors. Based on this, Class 1 was classified as “more adherent” and Class 2 was classified as “less adherent”. Conditional probabilities are provided in Table 3 and represented in Figure 1. As can be seen in Figure 1, the patterns of adherence were similar in both classes, with most participants reporting current nonsmoking status, and fewest participants reporting home BP monitor use.

Next, logistic regression analyses were conducted to compare characteristics of the “more adherent” class to the “less adherent” class (Table 4). “More adherent” patients were more likely to report full-time employment (odds ratio [OR] 0.79, 95% confidence interval [CI]: 0.64–0.96) and fewer problems with finances (OR 0.77, 95% CI: 0.62–0.96). For every 10-point increase in emotional well-being, represented by MCS scores, the odds of a patient being “more adherent” increased by 24% (OR 1.24, 95% CI: 1.01–1.53). Similarly, for every 10-point increase in physical well-being,

Table 1 Baseline patient characteristics

n	636
Age (mean \pm SD)	61.25 \pm 12.32 (range 25–92)
Gender, n (%)	420 female (66%)
Race	48.4% White, 49.0% African American
Baseline SBP, mmHg	125.0 \pm 17.7
Baseline DBP, mmHg	71.3 \pm 10.8
Current smokers, n (%)	130 (16.4%)
Adherence to dietary recommendations, n (%)	346 (55.1%)
Adherence to exercise recommendations, n (%)	252 (39.62%)
Adherence to medication, n (%)	371 (58.4%)
BP monitor use, n (%)	84 (13.2%)

Abbreviations: BP, blood pressure; SBP, systolic blood pressure; DBP, diastolic blood pressure; SD, standard deviation.

Table 3 Conditional probabilities of class membership in the two-class mode

	Class 1: “More adherent”	Class 2: “Less adherent”
Takes medications as prescribed	0.755	0.495
Does not have difficulty adhering to dietary recommendations	0.707	0.469
Exercises at least 30 min/day, most days/week	0.634	0.272
Does not currently smoke cigarettes	0.977	0.763
Uses home blood pressure monitor	0.320	0.034

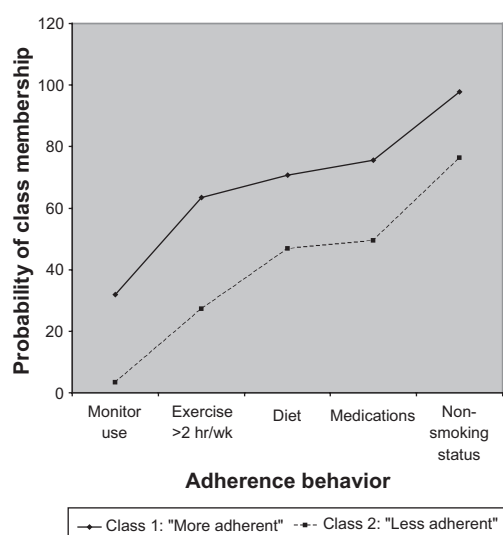


Figure 1 Conditional probabilities of being adherent to the five recommendations, given class membership.

represented by PCS scores, the patients' odds of being "more adherent" increased by 30% (OR 1.30, 95% CI: 1.06–1.59). Age, race, marital status, gender, or education levels did not appear to impact class membership. No differences in SBP, DBP, or BP control were found.

Discussion

This study sought to demonstrate the presence of subtypes of hypertensive patients who are either more or less adherent across several hypertension-related self-management behaviors. Two distinct subtypes of hypertensive patients were identified based on adherence to five self-management recommendations. Previous investigations have used LCA to show that health behaviors may be multidimensional.³⁴ However, to our knowledge, this is the first investigation to use LCA and demonstrated the presence of subtypes of adherence in hypertensive patients.

The next goal was to characterize the two classes. Previous investigations have focused on many factors that might impact nonadherence, with mixed results. Age, gender, and personality traits have all been examined as potential predictors or correlates of adherence, with weak and inconsistent results.^{20,35} Race has been more consistently predictive of adherence, with Whites demonstrating better adherence than African Americans.³⁶ It should be noted that the vast majority of these studies have focused on medication adherence, with little attention to other self-management behaviors. In this study, age, race, marital status, and gender were not related to class membership. These results may provide further

Table 4 Odds ratio of being a member of the "more adherent" class

Variable	OR	95% CI
Age (per 10-year increase)	1.16	0.97–1.40
Race (White versus Non-White)	1.34	0.86–2.12
Married (married versus unmarried)	1.31	0.80–2.06
Gender (female versus male)	0.77	0.48–1.24
Education (high school versus no high school)	1.12	0.97–1.30
SBP (per 5 mmHg increase)	1.04	0.98–1.11
DBP (per 5 mmHg increase)	0.96	0.87–1.07
SBP control	0.84	0.51–1.40
DBP control	0.77	0.48–1.23
SF-12 Physical Component Summary Score (per 10-point increase)	1.30*	1.06–1.59
SF-12 Mental Component Summary Score (per 10-point increase)	1.24*	1.01–1.53
Number of cohabitants	0.88	0.73–1.05
Instrumental support	1.11	0.45–2.73
Amount of contact with social support	1.05	0.70–1.58
Perceived stress	0.89	0.72–1.09
Employment status	0.79*	0.64–0.96
Financial situation	0.77*	0.62–0.96
Perceived seriousness of hypertension	0.88	0.59–1.33
Worry about hypertension	0.96	0.88–1.06
Availability of leisure time	1.06	0.91–1.24

Abbreviations: OR, odds ratio; CI, confidence interval; DBP, diastolic blood pressure; SBP, systolic blood pressure. * $p < 0.05$.

insight into the role of these traditional demographic variables in adherence as a latent class and might explain the contradictory results that have been documented in the literature when adherence behavior is examined individually.

Being more adherent was related to better physical and emotional well-being. We have previously reported that emotional well-being is related to adherence to diet and exercise and smoking recommendations.²⁰ These findings provide further support that lower emotional well-being, even if it does not meet clinical significance, may adversely influence adherence to all recommendations. This underscores the importance of assessing emotional well-being in primary care settings.

Results suggest that lack of employment and financial difficulties may be important barriers to optimally managing hypertension. Financial constraints may limit patients' ability to purchase medications. In addition, the necessity of working extra hours for income may limit the time patients can devote to exercise or limit access to resources such as gyms, personal trainers, or dietitians. Improving access to health care, which is a goal of "Healthy People 2010", may be critical in reducing the burden on both the individual and society as a whole.

Contrary to our hypothesis, adherence subtypes were not related to BP levels or BP control. The relationship between adherence and outcomes is complex, and may be influenced by dose, efficacy of treatment, response rates, and understanding of the disease.³⁷ Past investigations suggest that the impact of adherence on disease-specific medical outcomes may be limited even when the impact on other indices, such as emotional well-being, may be strong.^{10,37,38} Several explanations of our results may be posited. First, the sample in this study consisted of patients with well-controlled hypertension, as can be seen from the mean and SD of the sample BP values. Second, the sample reported high adherence to medications, perhaps the single most important factor in controlling hypertension in the short term. Third, our ability to detect causality may be hampered because adherence and BP were measured concurrently in this cross-sectional study. Given all these factors, the lack of relationship between the adherence classes and BP may illustrate that even within a highly adherent, well-controlled group of hypertensive patients, adherence behaviors may covary. Studies interested in understanding adherence should recognize that individually treating each behavior as unique ignores this covariance and likely inflates the Type I error. It is possible that previous studies linking individual behaviors to BP may not be as robust, given the potential of Type I error that results from multiple comparisons. Because this study utilized a novel approach to examining adherence, rigorous investigations are needed to examine the impact of nonadherence to multiple behaviors on BP levels in studies specifically designed to answer this question.

Our findings support the notion that the focus of adherence interventions can be potentially broadened to target multiple behaviors. Historically, interventions aimed at improving adherence target behaviors individually (eg, smoking cessation and exercise regimens). Inevitably, interventions targeting one behavior will be limited in their impact.³⁹ These interventions may also be time-consuming and costly, and may not reflect the relationship between various adherence behaviors. In reality, making changes to one health behavior may serve as a catalyst to making other lifestyle changes.⁴⁰ Based on such findings, recent interventions have focused on multiple health behavior changes,⁴¹ ie, interventions that may impact multiple behaviors simultaneously. These interventions may demonstrate an overall benefit even though their impact on an isolated behavior may be too small to detect if examined individually.

This study has some limitations. First, the cross-sectional nature of the data limits our ability to examine the predictive validity of the two subtypes. It is not clear whether these adherent/nonadherent subtypes would impact hypertension and other related outcomes prospectively. Second, the cross-sectional nature of the data does not allow us to determine predictors of class membership. Third, the use of multiple single-item measures raises questions regarding reliability and validity. Results of these analyses should be considered hypotheses-generating, given the potential psychometric difficulties of single-item measures. Fourth, our population was both relatively well-controlled and relatively adherent. Predictors of adherence behavior may not be the same in a more nonadherent population.

Despite these limitations, this study makes an important contribution to the extensive adherence literature in hypertensive patients. It proposes a unique way of characterizing hypertensive patients that may be both methodologically sound and clinically relevant. It also highlights the complexity inherent in obtaining adequate hypertension control. Further investigations are necessary to replicate these results, especially given the previous literature negating the presence of an “adherent personality”. Future investigations should focus on the predictive validity of the two subtypes as well as predictors of class membership. If replicated, these results provide support for designing interventions aimed at multiple health behaviors in the millions of Americans currently treated for hypertension.

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