Association of adherence measured by self-reported pill count with achieved blood pressure level in hypertension patients: a cross-sectional study

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Abstract

Background: Medication adherence plays an essential role in controlling blood pressure to reduce morbidity and mortality of hypertension disease. Thus, this study aimed to determine the association of medication adherence measured by self-reported pill count with blood pressure levels among patients at several community health centers in Surabaya.

Methods: Adherence was assessed using the pill count method by comparing the total number of antihypertension drugs taken with the prescribed drugs. The inclusion criteria involved hypertensive patients who received antihypertension drugs, specifically adults and elderly, except the pregnant woman. The patient blood pressure was measured by healthcare workers in the targeted community health centers. Descriptive and multivariable logistic regression analyses were performed to assess factors associated with medication adherence with blood pressure levels.

Results: A total of 264 hypertensive outpatients participating in this study, 77.65% of participants were adherent to antihypertensive drugs based on the pill count method, and 40.91% of participants had controlled blood pressure. Patients with uncontrolled blood pressure were about six times (adjusted odds ratio [AOR]: 6.15; 95% confidence interval [CI]: 2.694–14.039; \( P = 0.000 \)) more likely to have non-adherent medication than patients with controlled blood pressure. Reciprocally, non-adherent participants (pill count < 80%) were about six times (AOR: 6.081; 95% CI: 2.672–13.838; \( P = 0.000 \)) more likely to have uncontrolled blood pressure compared to adherent patients (pill count \( \geq 80\% \)). Age less than 40 years old (AOR: 5.814; 95% CI: 1.519–22.252; \( P = 0.01 \)) and having middle school educational level (AOR: 0.387; 95% CI: 0.153–0.974; \( P = 0.045 \)) were found to be independent factors associated with uncontrolled blood pressure.

Conclusions: The result showed that non-adherence to antihypertension drugs is associated with uncontrolled blood pressure. Then, age could be associated with uncontrolled blood pressure. Thus, pharmacists and other healthcare providers should pay attention to improving medication adherence and maintaining the controlled blood pressure.

Keywords: Hypertension, Blood pressure, Medication adherence, Pill count, Antihypertensive drug
Background
Hypertension is a non-communicable disease that is currently increasing in prevalence. Based on a national population survey in 2018, the prevalence of hypertension in Indonesia was 34.1% (age ≥18 years old) [1]. Furthermore, data from the World Health Organization (WHO) showed that around 1.13 billion people had hypertension in 2015, meaning that one out of three people worldwide was potentially diagnosed with hypertension. The hypertension case continues to increase and will be projected to 1.5 billion in 2025. Then, 9.4 million people will potentially die from hypertension and its complications every year [2]. In Southeast Asia, hypertension is a risk factor causing 1.5 million deaths per year [3].

Hypertension is defined as clinical blood pressure at or above 140/90 mmHg [4]. It is believed that managing the hypertension risk could be done by controlling the blood pressure. Controlled systolic blood pressure can reduce the risk of death, cardiovascular disease, stroke, and heart failure. One of the causes of low blood pressure control in hypertensive patients was the lack of adherence to the antihypertensive treatment [5, 6]. Although pharmacological therapy is available, medication adherence is known to be suboptimal because of many factors, such as age, knowledge about hypertension, motivation, and belief in treatment [7]. Other factors include therapeutic regimen [7, 8], the health system that guarantees health financing [9], and treatment information provided by health workers [10].

The importance of monitoring the adherence to antihypertensive drugs can help healthcare staff examine and control patients’ blood pressure. Thus, this study aimed to determine the association of medication adherence with the blood pressure of hypertensive patients. There are two methods of measuring adherence, direct and indirect methods [11, 12]. Direct method is carried out by observing the patients’ condition, such as measuring drug concentration in the patient’s blood samples. In contrast, indirect method for monitoring medication adherence is performed by giving an interview or a questionnaire, calculating the number of drugs, and monitoring drug use [13, 14]. This study was conducted using the pill count method (counting the remaining drug prescribed) to assess medication adherence.

Methods
Study design
This research was conducted in five community health centers (locally called Puskesmas), including Benowo, Jeruk, Ketabang, Tambak Rejo, and Gayungan. The community health center is the primary health care center covering a sub-district in Surabaya, East Java, Indonesia. Prospective data collection was done in 2 months (April and May 2020). Adherence was assessed by comparing the total number of prescribed antihypertension drugs and the remaining drugs when visiting the community health centers. A direct interview was performed to ask the patients about the number of remaining drugs and the sociodemographic characteristics (age, sex). Type of therapy and comorbidities were recorded from the patient’s medical record at community health centers. Blood pressure was measured by healthcare workers.

Participants recruitment
The study subjects were purposively selected by following the inclusion and exclusion criteria. The inclusion criteria involved all hypertensive outpatients, including adults and elderly, and currently on antihypertensive treatment. Meanwhile, we excluded patients aged <18 years, pregnant women, patients with mental disorders, and those diagnosed with hypertension for the first time.

Data analysis
The percentage of patient adherence was calculated by comparing the number of drugs consumed with the number of drugs that had to be consumed then multiplied by 100% [15–17]. In this study, a percentage of pill count <80% and 80–100% were considered non-adherent and adherent, respectively [15]. The blood pressure target in hypertensive patients refers to the hypertension guideline recommendation of Joint National Committee 8 (JNC 8). It further explains that hypertensive patients without comorbid diabetes mellitus and chronic kidney disease at the age of ≥60 years should achieve <150 mmHg systolic blood pressure and <90 mmHg blood pressure diastolic. Meanwhile, for patients aged <60 years, the target of achieving systolic blood pressure is <140 mmHg and systolic blood pressure is <90 mmHg. For hypertensive patients with comorbid diabetes mellitus and/or chronic kidney disease at all ages, the target is systolic blood pressure <140 mmHg and diastolic blood pressure <90 mmHg [18].

Statistical analysis
This study employed IBM SPSS ver. 20 (IBM Corp., Armonk, NY, USA) for the statistical analysis. A chi-square test was used to determine the association between demographic characteristics with medication adherence and blood pressure. Multivariable logistic regression models were fitted to identify the association of explanatory variables with the outcomes. Variables with a P-value less than 0.20 were considered for inclusion in a multivariable logistic regression analysis in which confounders could be controlled. The necessary assumptions of logistic regression were made by checking Hosmer and Lemeshow’s goodness of fittest statistics. Variables with a P-value less than 0.05 in the
multivariable logistic regression analysis were considered as statistically significant determinants. Adjusted odds ratio (AOR) with 95% confidence interval (CI) was calculated to measure the strength of the association between the explanatory variable and the outcome variable.

**Results**

A total of 264 patients on antihypertension treatment who participated in this study came from Benowo (40 patients), Jeruk (40 patients), Tambak Rejo (74 patients), Ketabang (40 patients), and Gayungan (70 patients). The majority of the study population was female (73.48%), aged 40–61 years old (51.13%), and educated up to high school (35.98%). Most subjects were prescribed a single antihypertensive agent (91.67%), and the remaining received a combination (8.33%). Regarding comorbidities, more than half of patients had diabetes and dyslipidemia (57.95%). The proportion of adherent and non-adherent patients were 77.65% and 22.35%, respectively. Then, according to the JNC 8 standard, patients were classified as having controlled (108 patients, 40.91%) and uncontrolled blood pressure (156 patients, 59.09%) (Table 1).

Tables 2 and 3 depicted the association of patients’ characteristics to medication adherence and blood pressure control, respectively (using the chi-square test). The result showed that medication adherence had association with blood pressure control only \((P = 0.000)\). Meanwhile, age and patient adherence could influence the blood pressure \((P = 0.005 \text{ and } P = 0.000, \text{ respectively})\). Then, sex, type of therapy, education level, and the presence of comorbid disease did not associate with adherence nor blood pressure.

In line with the previous finding, multivariable logistic regression also showed that patients with uncontrolled blood pressure were associated with poor medication adherence or vice versa (Tables 4 and 5; \(P = 0.000\)). Moreover, uncontrolled blood pressure was associated with those aged <60 years and educated up to middle school. Then, sex, type of therapy, and the presence of the comorbid disease did not associate with poor adherence or uncontrolled blood pressure (Table 5).

**Discussion**

This study examines the association of medication adherence with blood pressure control status among outpatients in primary health care facilities. It was evident that low adherence was significantly associated with uncontrolled blood pressure. Reciprocally, poor blood pressure control was significantly associated with non-adherence to antihypertension treatment. In line with this finding, several studies reported that highly adherent patients were more likely to have controlled blood pressure than those with lower adherence [19–21]. Although the proportion of adherent patients was high (77.65%), clinical consequences of suboptimal medication adherence are negligible, including uncontrolled blood pressure, accelerating disease progression, and increasing hospital admissions due to cardiovascular complications [22].

In this study, we could not find any significant association between medication adherence and patient characteristics (age, sex, type of therapy, educational level, and the presence of comorbidity) in both statistical analyses. In contrast, ample studies proved many independent predictors related to medication adherence. For instance, Khayyat et al. [21] reported that sex, age, and the presence of comorbid other diseases such as diabetes mellitus affect medication adherence. In another study by Kang et al. [23], it is stated that medication adherence

| Table 1 Percentage of baseline and clinical characteristics \((n = 264)\) |
|-----------------|-----------------|
| **Variable**    | **Frequency (%)** |
| **Age (yr)**    |                 |
| 20–40           | 19 (7.19)       |
| 41–60           | 135 (51.13)     |
| >60             | 110 (41.68)     |
| **Sex**         |                 |
| Female          | 194 (73.48)     |
| Male            | 70 (26.52)      |
| **Education level** |           |
| Elementary school | 73 (27.65)    |
| Middle school   | 37 (14.02)      |
| High school     | 95 (35.98)      |
| College         | 59 (22.35)      |
| **Comorbidity** |                 |
| No              | 154 (58.33)     |
| Comorbid presence | 110 (41.67)   |
| **Type of therapy (antihypertension drug)** | |
| Monotherapy     | 242 (91.67)     |
| Combination     | 22 (8.33)       |
| **Medication adherence (pill count method)** | |
| Adherent        | 205 (77.65)     |
| Non-adherent    | 59 (22.35)      |
| **Blood pressure level** |   |
| Controlled      | 108 (40.91)     |
| Uncontrolled    | 156 (59.09)     |

\(a\) The percentage of pill count <80% and 80–100% were considered non-adherent and adherent, respectively.

\(b\) Blood level pressure according to JNC 8 guideline: hypertensive patients without comorbid diabetes mellitus and chronic kidney disease at the age of \(\geq 60\) years should achieve \(<150\) mmHg systolic blood pressure and \(<90\) mmHg blood pressure diastolic. Meanwhile, for patients aged \(<60\) years, the target of achieving systolic blood pressure is \(<140\) mmHg and systolic blood pressure is \(<90\) mmHg. For hypertensive patients with comorbid diabetes mellitus and/or chronic kidney disease at all ages, the target is systolic blood pressure \(<140\) mmHg and diastolic blood pressure \(<90\) mmHg.
has an association with age and the presence of family members, but not with sex, usage duration, and blood pressure.

Then, the comorbidities accompanying hypertension were not associated with poor medication adherence and uncontrolled blood pressure (Tables 4 and 5). Opposed to this finding, several studies revealed that comorbidities such as diabetes mellitus, heart disease, and dyslipidemia affect patient adherence due to the consumption of more complex medications [24–26].

Patient adherence to their antihypertensive drugs indeed affects blood pressure control. However, other predictors of blood pressure control were found in the present study, such as age and education (Table 5). The patient’s age was inversely associated with uncontrolled blood pressure. The youngest group of patients (20–40 years old) were almost six times more likely to have uncontrolled blood pressure compared to those aged >60 (AOR: 5.809; 95% CI: 1.516–22.264; P = 0.010) (Table 5).

It may be assumed that uncontrolled blood pressure in these particular age groups correlates with their adherence level percentage. In the oldest patients group, adherence level was 81.82%, whereas in the age group of 41–60 years old was 75.55% and even lower in 20–40 years old with 73.68%, even though the difference was not significant. Furthermore, Choi et al. [24] stated that patients aged >50 years had high adherence to the consumption of antihypertensive drugs. Older patients usually have a caregiver who helps take their medication and maintain a healthy lifestyle, thus control blood pressure eventually [23]. Moreover, older age was associated with better medication adherence due to perceived vulnerability and disease severity [27, 28].

The level of education in this study is known not to affect medication adherence or blood pressure control based on the chi-square test. However, multivariable logistic regression found that patients educated up to midlevel school had higher blood pressure control than college graduates. Other studies also showed inconsistent results. Ayodapo et al. [29] reported that educational level has no considerable effect on medication adherence, while Adisa et al. [30] showed a statistically significant correlation between medication adherence and education. Generally speaking, patients now have

| Table 2 Sociodemographic and clinical characteristics of patients according to their medication adherence status (n = 264) |
|---------------------------------------------------------------|
| Variable                          | Adherent (n = 205) | Non-adherent (n = 59) | P-value |
| Age (yr)                          |                   |                      |         |
| 20–40                            | 14 (73.68)        | 5 (26.32)            | 0.552   |
| 41–60                            | 102 (75.55)       | 33 (24.45)           |         |
| >60                              | 89 (80.90)        | 21 (19.10)           |         |
| Sex                              |                   |                      | 0.145   |
| Female                           | 155 (79.9)        | 39 (20.1)            |         |
| Male                             | 50 (71.43)        | 20 (28.57)           |         |
| Comorbidity                      |                   |                      | 0.283   |
| No                               | 116 (75.32)       | 38 (24.68)           |         |
| Yes                              | 89 (80.91)        | 21 (19.09)           |         |
| Type of therapy (antihypertension drug) |               |                      | 0.265   |
| Monotherapy                      | 190 (78.51)       | 52 (21.49)           |         |
| Combination                      | 15 (68.18)        | 7 (31.82)            |         |
| Education level                  |                   |                      | 0.425   |
| Elementary school                | 53 (72.6)         | 20 (27.4)            |         |
| Middle school                    | 30 (81.08)        | 7 (18.92)            |         |
| High school                      | 77 (81.05)        | 18 (18.95)           |         |
| College                          | 45 (76.27)        | 14 (23.73)           |         |
| Blood pressure level             |                   |                      | 0.000   |
| Controlled                       | 100 (92.59)       | 8 (7.41)             |         |
| Uncontrolled                     | 105 (67.31)       | 51 (32.69)           |         |

Data are presented as number (%)

| Table 3 Sociodemographic and characteristic of patients according to their blood pressure control (n = 264) |
|---------------------------------------------------------------|
| Variable                          | Controlled (n = 108) | Uncontrolled (n = 156) | P-value |
| Age (yr)                          |                       |                        | 0.005   |
| 20–40                            | 3 (15.79)             | 16 (84.21)             |         |
| 41–60                            | 49 (36.30)            | 86 (63.70)             |         |
| >60                              | 56 (50.91)            | 54 (49.09)             |         |
| Sex                              |                       |                        | 0.302   |
| Female                           | 84 (76.36)            | 110 (23.64)            |         |
| Male                             | 24 (34.28)            | 46 (65.72)             |         |
| Comorbidity                      |                       |                        | 0.612   |
| No                               | 62 (40.26)            | 92 (59.74)             |         |
| Yes                              | 46 (41.82)            | 64 (58.18)             |         |
| Type of therapy (antihypertension drug) |               |                        | 0.365   |
| Monotherapy                      | 101 (41.74)           | 141 (58.26)            |         |
| Combination                      | 7 (30.43)             | 16 (69.57)             |         |
| Education level                  |                       |                        | 0.099   |
| Elementary school                | 26 (36.62)            | 47 (63.38)             |         |
| Middle school                    | 22 (59.46)            | 15 (40.54)             |         |
| High school                      | 37 (38.95)            | 58 (61.05)             |         |
| College                          | 23 (38.99)            | 36 (61.02)             |         |
| Medication adherence (pill count) |                       |                        | 0.000   |
| ≥80%                             | 98 (47.80)            | 107 (52.20)            |         |
| <80%                             | 10 (16.95)            | 49 (83.05)             |         |

Data are presented as number (%)
unlimited access to information regarding hypertension and the importance of medication adherence from websites, mobile applications, and health workers or pharmacists.

The success of hypertension therapy is influenced by adherence and other factors, such as lifestyle, physical activity, diet, sleep patterns, body mass index, smoking status, and stress [31]. Those factors were not observed in the present study due to data availability and time limitations. Another limitation involves the measuring adherence method (self-reported pill count). This method is subjective and may be inaccurate because the patient might remove the drugs from the container in anticipation of adjusting the number of the drug according to the medication schedule [7, 11, 17]. Besides, the number of respondents was limited, and the distribution of Puskesmas did not represent the whole city. So, it is necessary to add the number of respondents from many more Puskesmas for future study.

**Conclusions**

We found a significant association between medication adherence and blood pressure control. In addition, age was found to be an independent predictor that affects blood pressure. Thus, pharmacists and other healthcare providers should pay attention to improving medication adherence and maintaining controlled blood pressure, particularly for patients below 40 years old.

**Abbreviations**

WHO: World Health Organization; SBP: Sistole blood pressure; DBP: Diastole blood pressure; AOR: Adjusted odds ratio; JNC: Joint National Committee; PDC: Proportion of days covered; MEMS: Medication Electronic Monitoring System

**Supplementary information**

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**Additional file 1.**

**Additional file 2.**

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Not applicable.

**Authors’ contributions**

Concept: IE, EIL; Design: IE, EIL, RL; Supervision: IE, RL; Resources: IE, EIL; Materials: IE, EIL; Data collection and processing: IE, EIL, RL; Analysis and interpretation: IE, EIL, RAP; Literature search: IE, RL; Writing: IE, EIL, RL, RAP; Critical reviews: IE, RL, RAP. The author(s) read and approved the final manuscript.

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**Table 4** Multivariable logistic regression results of factors associated with poor medication adherence

| Variable                        | AOR (95% CI)     | P-value |
|---------------------------------|------------------|---------|
| Age (yr)                        |                  |         |
| 20–40                           | 0.963 (0.289–3.211) | 0.951   |
| 41–60                           | 1.309 (0.637–2.609) | 0.444   |
| >60                             | Ref.             | -       |
| Sex                             |                  |         |
| Female                          | 0.596 (0.305–1.177) | 0.316   |
| Male                            | Ref.             | -       |
| Comorbidity                     |                  |         |
| No                              | Ref.             | -       |
| Yes                             | 0.674 (0.351–1.294) | 0.236   |
| Type of therapy (antihypertension drug) |                  |         |
| Monotherapy                     | Ref.             | -       |
| Combination                     | 1.993 (0.695–5.716) | 0.199   |
| Education level                 |                  |         |
| Elementary school               | 1.783 (0.464–4.262) | 0.193   |
| Middle school                   | 1.759 (0.584–5.298) | 0.315   |
| High school                     | 0.894 (0.377–2.121) | 0.799   |
| College                         | Ref.             | -       |
| Blood pressure level            |                  |         |
| Controlled                      | Ref.             | -       |
| Uncontrolled                    | 6.176 (2.785–14.099) | 0.000   |

The reference category is adherent (percentage pill count ≥ 80%)

AOR, adjusted odds ratio; CI, confidence interval

**Table 5** Multivariable logistic regression results of factors associated with uncontrolled blood pressure level

| Variable                        | AOR (95% CI)     | P-value |
|---------------------------------|------------------|---------|
| Age (yr)                        |                  |         |
| 20–40                           | 5.814 (1.519–22.252) | 0.010   |
| 41–60                           | 2.008 (1.146–3.520) | 0.015   |
| >60                             | Ref.             | -       |
| Sex                             |                  |         |
| Female                          | 0.751 (0.397–1.419) | 0.378   |
| Male                            | Ref.             | -       |
| Comorbidity                     |                  |         |
| No                              | Ref.             | -       |
| Yes                             | 0.933 (0.540–1.612) | 0.804   |
| Type of therapy (antihypertension drug) |                  |         |
| Monotherapy                     | Ref.             | -       |
| Combination                     | 1.445 (0.519–4.026) | 0.373   |
| Education level                 |                  |         |
| Elementary school               | 1.147 (0.589–2.539) | 0.669   |
| Middle school                   | 0.387 (0.153–0.974) | 0.046   |
| High school                     | 1.158 (0.563–2.382) | 0.726   |
| College                         | Ref.             | -       |
| Medication adherence (pill count) |                |         |
| ≥80%                            | Ref.             | -       |
| <80%                            | 6.081 (2.672–13.838) | 0.000   |

The reference category is controlled blood pressure

AOR, adjusted odds ratio; CI, confidence interval
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Availability of data and materials
The datasets used and/or analyzed during the current study are available from the corresponding author on reasonable request.

Declarations

Ethics approval and consent to participate
Ethical approval was obtained from the Faculty of Public Health Ethics Committee at Airlangga University, Surabaya, Indonesia (No. 98/BA/KEPK/2020). All subjects were provided written informed consent before participation.

Consent for publication
Not applicable.

Competing interests
There is no conflict of interest in this study and publication.

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