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Article Title (within 20 words without	Effectiveness of Clinical Pharmacist Interventions on
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	Treatment Outcomes for Acute Myocardial
	Infarction Patients: A Before-After Study at a
	Vietnamese hospital
Running Title (within 10 words)	Clinical pharmacist intervention in the treatment of myocardial infarction
Author	Nhu Quynh Tran ^{1,2} , Tan Van Nguyen ^{3,4} , Quynh Thi Huong Bui ^{1,2,*}
ORCID (for more information, please visit https://orcid.org)	¹ Deparment of Clinical Pharmacy, Faculty of Pharmacy, University of Medicine and Pharmacy at Ho Chi Minh City, Ho Chi Minh City, Vietnam ² Department of Pharmacy, Thong Nhat Hospital, Ho Chi Minh City, Vietnam ³ Department of Geriatrics and Gerontology, University of Medicine and Pharmacy at Ho Chi Minh City, Ho Chi Minh City, Vietnam ⁴ Department of Interventional Cardiology, Thong Nhat Hospital, Ho Chi Minh City, Vietnam Tran Quynh Nhu (https://orcid.org/0009-0009-0379-5826) Tan Van Nguyen (https://orcid.org/0000-0002-0234-6596) Quynh Thi Huong Bui (https://orcid.org/0000-0003-3451-4870)
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Authors' contributions	Conceptualization: Tran Quynh Nhu, Tan Van Nguyen,
Please specify the authors' role using this	Quynh Thi Huong Bui
form.	Data curation: Tran Quynh Nhu
Authors can't change and add items, but you	Formal analysis: Tran Quynh Nhu
can delete items that are not applicable.	Methodology: Tran Quynh Nhu, Tan Van Nguyen,
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	Review Board of Thong Nhat Hospital, Ho Chi Minh
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	2022). Patients' personal information was kept
	confidential and used only for research purposes. All
	procedures followed the ethical standards set by
	institutional and national research committees. However,
	this did not interfere with the treatment process.
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5 CORRESPONDING AUTHOR CONTACT INFORMATION

For the corresponding author (responsible for correspondence, proofreading, and reprints)	Fill in information in each box below
First name, middle initial, last name	Quynh, Thi Huong, Bui
Email address – this is where your proofs will be sent	bthquynh@ump.edu.vn
Secondary Email address	
Address	Deparment of Clinical Pharmacy, Faculty of Pharmacy, University of Medicine and Pharmacy at Ho Chi Minh City, Vietnam
Cell phone number	(+84)912261353
Office phone number	
Fax number	

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10 **ABSTRACT**

- 11 Introduction: Acute myocardial infarction (AMI) is a leading cause of global morbidity and
- mortality. Clinical pharmacist intervention offers a promising approach to improve prescription
- appropriateness and treatment outcomes. This study evaluated the impact of this intervention in
- 14 treatment for AMI patients.
- 15 **Methods:** A retrospective before-and-after study was conducted on all AMI patients at the
- 16 Department of Interventional Cardiology, comparing two phases. The *pre* phase was designed
- without clinical pharmacist intervention (August 1, 2019, to December 31, 2019) and the *post*
- phase with the participation of clinical pharmacists in the prescription process (August 1, 2022,
- 19 to December 31, 2022); with 6-month post-AMI follow-up periods in each phase. The impact
- 20 of interventions was evaluated by comparing appropriateness of prescription, treatment
- outcomes, and adverse drug events (ADEs) between the two phases.
- Results: The study included 183 and 211 patients in the *pre* and *post* phases, respectively. The
- overall rates of prescription appropriateness were significantly higher in the *post* phase (85.8%)
- vs. 48.6%, p < 0.001). The mortality rates within 6 months of AMI discharge in the two phases
- were 18.6% and 16.5%, respectively (p = 0.604). The proportions of patients who experienced
- ADEs were 57.4% and 56.4%, respectively (p = 0.845). Clinical pharmacist interventions were
- associated with a higher rate of overall prescription appropriateness (OR: 6.734; 95% CI: 4.098
- -11.065; p < 0.001).
- 29 **Conclusions**: Clinical pharmacist interventions significantly improved the appropriateness of
- prescription for AMI treatment but did not reduce occurrence of mortality or ADE.
- 31 **Keywords:** Acute myocardial infarction; Prescription Appropriateness; Clinical pharmacist
- 32 Intervention

33 1. INTRODUCTION

- Acute myocardial infarction (AMI) is an ischemic syndrome-induced myocardial necrosis [1].
- 35 AMI is the leading cause of morbidity and mortality worldwide, responsible for over 15% of
- annual mortality [2] and creates significant economic burden on the society [3]. Currently, the
- 37 main treatment approach for AMI focuses on optimal medical therapy, percutaneous coronary
- 38 intervention (PCI) and coronary artery bypass grafting to restore perfusion, prevent further
- myocardial necrosis and severe cardiovascular events [1,4,5]. The incidence of AMI and rates
- 40 of associated mortality have decreased in some developed countries [2]. This is probably

41 influenced by the innovation and strengthening of healthcare systems, and advances in treatment 42 management [2]. However, patients with AMI are often treated with polypharmacy [4,5], which 43 increases the risk of adverse drug events (ADEs). ADEs are associated with escalated morbidity 44 and mortality, prolonged hospitalization, and increased healthcare costs [6]. Therefore, during 45 therapy, patients must be carefully monitored for the benefits and risks from drug regimens, to optimize their treatment effectiveness and safety. Adherence to clinical practice guidelines for 46 47 the treatment of AMI and prevention of secondary atherosclerotic cardiovascular events after 48 AMI have shown to reduce the risk of recurrent cardiovascular events, hospital readmission, 49 and incidence of death [1,4,5]. However, data from the United Kingdom Myocardial Ischemia 50 National Audit Project showed that approximately half of patients did not receive the 51 recommended treatment after acute coronary syndrome (ACS) [7]. Clinical pharmacist 52 intervention is a promising approach to promote appropriate prescription practices. These 53 interventions, which include medication adjustments, patient counselling and monitoring, 54 associated with reduced mortality rate, improved adherence, and better clinical outcomes [8-55 11]. Clinical pharmacists play a role in supporting prescription decisions at hospital admission, 56 in-hospital phase and post-discharge follow-up [8,9]. Several studies were conducted to evaluate 57 the effectiveness of clinical pharmacist interventions in patients with AMI [10,11]. 58 Thong Nhat Hospital is a Grade 1 general hospital under the Ministry of Health specialized in 59 geriatrics. Located in Ho Chi Minh City, the largest metropolitan area and a major healthcare 60 hub in southern Vietnam, the hospital holds a pivotal role in providing advanced medical 61 services [12]. The Department of Interventional Cardiology, recognized as one of the leading 62 cardiovascular intervention centres in the region, commits to the management of patients with 63 AMI and other complex cardiovascular diseases [13]. Since January 2020, a clinical pharmacist 64 has been working as a member of a multidisciplinary team in the department to ensure the 65 appropriateness of prescriptions. This could improve the quality of treatment and reinforce the safety of inpatients. This study aimed to assess the effectiveness of clinical pharmacist 66 67 interventions on improving the appropriateness of prescriptions and treatment outcomes in 68 patients with AMI.

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2. MATERIALS AND METHODS

2.1. Study setting

- A retrospective before-and-after study was conducted on inpatients diagnosed with and treated
- for AMI, including ST-segment elevation myocardial infarction (STEMI) and non-ST-segment
- elevation myocardial infarction (NSTEMI), at the Department of Interventional Cardiology,
- 75 Thong Nhat Hospital. The study consisted of two phases. In the pre-intervention phase (or the
- pre phase; from August 1st, 2019, to December 31st, 2019; with a 6-month post-AMI follow-up),
- 77 clinical pharmacists did not intervene in physician's decisions regarding medication
- 78 prescriptions. In the intervention phase (or the *post* phase; from August 1st, 2022, to December
- 79 31st, 2022, with a 6-month post-AMI follow-up), clinical pharmacists actively participated in
- 80 the decision-making process for prescriptions.

81 **2.2. Study population and sampling**

- 82 The study sampled all patients aged 18 or older who were diagnosed and treated for AMI
- 83 (including STEMI and NSTEMI with ICD codes I21 or I22) at the Department of Interventional
- 84 Cardiology, Thong Nhat Hospital, Ho Chi Minh City, Vietnam, during the two study phases.
- 85 The exclusion criteria were patients who were transferred to another hospital, refused treatment,
- or lost contact within 6 months after AMI. Furthermore, if patients were hospitalized more than
- once for AMI at the Department of Interventional Cardiology during the same study phase, the
- study only collected data from the first hospitalization.

2.3. Study outcomes

- 90 The data collection considered information on patient characteristics, number of drugs per day,
- 91 appropriateness of prescription (during hospitalization and discharge), treatment outcomes, and
- 92 ADEs. Patient characteristics included age, sex, AMI classification, revascularization strategy
- 93 comorbidity, number of comorbidities, Charlson Comorbidity Index (CCI), estimated
- 94 glomerular filtration rate (eGFR), number of drugs per day and the length of hospital stay. The
- 95 appropriateness of prescription included the number of drug-related problems (DRPs),
- 96 appropriateness of indications, dosage, route of administration, and the variables were derived
- 97 for overall appropriateness. The treatment outcome was the mortality rate within 6 months after
- AMI. ADEs included those that appeared during hospitalization, but not at the time of admission.
- 99 The impact of the clinical pharmacist interventions was evaluated by comparing indicators pair-
- wised from the *pre* and *post* phases. The evaluation focused on two key aspects: the primary
- outcome, which was the appropriateness of prescription; and the secondary outcomes, which
- included treatment outcomes and the occurrence of ADEs. This approach allowed for a

103 comprehensive analysis of the pharmacist's contribution to improving medication use and

patient care.

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Definition

106 Clinical pharmacist interventions

107 The clinical pharmacist intervention was defined as the intervention made by a graduated 108 clinical pharmacist with at least three years of professional experience; who was assigned to 109 work in the Department of Interventional Cardiology for at least 4 hours/day, 5 days/week and 110 participated in the care of all patient cases, including patients with AMI, alongside doctors, 111 nurses, and other healthcare professionals; specifically, those who participated in staff meetings, 112 examined patients with a physician, reviewed medical records, assessed the appropriateness of 113 prescription to treat AMI, monitored for potential side effects, and provided recommendations 114 to optimize medication therapy. The clinical pharmacist may also share further rationale 115 information to both patients and healthcare staff regarding medication management after AMI. 116 The clinical pharmacist reviewed the medical records from the previous weekend on the first 117 day of the following week and continued to intervene if any issues were identified. In case of 118 emergency, doctors could directly contact the on-duty pharmacists in the pharmacy department. 119 The interventions by the clinical pharmacist must be recorded in an electronic archive housed 120 on the hospital's secure internal server. To ensure confidentiality and data security, the folder 121 was password-protected, and access was restricted to assigned clinical pharmacists only.

122 The appropriateness of prescription

123 The appropriateness of AMI prescriptions was evaluated during hospitalization based on specific criteria. These included the appropriateness of indications, dosage, route of 124 125 administration, and overall appropriateness of prescription for AMI management (such as 126 nitrates, opioid analgesics, anticoagulants, antiplatelets, beta-blockers, calcium channel 127 blockers - CCBs, angiotensin-converting enzyme inhibitors - ACEis/angiotensin receptor 128 blockers – ARBs, aldosterone antagonists, statins, and proton pump inhibitors – PPIs). These 129 evaluations were conducted in accordance to the "Vietnam Minister of Health Guidelines for 130 Diagnosis and Treatment of Acute Coronary Syndrome 2019 [1]" and "Vietnamese National 131 Drug Formulary 2018 [14]".

Indications were considered appropriate if the prescribed medications aligned with guideline recommendations and had no contraindications. The appropriateness of dosage was assessed to

- ensure that all key drugs for the treatment of AMI were prescribed at reasonable doses that
- aligned with clinical guidelines. Similarly, the route of administration was evaluated to confirm
- that the prescribed method of drug delivery was suitable and consistent with the recommended
- practice for AMI management.
- 138 The overall appropriateness of prescription was defined as the fulfilment of all three criteria –
- appropriateness of indications, dosage, and route of administration. Notably, the dosage and
- route of administration were assessed only if the indication was deemed appropriate.
- 141 A DRP was identified if patients were prescribed at least one drug to treat AMI with an
- inappropriate indication, dosage, or route of administration. Indications, contraindications, and
- dosages applied to assess the rational use of certain drugs in the treatment of AMI are presented
- in Appendix 1.
- 145 Treatment outcomes
- The study recorded the mortality rate within six months following AMI, which included two
- 147 components. In-hospital mortality was defined as death that occurred during the patient's
- hospital stay, prior to discharge. Post-discharge mortality was recorded in patients who survived
- to discharge, up to six-month follow-up after the onset of AMI to capture any subsequent deaths.
- Data on mortality was obtained from the hospital information system and the electronic portal
- of the Vietnam Social Insurance, ensuring comprehensive and accurate documentation.
- 152 Adverse drug events
- 153 The study considered ADEs that occurred during hospitalization, but were not presented at the
- time of admission. ADEs were recorded by pharmacists, physicians or nurses, either from the
- medical records or by direct observation. In both phases, all ADEs were registered in the
- hospital's ADE surveillance network. This network collects information and reports to The
- National Centre of Drug Information and Adverse Drug Reactions Monitoring in Vietnam.
- 158 The ADE aetiology was assessed using the Naranjo algorithm. Both "certain" and "likely" (≥ 5
- points) ADEs were recorded. ADEs with Naranjo scores of under 5 were excluded.
- 160 The criteria for ADE diagnosis in this study are presented in Appendix 2.

161 **2.4. Statistical method**

- All data were analysed using the Statistical Package for the Social Sciences (SPSS) software,
- version 20.0, with the significance threshold at p < 0.05. Descriptive statistics were used to
- summarize the data. Variables with normal distribution were presented as means \pm standard

- deviation (SD), and those with non-normal distribution were summarized in medians with
- interquartile ranges (IQR). Categorical variables (such as AMI classification, sex, comorbidities,
- appropriateness of prescription, type of DRPs, treatment outcomes, and ADEs) were presented
- as frequency and percentage.
- 169 Inferential analyses were performed to compare variables between the two groups. Categorical
- variables were analysed using the Chi-square test. When more than 20% of expected cell counts
- were below 5, Fisher's exact test was applied. Continuous variables were compared using
- 172 Student's t-test for normally distributed data or the Mann-Whitney U test for data with non-
- 173 normal distribution.
- Multivariable logistic regression analysis, employing the backward elimination method, was
- 175 conducted to identify factors associated with the appropriateness of prescription for treating
- 176 AMI at the hospital. The dependent variable was overall prescription appropriateness, while
- independent variables included age, dyslipidaemia, history of coronary heart disease, number of
- drugs per day, and whether clinical pharmacist intervention was provided (yes/no). Independent
- variables were examined for potential univariate associations with statistical significance
- threshold of 80%, and multicollinearity prior to regression model entry. Additionally, the
- independent variables were examined for multicollinearity. Variance inflation factors (VIFs)
- and Hosmer-Lemeshow test were used for model selection.

183 **2.5. Ethical considerations**

- The study protocol followed the ethical standards set by institutional and national research
- committees, and was approved by the Institutional Review Board of Thong Nhat Hospital, Ho
- 186 Chi Minh City, Vietnam (60/2022/BVTN-HDYD September 20th, 2022). Patient's personal
- information was kept confidential and used only for research purposes.

188 **3. RESULTS**

189 3.1. Characteristics of the study population

- The study included 394 patients, of which 183 patients were in the *pre* phase and 211 in the *post*
- phase. A total of 180 (45.7%) patients were diagnosed with STEMI, whereas 214 (54.3%) were
- diagnosed with NSTEMI. There was no difference in the AMI classification between the two
- phases (p = 0.081). The majority of the study population were elderly patients (\geq 60 years old),
- with a median age of 66 (57-79) years for all study populations. The age group of 75 years and
- above accounted for 32.5%, with a statistically significant difference between the two phases,

- primarily observed in this age group. The proportion of males was higher than females (66.5%
- vs. 33.5%). The rate of patients undergoing PCI in the study was recorded as 54.1% in phase 1
- and 61.1% in phase 2 (p = 0.158). The most common comorbidities in both study phases were
- 199 hypertension, heart failure, and dyslipidaemia. The median length of hospital stay in the two
- 200 phases was 9 (IQR: 6 14) days and 8 (IQR: 6 12) days, respectively (p = 0.315).
- 201 Characteristics of the study population are summarized in Table 1.
- 202 [Insert Table 1 here]
- 203 **3.2.** The effectiveness of clinical pharmacist interventions
- 3.2.1. The appropriateness of AMI prescription during hospitalization
- The overall prescription appropriateness for AMI treatment was higher in the intervention phase
- 206 (85.8%) than in the pre-intervention phase (48.6%).
- The number of DRPs in the *post* phase decreased significantly compared with that in the *pre*
- 208 phase. One occurrence of DRP accounted for the highest proportion of cases in both phases, and
- 209 there were no cases with more than two DRPs in the *post* phase. In the *pre* phase, the rate of
- inappropriate indications and dosages of PPIs were the highest and decreased significantly in
- 211 the *post* phase. Details regarding the appropriateness of prescribing medications in both study
- 212 phases are presented in Table 2. The sub-analysis, excluding DRPs related to PPI prescribing,
- 213 indicated that the rates of overall prescription appropriateness of primary medications for AMI
- 214 treatment (excluding PPIs) in *pre* and *post* phases were 86.9% and 91.5%, respectively (p =
- 215 0.141, Appendix 3).
- 216 [Insert Table 2 here]
- 217 Appendix 4 presents the selection process for the multivariate model based on univariate
- 218 regression analysis, and demonstrates the goodness of fit for the chosen model through Variance
- 219 Inflation Factor (VIF) assessment and the Hosmer-Lemeshow test. The results of the
- 220 multivariable logistic regression analysis indicated that clinical pharmacist interventions were
- associated with a higher rate of overall prescription appropriateness for AMI treatment in the
- 222 hospital (Table 3). In contrast, a history of coronary artery disease and an increased number of
- drugs used per day were associated with a lower rate of appropriate prescriptions (p < 0.05).
- 224 [Insert Table 3 here]
- 225 3.2.3. Treatment outcomes

- The 6-month post-AMI mortality rate in the *pre* phase was 18.6%, which was higher than that
- in the *post* phase (16.6%). However, the difference was not statistically significant (Table 4).

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- 229 [Insert Table 4 here]
- 230 3.2.2. Adverse drug events
- A total of 105 patients (57.4%) in the *pre* phase and 119 patients (56.4%) in the *post* phase
- experienced at least one ADE during their hospital stay (p = 0.845). The most common ADEs
- 233 recorded in this study were electrolyte disorders (including hyperkalaemia, hypokalaemia,
- 234 hypernatremia, and hyponatremia), bleeding, and acute kidney injury, with the rate of each ADE
- exceeding 15% (Table 5).
- 236 [Insert Table 5 here]

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4. DISCUSSIONS

- In this study, the prevalence of STEMI and NSTEMI were 45.7% and 54.3%, respectively.
- These results are similar to those from the study by Kim et al. in 2021 [15], which reported 43.4%
- of patients with STEMI and 56.6% of patients with NSTEMI. The majority of the study
- 242 population was elderly patients (67.3%), age is one of the main factors contributing to the
- increased risk of death [9,16] and bleeding in patients with AMI [16,17]. In both phases of our
- study, a higher proportion of males (> 65%) than females (< 35%) were recorded. Several
- studies have explored the differences in cardiovascular outcomes between genders. Findings in
- Australia by Nedkoff et al. in 2023 [18] indicated that the risk of major adverse cardiovascular
- events, cardiovascular-related deaths, and all-cause mortality were generally higher in females.
- Score of one in CCI was most common in the study, followed by 2, 0, 3, and \geq 4 points.
- 249 Hypertension (94.2%), dyslipidaemia (75.4%), and heart failure (59.9%) were common
- comorbidities in both study phases. Comorbidities and polypharmacy are associated with poor
- 251 patient adherence to medications [19].
- 252 Compared to the *pre* phase, the *post* phase showed a statistically significant increase in the
- overall appropriateness of prescribing AMI medications during hospitalization (48.6% vs.
- 85.5%, p < 0.001). In particular, the appropriateness of the indications and dosages significantly
- 255 increased during the *post* phase. Most DRPs in the *pre* phase were inappropriate indications and
- 256 PPI doses, which decreased significantly during the *post* phase. In particular, rabeprazole was

repeatedly prescribed despite not being recommended for the prevention of NSAID-induced ulcers, and pantoprazole 40 mg once daily was also frequently used for ulcer prophylaxis, even though this dosage was considered inappropriate [14]. This showed that the DRPs in the treatment of AMI in the Department of Interventional Cardiology had been effectively intervened by clinical pharmacists. The sub-analysis results indicated that the prescribing of primary medications for AMI treatment (excluding PPIs) yielded a high level of appropriateness, with an overall rate of 86.9% in the *pre* phase and no significant difference in the intervention phase. As the Department of Interventional Cardiology at Thong Nhat Hospital is a major intervention centre in Vietnam, physicians demonstrated a high level of competency in providing indications for AMI treatment. In comparison, a study by Gona et al. in 2021 [20] showed that 52% of patients with ACS had DRPs, with the majority of DRPs related to drug selection (34.05%) and dose selection (26.97%). This is the higher DRP rate than observed in both phases of our study. In general, differences in study populations, areas, and criteria of appropriateness led to difficulties in comparing studies. The effect of the intervention was also proven through the logistic regression analysis results, with intervention from clinical pharmacists as a factor related to increasing the appropriateness of prescribing medications to treat AMI at the hospital (OR: 6.734; 95% CI: 4.098 - 11.065; p < 0.001). The importance of clinical pharmacists in the detection, resolution, and prevention of DRPs, and their contribution in improving patient outcomes and optimizing healthcare costs have been demonstrated in some other studies on cardiovascular patients [20,21]. Conversely, the results from the logistic regression equation showed that a history of coronary heart disease and an increased number of medications used per day were associated with a reduced rate of appropriate medication use in the treatment of AMI. This may be explained by clinical complexity, as patients with coronary heart disease often have multiple comorbidities and are frequently on several medications which can complicate treatment decisions [22]. In summary, the available data and main findings of this study suggested that clinical pharmacist interventions can have a positive impact on improving the rates of prescription appropriateness in treatment for patients with AMI. Previous studies have reported varying rates of major bleeding in patients with AMI, ranging from 0.39% to 10.8% [17,23,24]. There are several reasons for the variation in bleeding rates, including differences in bleeding definition, patient characteristics, and treatment therapy. Bleeding is a considerable adverse event associated with short-term, long-term, and thrombotic

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events, such as AMI and stroke. Additionally, interruption of antithrombotic therapy due to bleeding is independently associated with increased mortality [25]. Thirteen ADEs other than bleeding were recorded, with rates ranging from 0.5% to 30.2%. In a study by Mahadevappa et al. in 2022 [26], 20.7% of the patients experienced 30 different ADRs. The six-month post-AMI mortality rates in Soldati et al. in 2021 [19], Takeji et al. in 2021 [27], and Kumar et al. in 2021 [28] ranged from approximately 8% to 11%, which were lower than in our study. A cohort study assessing long-term survival following AMI in Australia and New Zealand found that the probability of survival decreased rapidly within the first year after AMI and declined gradually thereafter [29]. For many years, post-AMI mortality rates in different settings are varied because of differences in designated institutions and study populations. In general, our study did not observe any difference in the incidence of ADEs and the mortality rate after 6 months of AMI treatment between the two phases. This may be due to the overall high level of appropriateness in medication use across both phases, except for PPIs, which likely limited the occurrence of ADEs and the mortality rate. Although this study demonstrated the effectiveness of clinical pharmacist interventions in treating patients with myocardial infarction, we acknowledge several limitations, including the following. The study was conducted at a single centre, limiting its ability to reflect the diversity in culture, race, gender, and age of AMI patients. The before-and-after design with a lengthy gap between data collection points raised the possibility that prescribing improvements over time may have occurred independently of pharmacist intervention. The evaluation of "prescription appropriateness," based on indications, dosages, and administration routes, may not fully reflect the complexity of clinical pharmacy practice. DRPs in the *pre* phase may have been underreported due to the absence of a clinical pharmacist. Finally, the use of two distinct populations in the pre and post phases could introduce bias due to demographic and baseline differences. However, multivariable logistic regression was used to adjust for these factors, minimizing their impact and ensuring a robust evaluation of the interventions.

5. CONCLUSIONS

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- Our study demonstrated that clinical pharmacist interventions significantly improved the appropriateness of prescribing medications to treat AMI but did not reduce mortality or ADE. Specifically, clinical pharmacy has played a crucial role in improving the appropriateness of
- 318 indications and dosages for PPI prescriptions. These findings support enhancing the role of

- 319 clinical pharmacists in ensuring the appropriateness of AMI prescription to minimize the rate of
- 320 DRPs and optimize treatment outcomes.
- 321 **6. COMPETING INTERESTS**
- No potential conflict of interest relevant to this article was reported.
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Table 1. Patient's characteristics

	All	Pre-	Intervention	
Characteristics	(N=394)	intervention	$(n^2 = 211)$	p-value
A () 1' (IOD)	(((57, 70)	$\frac{(n^1 = 183)}{67.(5692)}$	(5 (57, 74)	0.025#
Age (years), median (IQR)	66 (57 – 79)	67 (56 – 83)	65 (57 – 74)	0.025#
Age group, n (%)				
< 50	47 (11.9)	23 (12.6)	24 (11.4)	_
50 - <75	219 (55.6)	84 (45.9)	135 (64.0)	0.001*
≥75	128 (32.5)	76 (41.5)	52 (24.6)	_
Sex, n (%)				
Male	262 (66.5)	120 (65.6)	142 (67.3)	_ 0.718*
Female	132 (33.5)	63 (34.4)	69 (32.7)	- 0.716
AMI classification, n (%)		40		
STEMI	180 (45.7)	75 (41.0)	105 (49.8)	_ 0.081*
NSTEMI	214 (54.3)	108 (59.0)	106 (50.2)	- 0.081
Revascularization strategy, n (%)	. 01			
PCI	228 (57.9)	99 (54.1)	129 (61.1)	0.158*
CABG	0 (0)	0 (0)	0 (0)	-
Comorbidities, n (%)				
Hypertension	371 (94.2)	172 (94.0)	199 (94.3)	0.891*
Dyslipidemia	297 (75.4)	114 (62.3)	183 (86.7)	< 0.001*
Heart failure	236 (59.9)	103 (56.3)	133 (63.0)	0.173*
Diabetes	146 (37.1)	57 (31.1)	89 (42.2)	0.024*
Coronary heart disease	96 (24.4)	46 (25.1)	50 (23.7)	0.740*
Chronic kidney disease	54 (13.7)	19 (10.4)	35 (16.6)	0.074*
History of stroke	29 (7.4)	15 (8.2)	14 (6.6)	0.554*
Atrial fibrillation	19 (4.8)	11 (6.0)	8 (3.8)	0.305*
Peripheral artery disease	2 (0.5)	2 (1.1)	0 (0)	0.215**
Number of comorbidities, median (IQR)	5 (3 – 6)	4 (3 – 6)	5 (4 – 6)	< 0.001*

83 (21.1)	35 (19.1)	48 (22.7)	
148 (37.6)	69 (37.7)	79 (37.4)	_
91 (23.1)	45 (24.6)	46 (21.8)	0.916*
38 (9.6)	18 (9.8)	20 (9.5)	_
34 (8.6)	16 (8.7)	18 (8.5)	_
65.3 ± 27.0	63.2 ± 27.9	67.0 ± 26.1	0.165##
11 (9 – 14)	11 (9 – 14)	11 (9 – 13)	0.943#
8 (6 – 13)	9 (6 – 14)	8 (6 – 12)	0.315#
	148 (37.6) 91 (23.1) 38 (9.6) 34 (8.6) 65.3 ± 27.0 11 (9 – 14)	$148 (37.6)$ $69 (37.7)$ $91 (23.1)$ $45 (24.6)$ $38 (9.6)$ $18 (9.8)$ $34 (8.6)$ $16 (8.7)$ 65.3 ± 27.0 63.2 ± 27.9 $11 (9-14)$ $11 (9-14)$	$148 (37.6)$ $69 (37.7)$ $79 (37.4)$ $91 (23.1)$ $45 (24.6)$ $46 (21.8)$ $38 (9.6)$ $18 (9.8)$ $20 (9.5)$ $34 (8.6)$ $16 (8.7)$ $18 (8.5)$ 65.3 ± 27.0 63.2 ± 27.9 67.0 ± 26.1 $11 (9 - 14)$ $11 (9 - 14)$ $11 (9 - 13)$

^{#:} Mann-Whitney U test; ##: Student's T test; *: Chi-square test; **: Fisher's exact test

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Table 2. The appropriateness of prescribing drugs to treat acute myocardial infarction

Annuantiatanaga	All	Pre-intervention	Intervention	n volue
Appropriateness	(N = 394)	$(n^1 = 183)$	$(n^2 = 211)$	p-value
The appropriateness of indications, n (%)	346 (87.8)	145 (79.2)	201 (95.3)	< 0.001*
The appropriateness of the dose, n (%)	314 (79.7)	121 (66.1)	193 (91.5)	< 0.001*
The appropriateness of the route of administration, n (%)	380 (96.4)	173 (94.5)	207 (98.1)	0.056*
Overall appropriateness, n (%)	270 (68.5)	89 (48.6)	181 (85.8)	< 0.001*
The number of DRPs median (IQR)	0 (0 – 1)	1 (0 – 1)	0 (0 – 0)	< 0.001#
- 0 DRP, n (%)	270 (68.5)	89 (48.6)	181 (85.8)	
- 1 DRP, n (%)	99 (25.1)	71 (38.8)	28 (13.3)	< 0.001*
- ≥ 2 DRPs, n (%)	25 (6.3)	23 (12.6)	2 (0.9)	
Inappropriate indication, n (%)				
- Contraindications of ACEi/ARB	4 (1.0)	4 (2.2)	0 (0)	0.045**
- Contraindications of spironolacton	10 (2.5)	3 (1.6)	7 (3.3)	0.350**
- Contraindications of statin	7 (1.8)	5 (2.7)	2 (0.9)	0.258**

Annuantiatanaga	All	Pre-intervention	Intervention	n volue	
Appropriateness	(N = 394)	$(n^1 = 183)$	$(n^2 = 211)$	p-value	
- Incorrect/or lack of PPI indication	31 (7.9)	30 (16.4)	1 (0.5)	< 0.001**	
Inappropriate dose, n (%)					
- Inappropriate dose of anticoagulant	16 (4.1)	7 (3.8)	9 (4.3)	0.825*	
- Inappropriate loading dose of	4 (1.0)	4 (2.2)	0 (0)	0.046**	
clopidogrel	4 (1.0)	4 (2.2)	0 (0)	0.040	
- Inappropriate loading dose of statin	4 (1.0)	4 (2.2)	0 (0)	0.046**	
- Inappropriate dose of beta-blocker	1 (0.3)	1 (0.5)	0(0)	0.464**	
- Inappropriate dose of ACEi/ARB	3 (0.8)	1 (0.6)	2 (0.9)	1.000**	
- Inappropriate dose of PPI	62 (15.7)	56 (30.6)	6 (2.8)	< 0.001*	
Inappropriate route of administration, r	n (%)	~O)			
- Inappropriate route of administration	14 (3.6)	10 (5.5)	4 (1.9)	0.056**	
of PPI	17 (3.0)	10 (3.3)	+ (1.7)	0.030	

Abbreviation: ACEi: Angiotensin-converting enzyme inhibitors; ARB: Angiotensin II receptor blockers; PPI: proton pump inhibitors

Table 3. Factors related to the appropriateness of prescribing during hospitalization by multivariable logistic regression analysis

Factors	p-value*	OR	95% CI
Interventions (yes)	< 0.001	6.734	4.098-11.065
History of coronary heart disease	0.005	0.466	0.273-0.797
Number of drugs per day	0.043	0.932	0.870-0.998
* Multivariable logistic regression			

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Table 4. The treatment outcomes of the study population

_	All	Pre-intervention	Intervention	n valua	
	Outcome	(N=394)	$(n^1 = 183)$	$(n^2 = 211)$	p-value
Survival		325 (82.5)	149 (81.4)	176 (83.4)	0.604*

^{*:} Mann-Whitney U test; *: Chi-square test; **: Fisher's exact test

Mortality at 6 months after AMI	69 (17.5)	34 (18.6)	35 (16.6)	0.604*
- In-hospital mortality	4 (1.0)	3 (1.6)	1 (0.5)	0.341**
- Post-discharge mortality $(n^1 = 180, n^2 = 210)$	65 (16.7)	31 (17.2)	34 (16.2)	0.785*
*: Chi-square test; **: Fisher's exact	t test			

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Table 5. Adverse drug events in the study

ADEa (0/)	All	Pre-intervention	Intervention	1
ADEs, n (%)	(N=394)	$(n^1 = 183)$	$(n^2 = 211)$	p-value
At least one ADE	224 (56.9)	105 (57.4)	119 (56.4)	0.845*
Electrolyte disorder	119 (30.2)	68 (37.2)	51 (24.2)	0.005*
Bleeding	105 (26.6)	51 (27.9)	54 (25.6)	0.610*
Acute kidney injury	75 (19.0)	29 (15.8)	46 (21.8)	0.133*
ADEs on the digestive system	37 (9.4)	16 (8.7)	21 (10.0)	0.731*
Acute liver injury	27 (6.9)	13 (7.1)	14 (6.6)	0.854*
Headache	14 (3.6)	8 (4.4)	6 (2.8)	0.414*
Hypotension	11 (2.8)	8 (4.4)	3 (1.4)	0.076**
Mental disorder	9 (2.3)	5 (2.7)	4 (1.9)	0.739**
Cough	9 (2.3)	6 (3.3)	3 (1.4)	0.314**
Anaphylaxis	7 (1.8)	5 (2.7)	2 (0.9)	0.258**
Hypoglycaemia	6 (1.5)	2 (1.1)	4 (1.9)	0.690**
Thrombocytopenia	4 (1.0)	2 (1.1)	2 (0.9)	1.000**
Hypertension	3 (0.8)	2 (1.1)	1 (0.5)	0.599**
Dyspnoea	2 (0.5)	0 (0)	2 (0.9)	0.501**

^{*:} Chi-square test; **: Fisher's exact test

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