# Prevalence of depression among heart failure inpatients and its associated socio-demographic factors: implications for personal-and family-based treatment management in health facilities in Vietnam 

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#### Abstract

OBJECTIVE: Our objective is to identify the prevalence of depression among inpatients with heart failure (HF), and to ascertain the factors associated with the depression from a wide spectrum of sociodemographic variables.

MATERIALS AND METHODS: We conducted a hospital-based cross-sectional survey of prospectively collected data in inpatients with a diagnosis of HF at Vietnam National Heart Institute, Bach Mai Hospital (Hanoi, Vietnam) from July 2020 to July 2021. A sample size of 128 inpatients with HF was finally included. Primary outcome variable was depression ICD-10.

RESULTS: The mean age was 62.34 (SD = 14.76). The sex ratio was 66 males to 62 females. The overall prevalence of depression ICD-10 was $46.88 \%$ among HF inpatients. The proportion of the depressed patients fluctuated between $37.21 \%$ and $\mathbf{8 3 . 3 3 \%}$ by NYHA heart failure classification. Compared to the depressive prevalence among patients with NYHA class II, the odds were 8.43 times higher for those with NYHA class IV (OR univariate 8.43; 95\% Cl 1.6343.46). Patient's age was significantly associated with increased prevalence of depression (OR multivariate 1.07 ; $95 \% \mathrm{Cl} 1.003-1.14$ ). It was significantly higher odds of depression in HF patients who felt sadness after a diagnosis of an illness (OR multivariate 18.02, 95\% CI: 4.2177.08). Individuals with higher household economic status were less likely to be diagnosed with depression compared to those with lower household economic status (OR multivariate


$0.15,95 \% \mathrm{Cl}: 0.02-0.92)$. The odds of depression were significantly higher in HF patients who reported with family conflict (OR multivariate 23.45, 95\% CI: 1.29-423.55), and in those having the loss of a close family member (OR multivariate 38.62, $95 \%$ CI: 1.41-1055.98).

CONCLUSIONS: The prevalence of depression by ICD-10 was relatively high. Age of patient, sadness after a diagnosis of an illness, household economic status, family conflict and loss of a close family member were significantly associated with the depression among HF inpatients. Present results suggest a need for a disease management program addressing both psychological and HF aspects with the aim of improving health outcomes for the inpatients in Vietnam health facilities.

## Key Words:

Depression, Heart failure, Stress factor, Sociodemographic

## Introduction

Heart failure (HF), the final stage of various heart disease, known to be a syndrome characterized by left ventricular dysfunction, exercise intolerance, reduced Quality of Life (QOL), and markedly shortened life expectancy ${ }^{1,2}$. Depression is not only the prevalent psychological complication mainly contributing to the rehospitaliza-
tion and death in HF patients as well as impaired QOL ${ }^{3-6}$. Hence, clinicians and researchers still consider the depression as an important outcome to evaluate HF therapy. A variety of current evidence ${ }^{3,7}$ showed the association between depression and HF. The most recent systematic review ${ }^{3}$ indicated the overall prevalence of depression among HF patients has been increasing globally in recent years, with the 1.5 times higher prevalence in women compared to men. The prevalence of this psychological problem was higher in countries with low economic income ( $56.7 \%$ ) than high-income countries (39.2\%) ${ }^{3}$.

Despite the negative effects of comorbid depressive symptoms in HF patients, depression was frequently unrecognized and undetected in people with HF because many of the physical symptoms are similar to those of HF. Sometimes a patient's sadness is perceived by doctors, caregivers and even the patient as a normal response of someone who is suffering from a chronic physical illness. Therefore, most of the symptoms of depression in them have not been diagnosed, especially clinical practice condition in Vietnam hospitals. The number of studies on depressive symptoms in HF patients was growing in some countries in Asia, but relatively little is known about depression in Vietnamese patients with HF and no adequate publication was reported from Vietnam. To the best of our knowledge, there are no intervention programs for this psychiatric problem in HF inpatients in lower middle income countries, as well as in resource-scare hospital settings. The purpose of this study was, therefore, to identify the prevalence of depression among inpatients with HF, and to ascertain in this population associated factors for the depression from a wide spectrum of sociodemographic variables, thereby, elaborating implications for personal and fami-ly-based management in health facilities.

## Patients and Methods

## Study Design and Setting

From July 2020 to July 2021, a cross-sectional survey was conducted on inpatients with a diagnosis of HF at Vietnam National Heart Institute, Bach Mai Hospital (Hanoi, Vietnam). The patients with HF of both sexes who met the following inclusion criteria were included:

- Having the ability to understand, read, speak, and write in Vietnamese.
- Patients of both sexes at the age of 18 years or above.
- Received inpatient treatment at Vietnam National Heart Institute, Bach Mai Hospital.
- Previously diagnosed with HF.

This single-centre study was approved by Institutional Review Board of Hanoi Medical University. All study patients with HF were voluntary, and they could leave the study at any time. Informed consent was obtained from all subjects in a language that they understood (Vietnamese language) and their signature.

## Sample Size and Sampling

The sample size of the study was calculated according to the following formula.

$$
\mathrm{n}=Z_{\left(1-\frac{\alpha}{2}\right)}^{2} \frac{\mathrm{p}(1-\mathrm{p})}{\mathrm{d}^{2}}
$$

n : number of participants, $\mathrm{Z}(1-\mathrm{a} / 2)=1.96$ (confidence interval: $95 \%$ ), d: $0.09, \mathrm{p}=48.5 \%$ which was recorded prevalence of depression among HF patients ${ }^{8}$. The research sample size was calculated as 118 . Finally, a total of 128 patients with HF were recruited. A convenient sampling was applied for the cohort of HF inpatients.

## Outcome Variable

Our outcome variable was depression ICD10 in a population of HF patients. Among 128 patients with HF, those with depression had a definitive diagnosis according to ICD-10 depression diagnostic criteria ${ }^{9}$ by three general practitioners known as the psychiatrists (Dr. N.N.T., Dr. V.-S.B. and Dr. V.-H.N.) with 10 years of experience. 128 patients were contacted by phone to schedule a consultation and a face-toface interview. The data of sociodemographic factors and stress factors were collected during the same interview by a group of psychiatrists and public health experts.

## Data Analysis

All analyses were performed with a software program (Stata ${ }^{\circledR} 15$; StataCorp LLC, College Station, TX, USA). First, descriptive statistics was applied for the sociodemographic profile of the sample. Second, depressive prevalence of the study sample was calculated with the frequency and percentage according to New York Heart

Association (NYHA) heart failure classification. Then, difference in the proportion of depression by NYHA classification was examined by $\chi^{2}$ test and their association was identified with an univariate logistic regression analysis. Third, difference in selected study characteristics between non-depressed group and depressed group were examined by $\chi^{2}$ test and Fisher's exact test. Finally, a multivariate model with logistic regression was utilized for analyzing the factors associated with ICD-10 depression. We set the level of statistical significance at 0.05 .

## Results

As shown in Table I, more than half of respondents aged 60 years old or over ( $59.38 \%$ ). The mean age was 62.34 ( $\mathrm{SD}=14.76$ ). The sex ratio for the entire sample size was 66 males to 62 females. Only $18.75 \%$ patients graduated from college/tertiary and higher. $75.78 \%$ of the patients married, and $20.31 \%$ of them widowed. The majority of patients was Kinh ethnicity (89.84\%). By household's wealth status, the proportion of the poor, the average and the wealthy were $17.19 \%$,

Table I. Sociodemographic profile of study patients with heart failure ( $\mathrm{N}=128$ ).

|  | Patients | \% |
| :---: | :---: | :---: |
| Age $\geq 60$ |  |  |
| No | 52 | 40.62 |
| Yes | 76 | 59.38 |
| Age (years) - Mean; SD (IQR) | 62.34; 14.76 (24-92) |  |
| Gender |  |  |
| Male | 66 | 51.56 |
| Female | 62 | 48.44 |
| Education |  |  |
| Illiteracy | 7 | 5.47 |
| Primary/Junior high school | 82 | 64.06 |
| High school | 15 | 11.72 |
| College | 19 | 14.84 |
| Tertiary and above | 5 | 3.91 |
| Ethnicity |  |  |
| Kinh | 115 | 89.84 |
| Minority | 13 | 10.16 |
| Occupation |  |  |
| Blue-collar worker | 78 | 60.94 |
| White-collar worker | 9 | 7.03 |
| Freelancer | 10 | 7.81 |
| Retirer | 31 | 24.22 |
| Marital status |  |  |
| Married | 97 | 75.78 |
| Single | 4 | 3.12 |
| Divorced | 1 | 0.78 |
| Widowed | 26 | 20.31 |
| Living area |  |  |
| Rural | 78 | 60.94 |
| Urban | 36 | 28.12 |
| Mountainous | 14 | 10.94 |
| Household economic status |  |  |
| Lower | 22 | 17.19 |
| Higher | 106 | 82.81 |
| Support from family caregivers in providing care |  |  |
| No | 9 | 7.03 |
| Yes | 119 | 92.97 |
| Impact of heart failure on occupation |  |  |
| Still continuing the current occupation | 26 | 20.31 |
| Do not continue the current occupation | 31 | 24.22 |
| Can do it but less | 71 | 55.47 |

As was shown in Figure 1, the most common traumatic stress factors were the sadness after a diagnosis of HF, which accounted for more than half of study sample ( $53.12 \%$ ). The second most common stress factor then was documented with economic difficulties (20.31\%).


Figure 1. Stress factors among study patients with HF ( $\mathrm{N}=$ 128). Calculated as the percentage; red $=$ YES; gray $=$ NO.
$75.78 \%$, and $7.03 \%$ respectively. Most of them received the support from family caregivers in providing care ( $92.97 \%$ ). Regarding the impact of HF on the occupation, more than half of the patient sample can do current occupation but less ( $55.47 \%$ ), $24.22 \%$ of them, reported to be unable to continue the current occupation and $20.31 \%$ of them were still continuing current occupation.

As was shown in Figure 1, the most common traumatic stress factors were the sadness after a diagnosis of HF, which accounted for more than half of study sample ( $53.12 \%$ ). The second most common stress factor then was documented with economic difficulties (20.31\%).

The overall prevalence of depression ICD10 was $46.88 \%$ among the patients with HF. The proportion of depression fluctuated between $37.21 \%$ and $83.33 \%$ by NYHA classification. The highest percentage of depression was observed among NYHA class IV. In the univariate logistic regression analysis, compared to the prevalence of depression among patients with NYHA class II, the odds were 8.43 times higher for those with NYHA class IV (OR 8.43; 95\% CI 1.63-43.46) (Table II).

There were statistically significant differences between non-depressed group and depressed group, observed in age $\geq 60(p=0.021)$, gender ( $p$ $=0.035)$, marital status ( $p=0.001$ ), family caregivers's support in providing care $(\mathrm{p}=0.038)$, and HF impact on occupation ( $p<0.001$ ) (Table III).

Patient's age was significantly associated with increased prevalence of depression (OR 1.07; 95\% CI 1.003-1.14). It was significantly higher odds of depression in HF patients who felt sadness after a diagnosis of an illness (OR $18.02,95 \% \mathrm{CI}$ : 4.21-77.08). Individuals with higher household economic status were less likely to be diagnosed with depression compared to those with lower household economic status (OR 0.15, 95\% CI: $0.02-0.92$ ). The odds of depression were significantly higher in HF patients who reported with family conflict (OR 23.45, 95\% CI: 1.29-423.55), and in those having the loss of a close family member (OR 38.62, 95\% CI: 1.41-1055.98). Overall effect of multivariate logistic regression model was computed as McFadden's pseudo-R square, which was $50.40 \%$ (Table IV).

## Discussion

Our study demonstrates that the onset of depression is a common occurrence in HF inpatients and five significant factors contributing to depression were identified to be age of HF patient, sadness after a diagnosis of an illness, higher household economic status, family conflict and loss of a close family member. Present results suggest a need for a disease management program addressing both psychological and HF aspects with the aim of improving health outcomes for the inpatients in Vietnam health facilities. These associated factors should be taken into account in planning hospitalization disease management programs as well as screening and intervention programs for the HF inpatients.

Table II. Prevalence of depression among HF patients according to NYHA heart failure classification.

|  | Overall <br> Count <br> (\% of Total) | NYHA II <br> Count <br> (\% of Total) | NYHA III <br> Count <br> (\% of Total) | NYHA IV <br> Count <br> (\% of Total) | $\boldsymbol{p}$-value |
| :--- | :---: | :---: | :---: | :---: | :---: |
| Non-depression | $68(53.12)$ | $27(62.79)$ | $39(53.42)$ | $2(16.67)$ | $0.018^{*}$ |
| Depression $60(46.88)$ $16(37.21)$ | $34(46.58)$ <br> OR (95\% CI) | REF | $1.47(0.68-3.17)$ | $8.43(1.63-43.46)^{*}$ |  |

*Significant at 0.05 ; Examined the difference by $\chi^{2}$ test; REF: reference subgroup; OR: odd ratio; $95 \%$ CI: $95 \%$ confidence interval.

Table III. Difference of socioeconomic characteristics among non-depressed group and depressed group.

| Selected variables | Depression |  | $p$-values |
| :---: | :---: | :---: | :---: |
|  | No ( $\mathrm{n}=68$ ) <br> Count (\% of Total) | Yes ( $\mathrm{n}=\mathbf{6 0}$ ) <br> Count (\% of Total) |  |
| Age $\geq 60$ |  |  | $0.021^{\text {C* }}$ |
| No | 34 (50.00) | 18 (30.00) |  |
| Yes | 34 (50.00) | 42 (70.00) |  |
| Gender |  |  | $0.035^{\text {C* }}$ |
| Male | 41 (60.29) | 25 (41.67) |  |
| Female | 27 (39.71) | 35 (58.33) |  |
| Education |  |  | $0.933{ }^{\text {F }}$ |
| Illiteracy | 3 (4.41) | 4 (6.67) |  |
| Primary/Junior high school | 44 (64.71) | 38 (63.33) |  |
| High school | 8 (11.76) | 7 (11.67) |  |
| College | 11 (16.18) | 8 (13.33) |  |
| Tertiary and above | 2 (2.94) | 3 (5.00) |  |
| Ethnicity |  |  | $0.956{ }^{\text {C* }}$ |
| Kinh | 61 (89.71) | 54 (90.00) |  |
| Minority | 7 (10.29) | 6 (10.00) |  |
| Occupation |  |  | 0.178** |
| Blue-collar worker | 47 (69.12) | 31 (51.67) |  |
| White-collar worker | 5 (7.35) | 4 (6.67) |  |
| Freelancer | 4 (5.88) | 6 (10.00) |  |
| Retirer | 12 (17.65) | 19 (31.67) |  |
| Marital status |  |  | $0.001{ }^{\mathrm{F} * *}$ |
| Married | 58 (85.29) | 39 (65.00) |  |
| Single | 3 (4.41) | 1 (1.67) |  |
| Divorced | 1 (1.47) | 0 (0.00) |  |
| Widowed | 6 (8.82) | 20 (33.33) |  |
| Living area |  |  | $0.883{ }^{\text {C }}$ |
| Rural | 42 (61.76) | 36 (60.00) |  |
| Urban | 18 (26.47) | 18 (30.00) |  |
| Mountainous | 8 (11.76) | 6 (10.00) |  |
| Household economic status |  |  | $0.083{ }^{\text {C }}$ |
| Lower | 8 (11.76) | 14 (23.33) |  |
| Higher | 60 (88.24) | 46 (76.67) |  |
| Support from family caregivers in providing care |  |  | $0.038^{\text {F* }}$ |
| No | 7 (10.29) | 2 (3.33) |  |
| Yes | 61 (89.71) | 58 (96.67) |  |
| Impact of heart failure on occupation |  |  | $<0.001^{\text {C**** }}$ |
| Still continuing the current occupation | 21 (30.88) | 5 (8.33) |  |
| Do not continue the current occupation | 7 (10.29) | 24 (40.00) |  |
| Can do it but less | 40 (58.82) | 31 (51.67) |  |

F: Fisher's exact test; C: $\chi^{2}$ test; *, **, ***Significant at $0.05,0.01$, and 0.001 , respectively.

Depressive symptoms have been linked to the development and progression of HF and other cardiovascular diseases. Also, depression is known as a poor prognostic marker in patients with a diagnosis of HF. In the present study, we found depression developed in $46.88 \%$ of HF patients in our institution, which was in line with the studies of inpatients and outpatients when this figure ranged from 9 to $60 \%$ in a previous report ${ }^{10}$ and was estimated to be higher two-to-three times higher than the general population ${ }^{11}$. High prevalence of depression in our study was noted
because this phenomenon is, to date, often less measured and missed in clinical practice conditions in Vietnam health facilities. A point need to be highlighted here: most studies documented the depressive patients by means of using the instruments of PHQ and/or BDI, when we documented the patients with depression through a definitive diagnosis according to ICD-10 depression diagnostic criteria. This point was also consistent with available evidence ${ }^{12}$, indeed, our analysis indicated HF patients having a diagnosis of depression with range from $37.21 \%$ in NYHA classes

Table IV. Factors associated with depression: the multivariable regression model.

| Selected variables | OR | SE | $p$-values | 95\% CI |  |
| :---: | :---: | :---: | :---: | :---: | :---: |
|  |  |  |  | Lower | Upper |
| Age | 1.07 | 0.03 | 0.039* | 1.003 | 1.14 |
| Gender |  |  |  |  |  |
| Male | REF |  |  |  |  |
| Female | 3.66 | 2.62 | 0.069 | 0.90 | 14.92 |
| Education |  |  |  |  |  |
| Illiteracy | REF |  |  |  |  |
| Primary/Junior high school | 0.47 | 0.65 | 0.591 | 0.03 | 7.28 |
| High school | 1.69 | 3.21 | 0.779 | 0.04 | 69.51 |
| College | 0.53 | 0.98 | 0.736 | 0.01 | 19.50 |
| Tertiary and above | 2.63 | 5.71 | 0.655 | 0.03 | 184.31 |
| Ethnicity |  |  |  |  |  |
| Kinh | REF |  |  |  |  |
| Minority | 1.95 | 2.36 | 0.579 | 0.18 | 20.88 |
| Occupation |  |  |  |  |  |
| Blue-collar worker | REF |  |  |  |  |
| White-collar worker | 0.67 | 0.91 | 0.773 | 0.04 | 9.40 |
| Freelancer | 6.34 | 8.56 | 0.171 | 0.44 | 89.39 |
| Retirer | 0.84 | 0.93 | 0.877 | 0.09 | 7.32 |
| Marital status |  |  |  |  |  |
| Married | REF |  |  |  |  |
| Single | 0.69 | 1.26 | 0.84 | 0.01 | 24.97 |
| Divorced | N/A |  |  |  |  |
| Widowed | 2.41 | 2.40 | 0.378 | 0.34 | 17.05 |
| Living area |  |  |  |  |  |
| Rural | REF |  |  |  |  |
| Urban | 1.60 | 1.62 | 0.638 | 0.22 | 11.61 |
| Mountainous | 1.53 | 1.76 | 0.71 | 0.16 | 14.65 |
| Household economic status |  |  |  |  |  |
| Lower | REF |  |  |  |  |
| Higher | 0.15 | 0.13 | 0.041* | 0.02 | 0.92 |
| Support from family caregivers in providing care |  |  |  |  |  |
| No | REF |  |  |  |  |
| Yes | 2.65 | 3.96 | 0.514 | 0.14 | 49.70 |
| Impact of heart failure on occupation |  |  |  |  |  |
| Still continuing the current occupation | REF |  |  |  |  |
| Do not continue the current occupation | 1.05 | 1.23 | 0.96 | 0.10 | 10.36 |
| Can do it but less | 2.11 | 2.01 | 0.432 | 0.32 | 13.64 |
| Sadness after a diagnosis of an illness |  |  |  |  |  |
| No | REF |  |  |  |  |
| Yes | 18.02 | 13.36 | $<0.001^{* *}$ | 4.21 | 77.08 |
| Economic difficulties |  |  |  |  |  |
| No | REF |  |  |  |  |
| Yes | 1.03 | 0.77 | 0.963 | 0.23 | 4.49 |
| Family conflict |  |  |  |  |  |
| No | REF |  |  |  |  |
| Yes | 23.45 | 34.62 | 0.033* | 1.29 | 423.55 |
| Loneliness |  |  |  |  |  |
| No | REF |  |  |  |  |
| Yes | 1.06 | 1.46 | 0.966 | 0.07 | 15.87 |
| Work pressure |  |  |  |  |  |
| No | REF |  |  |  |  |
| Yes | 40.82 | 79.80 | 0.058 | 0.88 | 1883.25 |
| The loss of a close family member |  |  |  |  |  |
| No | REF |  |  |  |  |
| Yes | 38.62 | 65.19 | 0.03* | 1.41 | 1055.98 |
| Pseudo R2 | 0.5040 |  |  |  |  |

REF: reference subgroup; OR: odd ratio; SE: standard error; $95 \% \mathrm{CI}$ : $95 \%$ confidence interval; * **Significant at 0.05 and 0.001 , respectively.

II and $83.33 \%$ in NYHA class IV, along with the increasing prevalence by NYHA functional class. Notably, the univariate logistic regression model indicated that NYHA class IV increased the odds of in-hospital depression by 8.43 -fold; while recent systematic review of Mandana Moradi et al ${ }^{3}$ showed that odds of depression with any severity in HF patients was significantly 2.5 times higher with NYHA class III, IV compared to with NYHA class I, II.

Older age was documented as one of contributing factors for depression in various patient populations ${ }^{13,14}$ and especially this problem was prevalent in older adults ${ }^{15}$. This point was reasonable when older adults were common in our sample size of HF patients. In the Vietnamese study population with HF, we saw that older age was significantly associated with greater prevalence of depression. It is consistent with previous studies ${ }^{16,17}$. According to the best review of the research team, not many studies have analyzed the association between patient age and depression with quantitative variable of age without dividing the cut-off value, such as 60 years old, 65 years old or 70 years old.

A low socioeconomic status has been directly related to more frequent mental health problems and poorer cardiovascular disease outcomes ${ }^{18-21}$. In particular, depression leads to poor outcomes, including increased risk of poor functional status, hospital readmission and mortality for HF patients, and long-term caring burden on family caregivers, consequentially, contributing to the higher healthcare costs imposed on the family and the health system ${ }^{1,5}$. Our evidence supported that HF patients with higher household economic status were at lower risk of depression compared to those with lower household economic status.

Stressful life events, such as the loss of a close family member, have been proved to be one of the most stressful life-events increasing the risk of affective disorders, of which, depressive symptoms ${ }^{22,23}$. One of the most important findings from this study was that patients experiencing loss of a close family member added significant burden and greater depressive prevalence to HF patients. It may be reasonable that these stress factors have been linked to a burden for family caregivers, while caring for HF patient is a considerable burden for family caregivers ${ }^{24,25}$. Besides, the most striking result of our study was that patients who reported with family conflict were significantly more depressive compared to those without family conflict.

Despite providing meaningful findings in HF inpatients under a real study picture in Vietnam's clinical practice, several limitations need be considered in the interpretation. In this study context, we only considered socio-demographic factors and common stress factors that increased the vulnerability to the presence of depression. Besides the limitation of constructed causal relationships, generalizations of present inference to other contexts should also be noted because the study sample may only be representative of the actual HF inpatient population in our institution. We suggest further studies on a larger sample size with a broad variety of clinical and laboratory potential risk factors as well as lifestyle habits and treatment situation.

## Conclusions

The current results for the first time provide preliminary support for a potential role of socio-demographic factors and common stress factors for the presence of depression in HF inpatients. With relatively high prevalence of depression, the study also indicated that patient age, sadness after a diagnosis of an illness, household economic status, family conflict and loss of a close family member were significantly independently associated with depression in the inpatients with a diagnosis of HF. A collaboration on a routine basis between cardiologists and psychiatrists is needed to explore the prevalence and severe of depression in HF patients. A small difference in identifing symptoms of depression may be sufficient to considerably influence the course of a chronic long-term condition as HF.

## Authorship Contribution Statement

Nguyen-Ngoc Tran, Van-San Bui, and VanHai Nguyen contributed to the conception of this work and the design of the study. Thi-Phuong-Nam Hoang, Van-Hai Nguyen and Ho-ang-Long Vo acquired data. Nguyen-Ngoc Tran and Hoang-Long Vo performed the statistical analyses. Nguyen-Ngoc Tran, Van-San Bui, Van-Hai Nguyen, and Hoang-Long Vo conducted the interpretation of the data and drafted the manuscript. Hoang-Thanh Nguyen and Minh-Tam Duong revised the work critically for important intellectual content. All authors have approved the submitted version of the manuscript.

## Conflict of Interest

The Authors declare that they have no conflict of interests.

## Acknowledgements

The authors sincerely thank Board of Directors of Vietnam National Heart Institute, Bach Mai Hospital (Hanoi, Vietnam) for their best support in implementing this study. We also express our thanks to the patients and their family for the assistance in the entire of data collection.

## Funding

This research received no specific grant from any funding agency, commercial or not-for-profit sectors.

## Availability of Data and Materials

The data presented in this study are available upon request from the corresponding author. Due to the protection of personal data, the data are not publicly available.

## Authors' Contribution

Nguyen-Ngoc Tran, Van-San Bui, and Van-Hai Nguyen contributed to the conception of this work and the design of the study. Thi-Phuong-Nam Hoang, Van-Hai Nguyen and Hoang-Long Vo acquired data. Nguyen-Ngoc Tran and Hoang-Long Vo performed the statistical analyses. Nguy-en-Ngoc Tran, Van-San Bui, Van-Hai Nguyen, and Ho-ang-Long Vo conducted the interpretation of the data and drafted the manuscript. Hoang-Thanh Nguyen and MinhTam Duong revised the work critically for important intellectual content. All authors have approved the submitted version of the manuscript.

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