

CHA₂DS₂-VASc Score: A Simple But Not Appropriate Risk Score for Predicting Mortality Risk in Patients with Heart Failure

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Despite recent advances in heart failure (HF) therapies, the prognosis for patients suffering HF remains poor, with a 5-year 50% mortality rate.¹ However, estimating patient survival probability based on risk profile will help to identify high-risk patients, and assist patients and their families in selecting the most appropriate treatments or facilitate end-of-life decisions. There are numerous factors associated with adverse outcomes in HF including demographic variables (e.g., advanced age, male, ischemic etiology, and resuscitated sudden death),^{2,3} clinical factors [e.g., hypotension, New York Heart Association (NYHA) functional class III-IV, and prior HF hospitalization],⁴ co-morbidities (e.g., anemia, diabetes, and renal dysfunction),⁵ imaging markers (e.g., ejection fraction, global longitudinal strain),^{6,7} and serum markers (e.g., hyponatremia, natriuretic peptides).⁷⁻⁹ However, no single parameter can be expected to reliably predict outcomes in HF. The combination of several variables in prognostic models has emerged as the most appropriate approach to predict outcomes in the heterogeneous HF population.¹⁰⁻¹²

In a systematic review of risk prediction models in HF between January 1995 to March 2013, 64 models from 48 studies were found.¹⁰ Of the 64 models, 43 models predicted death, 10 hospitalization, and 11 death or hospitalization.¹⁰ Despite the marked differences in clinical factors, demographic data, and choice of candidate variables, several variables emerged as consistent and strong predictors of mortality.¹⁰ These

variables included age, renal function, blood pressure, blood sodium, ejection fraction, male, NYHA functional class, diabetes, natriuretic peptides, weight or body mass index, and exercise tolerance.¹⁰ In this review,¹⁰ two prediction models for mortality were recommended: the Cardiac and Comorbid Conditions HF (3C-HF) score, and the Meta-Analysis Global Group in Chronic HF (MAGGIC) score.^{11,12} The 3C-HF score has a very good discriminatory ability for predicting 1-year mortality (C-statistic of 0.87), has been externally validated (C-statistic of 0.82), and enables risk calculations in a wide variety of cases of HF.¹¹ The MAGGIC integer score used information from 30 studies and 39,372 HF patients to derive a risk score for three-year mortality.¹² The large size of the study and the derivation of patients from wide geographic regions provide a distinctively robust and generalizable tool to assess mortality risk in HF.¹²

The CHA₂DS₂-VASc score [congestive heart failure, hypertension, age ≥ 75 years (doubled), diabetes, stroke/transient ischemic attack (doubled), vascular disease, age 65-74 years, female] is used clinically for stroke risk stratification in nonvalvular atrial fibrillation (AF).¹³ Recently, use of the CHA₂DS₂-VASc score in predicting stroke risk in AF has extended beyond AF and stroke risk predicting.¹⁴ In a cohort study of 42,987 HF patients not receiving anticoagulation, the CHA₂DS₂-VASc score was able to predict mortality in patients with and without AF, with a high negative predictive value (92% and 91% for patients with and without AF, respectively) at 1-year follow-up.¹⁴ The absolute risks of stroke, thromboembolism and death increased in a comparable manner in patients with and without AF, exhibiting a clear dose-response relationship.¹⁴ In addition, a higher CHA₂DS₂-VASc score has been shown to be associated with increased risks of acute stent thrombosis after percutaneous coronary intervention,¹⁵ mortality in acute pulmonary embolism,¹⁶ and no reflow phenomenon in

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ST-elevation myocardial infarction.¹⁷

In this issue of *Acta Cardiologica Sinica*, Yayla et al. conducted a prospective study to investigate the association between the CHA₂DS₂-VAsc score and mortality in 106 hospitalized HF patients.¹⁸ The authors have shown that the CHA₂DS₂-VAsc score was independently predictive of mortality in HF patients with reduced ejection fraction in a conditional stepwise regression model.¹⁸ In this study, 21 patients (20%) died within one year.¹⁸ Compared to the surviving patients, the patients who died were more likely to have a poorer NYHA functional class, higher values of urea, creatinine, B-natriuretic peptide, and CHA₂DS₂-VAsc score, as well as lower hemoglobin. In the multivariate regression analysis, the CHA₂DS₂-VAsc score and NYHA functional class III-IV were independent predictors of mortality.¹⁸ The authors observed that 1 point increase in the CHA₂DS₂-VAsc score was associated with a 2-fold increase in mortality (hazard ratio 2.13, 95% confidence interval 1.48-3.15). However, there are several points that require further attention.¹⁸ First, the patients in this study are younger (mean 60 y/o) than most of the HF studies (mean 67-70 y/o).¹⁰⁻¹² Therefore, the generalization of the results to other HF populations is questionable. Second, there was no significant age difference between the surviving and non-surviving patients. However, in the CHA₂DS₂-VAsc score, age is a significant risk marker, where a double weight is factored in for patients who are 75 y/o or older.¹³ It is not appropriate to apply the CHA₂DS₂-VAsc score for predicting mortality in this study cohort. Third, male rather than female was more likely to be associated with higher mortality risk in most of the prior HF studies.¹⁰ The association between stroke/transient ischemic attack and mortality in HF has not been previously documented.¹⁰ Therefore, it is not appropriate to apply the CHA₂DS₂-VAsc score for predicting mortality in HF.

Despite the factor that a CHA₂DS₂-VAsc score is a simple risk score and obtainable in most of the patients,¹⁴ the score was originally designed and validated for predicting stroke risk in nonvalvular AF.¹³ It is not appropriate to be used for mortality prediction in HF. For mortality prediction, we suggest evaluation of all possible factors to determine the independent risk factors, or use a prior established risk model for outcome prediction.^{11,12}

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REFERENCES

1. Mosterd A, Hoes AW. Clinical epidemiology of heart failure. *Heart* 2007;93:1137-46.
2. Lee DS, Stitt A, Austin PC, et al. Prediction of heart failure mortality in emergent care a cohort study. *Ann Intern Med* 2012;156:767-75.
3. Barlera S, Tavazzi L, Franzosi MG, et al. Predictors of mortality in 6975 patients with chronic heart failure in the gruppo italiano per lo studio della streptochinasi nell'infarto miocardico-heart failure trial proposal for a nomogram. *Circ Heart Fail* 2013;6:31-9.
4. Bayes-Genis A, de Antonio M, Galan A, et al. Combined use of high-sensitivity st2 and nt probnp to improve the prediction of death in heart failure. *Eur J Heart Fail* 2012;14:32-8.
5. Smith DH, Thorp ML, Gurwitz JH, et al. Chronic kidney disease and outcomes in heart failure with preserved versus reduced ejection fraction: the cardiovascular research network preserve study. *Circ Cardiovasc Qual Outcomes* 2013;6:333-42.
6. Curtis JP, Sokol SJ, Wang Y, et al. The association of left ventricular ejection fraction, mortality, and cause of death in stable outpatients with heart failure. *J Am Coll Cardiol* 2003;42:736-42.
7. Sengelov M, Jorgensen PG, Jensen JS, et al. Global longitudinal strain is a superior predictor of all-cause mortality in heart failure with reduced ejection fraction. *JACC Cardiovasc Imaging* 2015;8:1351-9.
8. Wright GA, Struthers AD. Natriuretic peptides as a prognostic marker and therapeutic target in heart failure. *Heart* 2006;92:149-51.
9. Kozdag G, Ertas G, Kilic T, et al. Elevated level of high-sensitivity c-reactive protein is important in determining prognosis in chronic heart failure. *Med Sci Monit* 2010;16:CR156-61.
10. Rahimi K, Bennett D, Conrad N, et al. Risk prediction in patients with heart failure: a systematic review and analysis. *JACC Heart Fail* 2014;2:440-6.
11. Senni M, Parrella P, De Maria R, et al. Predicting heart failure outcome from cardiac and comorbid conditions: the 3C-HF score. *Int J Cardiol* 2013;163:206-11.
12. Pocock SJ, Ariti CA, McMurray JJ, et al. Predicting survival in heart failure: a risk score based on 39372 patients from 30 studies. *Eur Heart J* 2013;34:1404-13.
13. Lip GY, Nieuwlaat R, Pisters R, et al. Refining clinical risk stratification for predicting stroke and thromboembolism in atrial fibrillation using a novel risk factor-based approach: the euro heart sur-

- vey on atrial fibrillation. *Chest* 2010;137:263-72.
14. Melgaard L, Gorst-Rasmussen A, Lane DA, et al. Assessment of the CHA₂DS₂-VASc score in predicting ischemic stroke, thromboembolism, and death in patients with heart failure with and without atrial fibrillation. *JAMA* 2015;314:1030-8.
 15. Heestermans AA, van Werkum JW, Zwart B, et al. Acute and sub-acute stent thrombosis after primary percutaneous coronary intervention for ST-segment elevation myocardial infarction: incidence, predictors and clinical outcome. *J Thromb Hhaemost* 2010;8:2385-93.
 16. Onuk T, Karatas MB, Ipek G, et al. Higher CHA₂DS₂-VASc score is associated with increased mortality in acute pulmonary embolism. *Clin Appl Thromb Hemost* 2016:1076029615627341.
 17. Ipek G, Onuk T, Karatas MB, et al. CHA₂DS₂-VASc score is a predictor of no-reflow in patients with ST-segment elevation myocardial infarction who underwent primary percutaneous intervention. *Angiology* 2016;67:840-5.
 18. Yayla C, Temizer O, Acar B, et al. The association between CHA₂DS₂-VASc score and mortality in patients with heart failure with reduced ejection fraction. *Acta Cardiol Sin* 2017;33:429-35.

