Cardiovascular disease and mental health: a dangerous duo?

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This editorial refers to 'Screening for anxiety and depression in clinical practice: translating scores from World Health Organization-5/Anxiety Symptom Scale-2/Major Depression Inventory-2 to Hospital Anxiety and Depression Scale', by N.F. Johnsen et al., https://doi.org/ 10.1093/eurjpc/zwad180.

Psychiatric disorders represent a major healthcare burden, affecting as many as one of five adults during their lifetime, with relevant consequences on cardiovascular care. Indeed, on top of being extremely prevalent among patients with cardiovascular disease, significantly impacting on quality of life and representing a major cause of morbidity, mental disorders constitute a major risk factor for developing major adverse cardiovascular events.^{1–3} Major depression, in particular, is associated with reduced quality of life and increased mortality, and even more so among individuals with established cardiovascular disease.⁴

Notably, among patients who suffered from myocardial infarction, major depression is as prevalent as 20%, ^{5,6} being associated with significantly worse outcomes.⁷ Furthermore, those with established cardiovascular disease and also receiving a diagnosis of major depression have a 2–4 times higher risk of recurrent adverse cardiovascular event at follow-up.^{7–10} Anxiety disorder as well is a widespread condition with a roughly 20% prevalence in the general population. The detrimental role of such specific psychiatric condition with respect to cardiovascular health has been demonstrated by a recent meta-analysis, showing that anxious persons have a risk of cardiac death roughly double with respect to general population.¹¹

Among the mechanisms able to explain the interaction among psychiatric and cardiac disorders, a pivotal role is thought to be played by dysregulation both of autonomous nervous system and the hypothalamic-pituitary-adrenal axis, with increasing levels of cortisol and cathecholamines, both plasmatic and urinary, and a decreased heart rate variability, with higher mean and average daily heart rate. Alongside such detrimental effects on physiology, it is worth mentioning the behavioural effect of major depression and anxiety, with higher tendency of the patient towards unhealthy lifestyles such as cigarette smoking, poor dietary habits, lack of physical activity, and low adherence to medical therapy. Such amount of evidence led American Heart Association in 2014 to mention depression among risk factors related to poor prognosis after acute coronary syndrome.¹²

Despite the prevalence and relevance of such disease, a recent questionnaire showed how diagnosing and treating depression is often felt as beyond the scope of everyday clinical practice, as nearly half of the colleagues felt not comfortable with making diagnosis of depression during clinical practice as well as treating depression. Nonetheless, screening for depression in cardiovascular patients has been advocated in recent international guidelines,^{13,14} and given the availability of efficacious treatment, is a worthy target to be pursued for the sake of improving patient's quality of life.

To this purpose, Johnsen et al. report in this issue of the Journal a timely and important study addressing the relevant issue on how to effectively screen patients with cardiovascular disease for anxiety and depression.¹⁵ As a matter of fact, the adoption of an effective screening tool should represent a trade-off between diagnostic accuracy and ease of use. Over 10000 patients with a cardiovascular diagnosis (either atrial fibrillation, heart failure, heart valve disease, or ischaemic heart disease) from the Danish National Patient Register were offered a dedicated survey including 51 questions, comprising validated questionnaires such as Anxiety Symptom Scale-2 (ASS-2), Hospital Anxiety and Depression Scale (HADS), Major Depression Inventory-2 (MDI-2), and World Health Organization-5 (WHO-5), thanks to the cooperation of several healthcare providers. After pilot testing, the questionnaire in 1% of the sample was sent over, receiving over 4000 responses. Such a strenuous effort made it possible to obtain conversion tables enabling cross-walk among HADS-A and WHO-5/ASS-2, on one hand, and HADS-D and WHO-5/MDI-2, on the other. Using bi-factor confirmatory factor analysis, it was possible to obtain translation tables to estimate HADS values specific to each cardiovascular condition, starting from different combinations of ASS, MDI-2, and WHO-5 (Figure 1).

Despite being clinically validated, with higher score values clearly linked to worse cardiovascular prognosis; HADS is worldwide protected by copyright laws, a feature that has severely limited its use. The availability of a questionnaire relatively easy to administer might certainly contribute to a more widespread use of such an important tool, minimizing the percentage of cardiovascular patients suffering from psychiatric disorders, with a significant impact both on quality

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SCREENING FOR ANXIETY/ DEPRESSION IN CARDIAC PATIENTS

HADS-A, WHO-5/ASS-2 HADS-D, WHO-5/MDI-2

ASSIGN, FHS QRISK3, SCORE

SCREENING FOR CARDIAC DISEASE

Figure 1 Balancing screening for anxiety or depression among patients with or at risk of cardiovascular disease, and screening for cardiovascular disease among individuals with or at risk of anxiety or depression. ASS-2, Anxiety Symptom Scale-2; FRS, Framingham Risk Score; HADS, Hospital Anxiety and Depression Scale; MDI-2, Major Depression Inventory-2; WHO-5, World Health Organization-5.

of life and prognosis. As the authors clearly acknowledge, the proposed test is the result of the combination of three different questionnaires, two of which are negatively phrased with the other one positively phrased, making it necessary for healthcare providers to avoid misunderstandings while administering the questionnaire. Other tools are available as well: whereas the European Society of Cardiology does not mention a specific questionnaire for depression and anxiety screening,¹² the American Heart Association endorses screening of all patients with coronary artery disease with Patient Health Questionnaire (PHQ)-2, and to test with 9-item PHQ-9 only those who test positive.^{13,14} Regardless of the chosen test, further effort is required in those cases in which the test results positive in order to properly provide high-quality psychiatric support: a multidisciplinary approach should be considered with referral of patients to mental health specialists, especially if there is substantial risk of harm either to self or others.

In conclusion, the improvement of survival and quality of life after diagnosis of cardiovascular disease has been made possible at the expense of a path that the patient has to go through, comprising repeated medical consultations, rigorous control of risk factors, and strict adherence to medical therapy, as even the most sophisticated drugs do not work if not assumed by the patient. Good mental health and positive patient– physician relationship are key to boost outcomes, benefiting the most from therapeutical advances in the cardiovascular field, and this holds even truer in the current pandemic era.¹⁶ For this reason, screening for depression and anxiety in cardiovascular patients is beneficial. Several validated clinical questionnaires are already available, and the study by Johnsen et al. adds a novel, free-to-use tool that might be implemented in everyday clinical practice, aiming to a patient-centred, holistic approach in the treatment of cardiovascular diseases.

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References

- Hasin DS, Sarvet AL, Meyers JL, Saha TD, Ruan WJ, Stohl M, et al. Epidemiology of adult DSM-5 major depressive disorder and its specifiers in the United States. JAMA Psychiat 2018;75:336–346.
- Yusuf S, Hawken S, Ounpuu S, Dans T, Avezum A, Lanas F, et al. Effect of potentially modifiable risk factors associated with myocardial infarction in 52 countries (the INTERHEART study): case-control study. *Lancet* 2004;**364**:937–952.
- Kewcharoen J, Tachorueangwiwat C, Kanitsoraphan C, Saowapa S, Nitinai N, Vutthikraivit W, et al. Association between depression and increased risk of readmission in patients with heart failure: a systematic review and meta-analysis. *Minerva Cardiol Angiol* 2021;69:389–397.
- Roncella A, Pristipino C. Psychotherapy for Ischemic Heart Disease: An Evidence-based Clinical Approach. Hauppauge, Switzerland: Springer International; 2016.
- Ziegelstein RC. Depression in patients recovering from a myocardial infarction. JAMA 2001;286:1621–1627.
- Thombs BD, Bass EB, Ford DE, Stewart KJ, Tsilidis KK, Patel U, et al. Prevalence of depression in survivors of acute myocardial infarction. J Gen Intern Med 2006;21:30–38.
- Meijer A, Conradi HJ, Bos EH, Anselmino M, Carney RM, Denollet J, et al. Adjusted prognostic association of depression following myocardial infarction with mortality and cardiovascular events: individual patient data meta-analysis. Br J Psychiatry 2013;203:90–102.
- Meijer A, Conradi HJ, Bos EH, Thombs BD, van Melle JP, de Jonge P. Prognostic association of depression following myocardial infarction with mortality and cardiovascular events: a meta-analysis of 25 years of research. *Gen Hosp Psychiatry* 2011;**33**:203–216.
- Nicholson A, Kuper H, Hemingway H. Depression as an aetiologic and prognostic factor in coronary heart disease: a meta-analysis of 6362 events among 146 538 participants in 54 observational studies. *Eur Heart J* 2006;27:2763–2774.

- Frasure-Smith N, Lespérance F, Talajic M. Depression and 18-month prognosis after myocardial infarction. *Circulation* 1995;91:999–1005.
- Roest AM, Martens EJ, de Jonge P, Denollet J. Anxiety and risk of incident coronary heart disease: a meta-analysis. J Am Coll Cardiol 2010;56:38–46.
- 12. Lichtman JH, Bigger JT Jr, Blumenthal JA, Frasure-Smith N, Kaufmann PG, Lespérance F, et al. Depression and coronary heart disease: recommendations for screening, referral, and treatment: a science advisory from the American Heart Association Prevention Committee of the Council on Cardiovascular Nursing, Council on Clinical Cardiology, Council on Epidemiology and Prevention, and Interdisciplinary Council on Quality of Care and Outcomes Research: endorsed by the American Psychiatric Association. *Circulation* 2008;**118**:1768–1775.
- Visseren FLJ, Mach F, Smulders YM, Carballo D, Koskinas KC, Bäck M, et al. 2021 ESC guidelines on cardiovascular disease prevention in clinical practice. Eur Heart J 2021;42: 3227–3337.
- 14. Lichtman JH, Froelicher ES, Blumenthal JA, Carney RM, Doering LV, Frasure-Smith N, et al.; American Heart Association Statistics Committee of the Council on Epidemiology and Prevention and the Council on Cardiovascular and Ctroke Nursing. Depression as a risk factor for poor prognosis among patients with acute coronary syndrome: systematic review and recommendations: a scientific statement from the American Heart Association. *Circulation* 2014;**129**:1350–1369.
- 15. Johnsen NF, Jensen SN, Christensen KB, Pedersen SS, Helmark C, Zwisler A-D, et al. Screening for anxiety and depression in clinical practice: translating scores from World Health Organization-5/Anxiety Symptom Scale-2/Major Depression Inventory-2 to Hospital Anxiety and Depression Scale. *Eur J Prev Cardiol* 2023;**30**:1689–1701.
- Mazza M, Marano G, Antonazzo B, Cavarretta E, Nicola M DI, Janiri L, et al. What about heart and mind in the COVID-19 era? *Minerva Cardiol Angiol* 2021;69: 222–226.