LETTER TO THE EDITOR

Red cell distribution width is a predictor of mortality in patients with major bleeding

A largura de distribuição dos eritrócitos é um preditor de mortalidade em pacientes com hemorragia grave

Dear Editor,

We have read with great enthusiasm the recently published article entitled "Impact of red blood cell distribution width on risk for bleeding events in patients with non-ST elevation acute coronary syndromes" by Gonçalves et al.1 In this well-presented article they aimed to determine the prognostic value of red blood cell distribution width (RDW) in patients with non-ST segment elevation acute coronary syndromes (NSTE-ACS), particularly regarding the risk of major bleeding. They concluded that RDW was an independent predictor of major bleeding in patients with non-ST segment elevation myocardial infarction (NSTEMI). They suggested that higher RDW values were associated with greater in-hospital mortality and were an independent predictor of in-hospital major bleeding in patients with NSTEMI. The ready availability of this parameter at no additional cost may encourage its wider use in clinical practice in the future. We thank the authors for their contribution.

It is known that anemia is an important predictor of short- and medium-term mortality after ACS.2 In the present study, there were significant differences between the third tertile of RDW compared to the other groups. Anemia can be an independent risk factor of ACS patients with major bleeding in the present study. RDW is a standard laboratory parameter that shows variation in red blood cell size on a standard hemogram and is usually used in assessing blood diseases. It has recently been reported as an independent predictor of all-cause long-term mortality in patients with coronary artery disease.3 However, conditions in the differential diagnosis of anemias can affect RDW and so this parameter might change in the presence of abnormalities such as thyroid disease, renal or hepatic dysfunction (creatinine >1.5 mg/dl or aspartate aminotransferase and alanine transaminase more than twice the upper limit of normal, respectively), inflammatory diseases, and use of medications which could influence RDW. On the other hand, it is also reported that an increased RDW is associated with nutritional deficiency (of iron, vitamin B12, and folic acid) and ethnicity.4

Recent clinical studies have considered the importance of bleeding events in the prognosis in patients with ACS. In fact, the risk for cardiovascular complications associated with bleeding is similar to that of a myocardial infarction (MI). Some bleeding risk scores are now available that dependably quantify the likelihood of an ACS patient experiencing a bleeding complication. There is a strong relationship between bleeding and some conditions including older age,5 renal impairment, female gender, and an invasive management approach. Although patients who tend to bleed frequently have more comorbidities than their non-bleeding counterparts, several lines of experimental and clinical evidence suggest an independent causal pathway for bleeding-associated cardiovascular risk. When possible, bleeding should be avoided, and strategies such as use of risk scores, novel antithrombotics (such as bivalirudin and fondaparinux), vascular closure devices and radial access may decrease major bleeding.6

Reduced glomerular filtration rate (GFR) may be also associated with adverse outcomes in patients with cardiovascular disease. The authors used the Cockcroft-Gault (CG) equation for GFR. However, compared to the Modification of Diet in Renal Disease (MDRD) formula, the CG equation can give lower GFR measurements in younger age groups but higher GFR measurements in older individuals. The value of GFR in predicting various cardiovascular endpoints in patients with MI is established. Compared to the CG formula, the MDRD formula is significantly more accurate in predicting the severity of coronary artery disease and two-year cardiovascular risk in patients admitted to the intensive care unit with MI.7 For this reason, it can be useful, and the results of the study might have been different if the authors had also used the MDRD formula.

In a previous study it was found that elevated levels of inflammatory molecules are markers of in-hospital major bleeding and also indicate an increased risk for the progression of NSTEMI. These molecules can be reduced by medications including antihypertensive therapy8 and aspirin. Additionally, as well as RDW, neutrophil/lymphocyte ratio, gamma-glutamyltransferase, C-reactive protein, mean platelet volume and uric acid are also easy methods to assess patients’ cardiovascular disease.9 Moreover,

---

1 At the time of publication, the authors of the article had not responded to requests for a response.

2174-2049/S - see front matter © 2013 Sociedade Portuguesa de Cardiologia. Published by Elsevier España, S.L. All rights reserved.
hypoalbuminemia on admission is also a strong independent predictor of long-term mortality and development of advanced HF in patients with STEMI undergoing percutaneous coronary intervention. These markers might be useful in clinical practice. Finally, it would have been better if the authors had defined how much time they specified for measuring RDW levels, since delaying blood sampling can cause abnormal results in RDW measurements.

In conclusion, we strongly believe that the findings of the current study will lead to further large-scale studies examining the relationship between RDW and in-hospital major bleeding in patients presenting with ACS. However, one should keep in mind that RDW alone, without other inflammatory markers, may not give precise information to clinicians about their patients’ inflammatory condition and prognostic indication. So, on this basis we think that it should be evaluated together with other serum inflammatory markers.

**Conflicts of interest**

The authors have no conflicts of interest to declare.

**References**


Sevket Balta*, Sait Demirkol, Murat Unlu, Turgay Celik

*Corresponding author.

**E-mail address:** drsevketb@gmail.com (S. Balta).