

1 **Abstract**

2 **Background**

3 Decrease in histidine-rich glycoprotein (HRG) was reported as a cause of dysregulation
4 of the coagulation-fibrinolysis and immune systems, leading to multi-organ failure, and
5 it may be a biomarker for sepsis, ventilator-associated pneumonia, preeclampsia, and
6 coronavirus disease 2019. However, the usefulness of HRG in perioperative
7 management remains unclear. This study aimed to assess the usefulness of HRG as a
8 biomarker for predicting postoperative complications.

9 **Methods**

10 This was a single-center, prospective, observational study of 150 adult patients who were
11 admitted to the intensive care unit after surgery. Postoperative complications were defined
12 as those having a grade II or higher in the Clavien–Dindo classification, occurring within
13 7 days after surgery. The primary outcome was HRG levels in the patients with and
14 without postoperative complications. The secondary outcome was the ability of HRG,
15 white blood cell, C-reactive protein, procalcitonin, and presepsin to predict postoperative
16 complications. Data are presented as number and median (interquartile range).

17 **Results**

18 The incidence of postoperative complications was 40%. The HRG levels on
19 postoperative day 1 were significantly lower in patients who developed postoperative
20 complications (n=60; 21.50 [18.12–25.74] $\mu\text{g}/\text{mL}$) than in those who did not develop
21 postoperative complications (n=90; 25.46 [21.05–31.63] $\mu\text{g}/\text{mL}$). The Harrell C-index
22 scores for postoperative complications were HRG, 0.65; white blood cell, 0.50; C-
23 reactive protein, 0.59; procalcitonin, 0.73; and presepsin, 0.73. HRG was independent
24 predictor of postoperative complications when adjusted for age, the presence of
25 preoperative cardiovascular comorbidities, American Society of Anesthesiologists
26 Physical Status Classification, operative time, and the volume of intraoperative bleeding
27 (adjusted hazard ratio=0.94; 95% confidence interval, 0.90–0.99).

28 **Conclusions**

29 The HRG levels on postoperative day 1 could predict postoperative complications.
30 Hence, HRG may be a useful biomarker for predicting postoperative complications.