Liver disease and mortality among patients with hip fracture: does gender matter?

Dear editor,

Several studies have revealed that the 1-year all-cause mortality rate was about 20% following hip fracture (HF), which highlights the importance of the prognostic impact of other comorbidities on mortality.\(^1\)

Recently, we read with great interest the study by Montomoli et al.\(^2\) The authors performed a nationwide cohort study to investigate the prognostic impact of liver disease on 30-day mortality and the 31–365-day mortality among patients with HF. They concluded that relative to patients without liver disease, patients with liver disease, especially those with liver cirrhosis, had higher 30-day mortality and 31–365-day mortality rate following HF. The research appears informative clinically. Thus, there are some issues that that should be addressed regarding this study.

First, we were confused regarding the method for the diagnosis of liver cirrhosis. Because the early phase of liver cirrhosis has no apparent signs and symptoms, diagnosis in some patients can be missed without the detailed inspection. Consequently, the prevalence of liver cirrhosis may be underestimated, which may affect the results.

Second, ~90% of patients with HF received hip surgery in this study. Much more patients were elderly and classified as high comorbidity level, which was the risk factor for surgical complication and related to the worsened prognosis.\(^3\) Nevertheless, the surgery-associated complications, such as chest infection, early failure pneumo-nia, and pulmonary embolism, were not reported in the study by Montomoli et al.\(^2\) Indeed, this information is critical in analyzing the association between liver disease and mortality in HF patients.

Third, the nonalcoholic fatty liver disease (NAFLD) is the most common liver disorder all over the world. Previous pieces of evidence suggested that the associations between NAFLD and mortality were different between men and women.\(^4\) NAFLD was associated with higher risk of all-cause mortality and death from cancer and cardiovascular disease in women; but these associations were not detected in men.\(^4\) This finding was also observed in the study by Montomoli et al,\(^2\) which showed that association of noncirrhotic liver disease and overall mortality was sex-based (HR: 1.10; 95% CI: 0.90–1.35 for man and HR: 1.36; 95% CI: 1.15–1.61 for women, respectively). The reason for this finding may be due to the levels of sex hormones, which is more closely associated with obesity, metabolic phenotype, and the degree of inflammation.\(^4\) Thus, further studies are warranted to investigate the mechanisms responsible for sex-based differences in the association between the liver disease and mortality.

Ling-Yue Zhao
Yi-Jun Luo
Jing Zhu
Department of Ambulatory Surgery, Shenzhen Nanshan District People’s Hospital, Shenzhen, Guangdong 518052, People’s Republic of China

Correspondence: Yi-Jun Luo
Department of Day Surgery, Shenzhen Nanshan District People’s Hospital, 89 Taoyuan Road, Shenzhen, Guangdong 518052, People’s Republic of China
Tel +86 755 513 1121
Fax +86 755 513 1121
Email nsluyijun@yeah.net
Disclosure
The authors report no conflicts of interest in this communication.

References
Authors’ reply
Jonathan Montomoli
Rune Erichsen
Henrik Gammelager
Alma B Pedersen
Department of Clinical Epidemiology, Aarhus University Hospital, Aarhus N, Aarhus, Denmark

Correspondence: Jonathan Montomoli
Department of Clinical Epidemiology, Aarhus University Hospital, Olof Palmes Allé 43-45, 8200 Aarhus N, Aarhus, Denmark
Tel +45 8716 7212
Fax +45 8716 7215
Email jonathan.montomoli@gmail.com

Dear editor
We thank Zhao et al for showing interest in our work.1 First, we agreed that misclassification of patients with asymptomatic liver disease may have hampered our results. Hence, some patients with liver disease are likely to have been categorized in the cohort without liver disease causing us to underestimate the prognostic impact of liver disease on hip fracture (HF).

Second, with respect to complications following HF, we agree that these intermediate steps are clinically relevant to study as the underlying cause for the increased mortality. However, studying complications after HF and subsequent mortality requires detailed mediation analyses and thus falls outside the scope of our study.

Third, and finally, we acknowledge that studying the impact of nonalcoholic fatty liver disease on HF prognosis would be interesting and important. Unfortunately, our data did not provide the opportunity to differentiate nonalcoholic fatty liver disease from other noncirrhotic liver diseases, and we were thus unable to investigate further. The fact that we observed differences in hazards ratios for men and women may be explained by underlying differences in hormone levels as suggested by Zhao et al, but this is purely speculative and not something our data provides basis for.

Disclosure
The authors report no conflicts of interest in this communication.

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