



OPEN ACCESS

EDITED AND REVIEWED BY
Katherine Samaras,
St Vincent's Hospital Sydney, Australia

*CORRESPONDENCE
Christiano Argano
✉ chargano@yahoo.it

RECEIVED 06 November 2023
ACCEPTED 14 November 2023
PUBLISHED 22 November 2023

CITATION
Argano C (2023) Editorial: Interactions
between NAFLD and cardiac conduction,
structure and function: recent advances
and treatments.
Front. Endocrinol. 14:1334227.
doi: 10.3389/fendo.2023.1334227

COPYRIGHT
© 2023 Argano. This is an open-access
article distributed under the terms of the
[Creative Commons Attribution License
\(CC BY\)](https://creativecommons.org/licenses/by/4.0/). The use, distribution or
reproduction in other forums is permitted,
provided the original author(s) and the
copyright owner(s) are credited and that
the original publication in this journal is
cited, in accordance with accepted
academic practice. No use, distribution or
reproduction is permitted which does not
comply with these terms.

Editorial: Interactions between NAFLD and cardiac conduction, structure and function: recent advances and treatments

Christiano Argano*

Department of Internal Medicine, National Relevance and High Specialization Hospital Trust ARNAS Civico, Di Cristina, Benfratelli, Palermo, Italy

KEYWORDS

NAFLD, MAFLD, atrial fibrillation, atrial remodeling, left ventricular diastolic dysfunction, monocyte-to-high-density lipoprotein ratio (MHR), heart failure, mortality

Editorial on the Research Topic

Interactions between NAFLD and cardiac conduction, structure and function: recent advances and treatments

Non-alcoholic fatty liver disease (NAFLD) represents the most common liver disease globally (1), and its prevalence has been rising along with the obesity and type 2 diabetes epidemic (2). According to a recent systematic review, the global prevalence of NAFLD increased from 25.3% in 1990-2006 to 38.0% in 2016-2019 (3). Consequently, NAFLD is becoming a major public healthcare challenge (4). It causes a deterioration of quality of life, which increases with disease progression and is worsened by comorbidities (5, 6), representing the leading cause of liver-related morbidity and mortality (7). NAFLD does not only affect the liver but was shown to be a multisystem disease with increased risk of type 2 diabetes, dyslipidaemia, metabolic syndrome, hypertension, kidney and cardiovascular diseases (8). Increasing literature data suggests that subjects with NAFLD show myocardial functional and structural changes, leading to cardiac remodelling and increased risk of heart failure (HF) (9). In particular, different studies showed associations between NAFLD and left ventricular and diastolic dysfunction (9, 10), right ventricular dysfunction (11) and left ventricular hypertrophy (12). Moreover, recent data showed that NAFLD is associated with an increased risk of incident atrial fibrillation (AF) (13) and subjects with higher hepatic fibrosis index were associated with increased AF risk (14).

Mechanisms that link NAFLD to cardiac functional and structural changes, apart from the common risk factors, have yet to be explored. In NAFLD, insulin resistance, inflammation and oxidative stress may lead to cardiac insulin resistance and cardiac fibrosis, determining an alteration in cardiac structure (15). Moreover, the increased myocardial fatty acid oxidation, which is less efficient than glucose metabolism, may contribute to the development of heart failure (16).

This Research Topic aimed to provide insight into several aspects of the connections between NAFLD cardiac conduction, structure and function, presenting novelties on the potential pathological mechanism at the basis of these relationships. Overall, four original articles and two meta-analyses have been published.

In this Research Topic, Peng et al. after underlining the importance of the current debate on renaming NAFLD to metabolic dysfunction associated fatty liver disease (MAFLD), aimed to examine whether MAFLD is associated with left ventricular diastolic dysfunction and cardiac remodelling. Additionally, the authors try to identify the impact of different subgroups (lean, overweight/obese and diabetes) and the severity of MAFLD (normal, mild, moderate, and severe hepatic steatosis). Left ventricular diastolic dysfunction was significantly more prevalent in the MAFLD group than in the normal group. The overweight and diabetes subgroups were significantly associated with cardiac alterations such as interventricular septum thickness, LV posterior wall thickness, left atrial diameter, relative wall thickness, and LV mass index. In addition, subjects with moderate to severe steatosis had higher risks for left ventricular diastolic dysfunction and cardiac remodelling.

Decoin et al. brought new insights into the association between NAFLD/MAFLD and atrial fibrillation. The authors showed that the presence of MAFLD patients at risk of liver fibrosis is associated with adverse atrial remodelling, particularly an increase in LA volume, impaired LA reservoir function, and increased low-voltage areas. In addition, the liver fibrosis scoring in MAFLD patients predicted AF recurrence after ablation. In MAFLD subjects with AF recurrence, high liver fibrosis scores presented a higher AF burden. The most significant conclusion to be drawn is that liver fibrosis scoring in MAFLD patients is associated with adverse atrial remodelling and AF recurrences following catheter ablation.

Wang et al. proposed a novel biomarker of inflammation and oxidative stress, such as Monocyte-to-high-density lipoprotein ratio (MHR), as a predictor of the risk of AF among NAFLD patients. A retrospective cross-sectional analysis in a single-center sample was performed. MHR was significantly higher in patients with NAFLD with AF in comparison with NAFLD subjects without AF. MHR could be a simple and practical new inflammatory index used to assess the risk of AF in the clinical management of NAFLD patients.

Zhou et al. performed an updated systematic review and meta-analysis of cohort studies to define better the correlation between NAFLD/MAFLD and the likelihood of developing AF. Current updated evidence showed that NAFLD may be linked to a slightly higher risk of developing AF, particularly among Asian populations and those diagnosed with NAFLD using fatty liver index criteria. Authors highlighted the concept that further studies should consider factors such as specific population, the severity of NAFLD/MAFLD, diagnostic methods of NAFLD and AF, and cardiometabolic risk factors to determine better the association between NAFLD and the risk of AF.

Qiu et al. performed a meta-analysis to explore the association between NAFLD and the risk of adverse outcomes in patients with

HF. A total of six studies involving 12,374 patients with HF were included for analysis, with a median follow-up duration of 2.5 years. The pooled analysis showed that, after adjusting for multiple cardiovascular risk factors, HF patients with NAFLD were associated with a significantly increased risk of major adverse outcomes, all-cause mortality and HF hospitalization or re-hospitalization.

Finally, Jiang et al. investigated the associations of NAFLD and its advanced fibrosis with heart failure with preserved ejection fraction (HFpEF) according to obesity, glycated haemoglobin A1c (HbA1c), blood pressure (BP), and low-density lipoprotein cholesterol (LDL-C) goal achievement status in 2,418 hospitalized patients with type 2 diabetes. This cross-sectional analysis showed that in patients with type 2 diabetes, simple steatosis was not associated with HFpEF risk compared with patients without steatosis, and advanced hepatic fibrosis was significantly associated with an increased risk of HFpEF, regardless of obesity status, HbA1c, BP, and LDL-C goal achievement status.

Taken together, the studies published in this Research Topic provide the reader with an increased understanding of interactions between NAFLD and cardiac conduction, structure and function, opening new scenarios in the evaluation and treatment of subjects suffering from NAFLD. Further studies are needed to translate these observations into the real-world management of NAFLD patients.

Author contributions

CA: Writing – original draft, Conceptualization.

Conflict of interest

The author declares that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

Publisher's note

All claims expressed in this article are solely those of the authors and do not necessarily represent those of their affiliated organizations, or those of the publisher, the editors and the reviewers. Any product that may be evaluated in this article, or claim that may be made by its manufacturer, is not guaranteed or endorsed by the publisher.

References

1. Teng ML, Ng CH, Huang DQ, Chan KE, Tan DJ, Lim WH, et al. Global incidence and prevalence of non-alcoholic fatty liver disease. *Clin Mol Hepatol* (2023) 29:S32–42. doi: 10.3350/cmh.2022.0365
2. Radu F, Potcovaru C-G, Salmen T, Filip PV, Pop C, Fierbințeanu-Braticievici C. The link between NAFLD and metabolic syndrome. *Diagnostics* (2023) 13(4):614. doi: 10.3390/diagnostics13040614
3. Younossi ZM, Golabi P, Paik JM, Henry A, Van Dongen C, Henry L. The global epidemiology of non-alcoholic fatty liver disease (NAFLD) and non-alcoholic steatohepatitis (NASH): a systematic review. *Hepatology* (2023) 77:1335–47. doi: 10.1097/HEP.0000000000000004
4. Allen AM, Lazarus JV, Younossi ZM. Healthcare and socioeconomic costs of NAFLD: A global framework to navigate the uncertainties. *J Hepatol* (2023) 79:209–17. doi: 10.1016/j.jhep.2023.01.026
5. McSweeney L, Breckons M, Fattakhova G, Oluboyede Y, Vale L, Ternent L, et al. Health-related quality of life and patient-reported outcome measures in NASH-related cirrhosis. *JHEP Rep* (2020) 2:100099. doi: 10.1016/j.jhepr.2020.100099
6. Corrao S, Natoli G, Nobili A, Mannucci PM, Pietrangelo A, Perticone F, et al. RePoSI Investigators. Comorbidity does not mean clinical complexity: evidence from the RePoSI register. *Intern Emerg Med* (2020) 15:621–8. doi: 10.1007/s11739-019-02211-3
7. Riazi K, Azhari H, Charette JH, Underwood FE, King JA, Afshar EE, et al. The prevalence and incidence of NAFLD worldwide: a systematic review and meta-analysis. *Lancet Gastroenterol Hepatol* (2022) 7:851–61. doi: 10.1016/S2468-1253(22)00165-0
8. Byrne CD, Targher G. NAFLD: a multisystem disease. *J Hepatol* (2015) 62:S47–64. doi: 10.1016/j.jhep.2014.12.012
9. Chiu LS, Pedley A, Massaro J, Benjamin EJ, Mitchell GF, McManus DD, et al. The association of non-alcoholic fatty liver disease and cardiac structure and function – framingham heart study. *Liver Int* (2020) 40:2445–54. doi: 10.1111/liv.14600
10. Petta S, Argano C, Colomba D, Cammà C, Di Marco V, Cabibi D, et al. Epicardial fat, cardiac geometry and cardiac function in patients with non-alcoholic fatty liver disease: association with the severity of liver disease. *J Hepatol* (2015) 62:928–33. doi: 10.1016/j.jhep.2014.11.030
11. Sunbul M, Kivrak T, Durmus E, Akin H, Aydin Y, Ergelen R, et al. Non-alcoholic steatohepatitis score is an independent predictor of right ventricular dysfunction in patients with non-alcoholic fatty liver disease. *Cardiovasc Ther* (2015) 33:294–9. doi: 10.1111/1755-5922.12145
12. Mantovani A, Zoppini G, Targher G, Golia G, Bonora E. Non-alcoholic fatty liver disease is independently associated with left ventricular hypertrophy in hypertensive Type 2 diabetic individuals. *J Endocrinol Invest* (2012) 35:215–8. doi: 10.1007/BF03345421
13. Cai X, Zheng S, Liu Y, Zhang Y, Lu J, Huang Y. Non-alcoholic fatty liver disease is associated with increased risk of atrial fibrillation. *Liver Int* (2020) 40:1594–600. doi: 10.1111/liv.14461
14. Park HE, Lee H, Choi S-Y, Kim HS, Chung GE. The risk of atrial fibrillation in patients with non-alcoholic fatty liver disease and a high hepatic fibrosis index. *Sci Rep* (2020) 10:5023. doi: 10.1038/s41598-020-61750-4
15. Ziolkowska S, Binienda A, Jabłkowski M, Szemraj J, Czarny P. The interplay between insulin resistance, inflammation, oxidative stress, base excision repair and metabolic syndrome in non-alcoholic fatty liver disease. *Int J Mol Sci* (2021) 22:11128. doi: 10.3390/ijms222011128
16. Aroor AR, Mandavia CH, Sowers JR. Insulin resistance and heart failure: molecular mechanisms. *Heart Fail Clin* (2012) 8:609–17. doi: 10.1016/j.hfc.2012.06.005