



Strategies for a two-step liver fibrosis assessment in clinical practice

To the Editor:

We read with great interest the article by Mansour *et al.*¹ which proposes to incorporate a 2-step fibrosis assessment (Fibrosis-4 [FIB-4] score then vibration controlled transient elastography [VCTE]) into routine annual care of patients with type 2 diabetes (T2DM) in primary care clinics. The authors conclude that this strategy "... significantly improves the identification of advanced liver disease in patients with T2DM" when compared to current standard of care. Indeed, the systematic screening for non-alcoholic fatty liver disease (NAFLD) in patients with T2DM is recommended by the European Association for the Study of the

Liver (EASL)² and the American Diabetes Association (ADA)³ and the use of non-invasive scoring systems has been shown to be the simplest and most accurate strategy to identify patients at high risk of advanced fibrosis.⁴ In line with the authors, we are confident that this management approach has great potential to improve clinical care.

Primary care providers (PCPs) and diabetologists represent the most important link in the chain of management of patients with NAFLD since they are the first medical point of contact for this population. However, patients with T2DM are complex, with multiple comorbidities, and can be a significant burden on

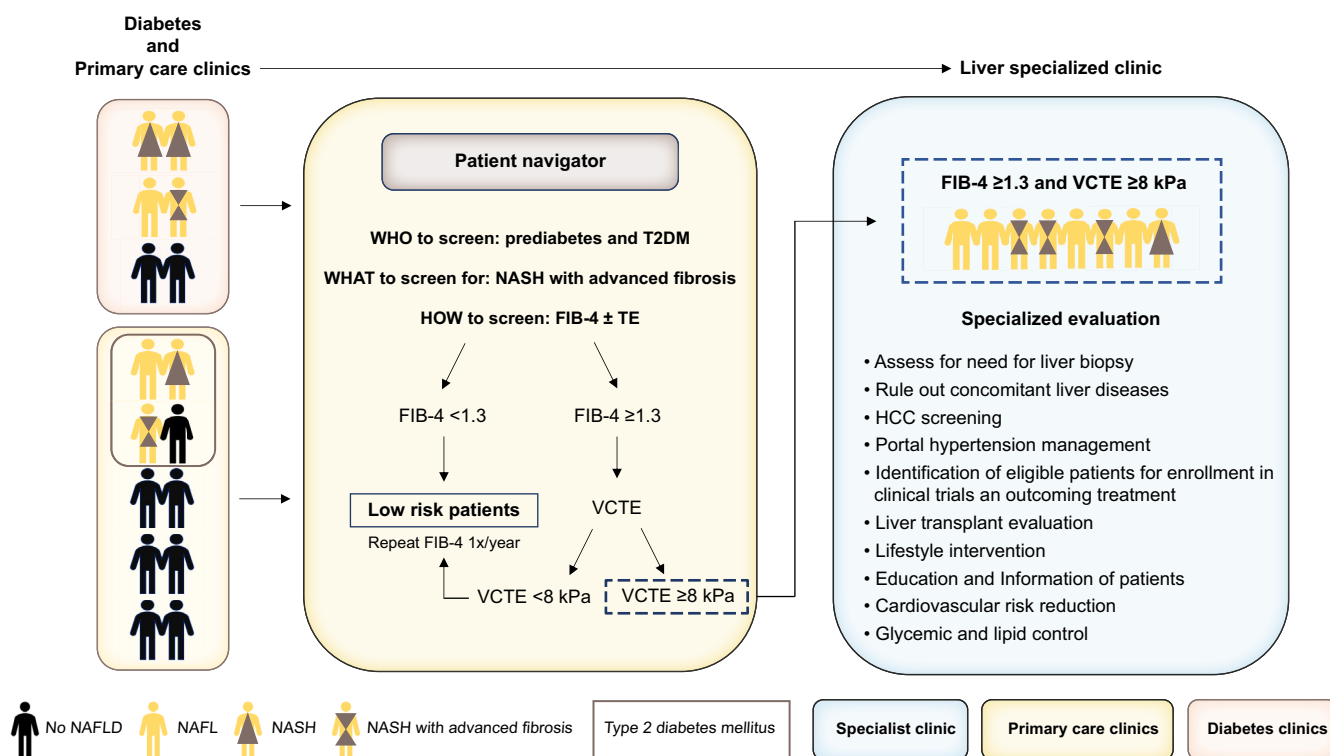


Fig. 1. Stepwise approach for screening, risk stratification and referral of NAFLD patients between primary care and diabetic clinics, and liver specialists.

The proposed algorithm involves a first step annual FIB-4 score (FIB-4) followed by VCTE for those with indeterminate or high-risk score (FIB-4 ≥ 1.3). Low risk patients (FIB-4 < 1.3 or VCTE < 8 kPa) can be followed-up by primary care providers for lifestyle changes and yearly calculation of FIB-4, while high risk patients (FIB-4 ≥ 1.3 and VCTE ≥ 8 kPa) should be referred to liver specialized clinics for further assessment and evaluation. NAFLD, non-alcoholic fatty liver disease; NAFL, non-alcoholic fatty liver; NASH, non-alcoholic steatohepatitis; FIB-4, Fibrosis-4 index; VCTE, vibration controlled transient elastography. Adapted from reference 7.

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primary care and diabetology clinics that are already overburdened.^{5,6} Hence, the optimal screening program should be simple and easily integrated into an existing workflow to improve long-term adherence and implementation.

We have recently published a workflow⁷ which proposes to incorporate the calculation of the FIB-4 into the already existing checklists used in the diabetic population. More specifically, we proposed the addition of platelet count to the checklist included in the 2020 guidelines published by the ADA – which included baseline and yearly transaminases – to allow the calculation of the FIB-4 and identify patients at high risk of advanced fibrosis. An indeterminate or high-risk score would then prompt additional evaluation with TE (Fig. 1).

Moreover, we propose the inclusion of a patient navigator into the care system of these patients. A patient navigator would co-manage the checklist with the PCP to increase adherence to screening for multiple complications of diabetes including non-alcoholic steatohepatitis (NASH), retinopathy, nephropathy, diabetic ulcers and ensure follow-up with subsequent testing and referral to specialists. Specifically in the screening for NASH, the patient navigator would have multiples

functions: i) flag patients who need labs for the calculation of FIB-4; ii) identify patients with indeterminate or high-risk FIB-4 scores (FIB-4 ≥ 1.3) who need referral to a specialized liver center and/or for VCTE; iii) follow-up to ensure that patients have undergone TE or had a specialist appointment. In our center, despite incorporating an automatic calculator into our electronic medical system, still roughly 40% of patients were missing data to calculate FIB-4, which highlights the key role of a patient navigator in prompting the PCP to order the labs needed. The inclusion of a patient navigator in the management of complex patients, such as patients with chronic liver diseases⁸ or diabetic patients⁹ has already been shown to be beneficial¹⁰ with an improvement of care, glycemic control, and better patient engagement.

In conclusion, we agree with the authors on the importance of incorporating a screening algorithm into the routine annual care of patients with T2DM in primary care clinics. We also believe that the inclusion of patient navigators in the complex management of patients with T2DM in primary care clinics and diabetic centers, as part of a multidisciplinary team, would greatly increase the utility of the screening algorithm.

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Conflict of interest

The authors declare no conflicts of interest that pertain to this work.

Please refer to the accompanying ICMJE disclosure forms for further details.

Authors' contributions

Conceptualization and draft of the manuscript: JVB. Critical revision of the manuscript for important intellectual content: JVB, ML.

Supplementary data

Supplementary data to this article can be found online at <https://doi.org/10.1016/j.jhepr.2021.100382>.

References

- [1] Mansour D, Grapes A, Herscovitz M, Cassidy P, Vernazza J, Broad A, et al. Embedding assessment of liver fibrosis into routine diabetic review in primary care. *JHEP Rep* 2021;3(4):100293. <https://doi.org/10.1016/j.jhepr.2021.100293>.
- [2] European Association for the Study of the Liver (EASL); European Association for the Study of Diabetes (EASD); European Association for the Study of Obesity (EASO). EASL-EASD-EASO Clinical Practice Guidelines for the management of non-alcoholic fatty liver disease. *J Hepatol* 2016;64:1388–1402. <https://doi.org/10.1016/j.jhepr.2015.11.004>.
- [3] American Diabetes Association. 4. Comprehensive medical evaluation and assessment of comorbidities: standards of medical care in diabetes-2020. *Diabetes Care* 2020;43:S37–S47. <https://doi.org/10.2337/dc20-S004>.
- [4] Armstrong MJ, Marchesini G. Referral pathways for NAFLD fibrosis in primary care - No longer a 'needle in a haystack. *J Hepatol* 2019;71(2):246–248. <https://doi.org/10.1016/j.jhepr.2019.05.010>.

- [5] Tai-Seale M, McGuire TG, Zhang W. Time allocation in primary care office visits. *Health Serv Res* 2007;42(5):1871–1894. <https://doi.org/10.1111/j.1475-6773.2006.00689.x>.
- [6] Altschuler J, Margolius D, Bodenheimer T, Grumbach K. Estimating a reasonable patient panel size for primary care physicians with team-based task delegation. *Ann Fam Med* 2012;10(5):396–400. <https://doi.org/10.1370/afm.1400>.
- [7] Vieira Barbosa J, Lai M. Nonalcoholic fatty liver disease screening in type 2 diabetes mellitus patients in the primary care setting. *Hepatol Commun* 2020;5(2):158–163. <https://doi.org/10.1002/hep4.1618>. eCollection 2021 Feb.
- [8] McBrien KA, Ivers N, Barnieh L, Bailey JJ, Lorenzetti DL, Nicholas D, et al. Patient navigators for people with chronic disease: a systematic review. *PLoS one* 2018;13(2):e0191980. <https://doi.org/10.1371/journal.pone.0191980>.
- [9] Horny M, Glover W, Gupte G, Saraswat A, Vimalananda V, Rosenzweig J. Patient navigation to improve diabetes outpatient care at a safety-net hospital: a retrospective cohort study. *BMC Health Serv Res* 2017;17(1):759. <https://doi.org/10.1186/s12913-017-2700-7>.
- [10] Hechter RC, Qian L, Luo Y, Ling Grant DS, Baxter R, Klein NP, et al. Impact of an electronic medical record reminder on hepatitis B vaccine initiation and completion rates among insured adults with diabetes mellitus. *Vaccine* 2019;37(1):195–201. <https://doi.org/10.1016/j.vaccine.2018.06.035>.

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