

Finally, we share the interest in exploring aspects, such as eating disorders, in this population, but that was beyond the scope of our study, which focused on comparing general intake profiles. Including multiple clinical, psychologic, and social dimensions in a single design may be ideal but is not always feasible or methodologically adequate. Our agreement as to the relevance of the relationship between eating disorders and IBS is demonstrated by the recent evaluation of said association by our working group.³ With respect to sample size, it was clearly a limitation of our study, as was the tool utilized and the age differences between groups, all of which we stated at the end of the manuscript.

In conclusion, we appreciate all constructive criticism, but we emphasize the importance of making pertinent observations, based on a thorough reading of the manuscript and with relevant references. Opinions or generalizations that lack robust evidence must be clearly distinguished from rigorous scientific analysis.

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Serum 7 α C4 as a pragmatic diagnostic tool for bile acid malabsorption in Mexico: Implications and future directions



El 7 α C4 sérico como herramienta diagnóstica práctica para la mala absorción de ácido biliar en México: implicaciones y direcciones futuras

Dear Editors,

We read with great interest the recent article by Mendoza-Domínguez et al.¹ titled “Real-world experience with the diagnosis of bile acid malabsorption (BAM) using serum 7-alpha-C4 and 48-h stool bile acids.” Their work offers crucial insights into diagnostic alternatives for BAM in Mexico, where 75SeHCAT testing remains unavailable.

The authors demonstrate that serum 7 α C4 levels correlate moderately with total fecal bile acids (TBA) and can identify BAM even when primary (PBA) and TBA levels are normal. This is clinically significant, as it suggests serum 7 α C4 is a reliable and cost-saving biomarker, with their data indicating an approximate 49% reduction in diagnostic costs compared with combined stool and serum testing.

However, we wish to highlight three key considerations for clinical translation and future research.

First, although serum 7 α C4 offers logistical and economic advantages over 48-h stool collection, its current utility remains limited by the necessity of shipping samples abroad for analysis, leading to delays and higher systemic costs. As the authors note, establishing local tandem mass spectrometry facilities for 7 α C4 quantification would accelerate diagnosis and enhance equitable access for public sector patients. Indeed, Camilleri et al.² previously emphasized that widespread clinical use of serum 7 α C4 requires standardized protocols and availability of high-throughput assays, which remain limited in low- and middle-income settings.

Second, the low prevalence of BAM among patients with functional diarrhea or IBS-D in their cohort contrasts with meta-analytic data indicating a prevalence of ~30% in IBS-D³ and up to 63.5% following cholecystectomy.⁴ This discrepancy may reflect underdiagnosis due to limited clinician awareness or referral bias toward patients with structural causes of diarrhea. Recent guidelines recommend consideration of BAM in all cases of chronic unexplained diarrhea, given its responsiveness to bile acid sequestrants.⁵

Third, while cholestyramine remains the primary treatment option in Mexico, tolerability issues and poor palatability often lead to non-adherence. Emerging therapies such as colesevelam, which have demonstrated better gastrointestinal tolerability profiles, warrant evaluation in Mexican populations.

Furthermore, while Mendoza-Domínguez et al.¹ found serum 7 α C4 to be elevated in patients with BAM despite normal fecal BA levels, combining serum 7 α C4 with single stool bile acid measurements may offer a pragmatic diagnostic approach to avoid the logistical difficulties of 48-h stool collection while maintaining diagnostic accuracy.

In sum, this timely study by Mendoza-Domínguez et al.¹ advances our understanding of BAM diagnosis in Mexico and underscores the need to integrate serum 7 α C4 testing into routine clinical practice, supported by local laboratory capacity and clinician education. Further research should establish national reference values for serum 7 α C4, validate combined serum-stool diagnostic algorithms, and evaluate newer sequestrants to optimize treatment outcomes.

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Declaration of competing interest

The authors declare there is no conflict of interest.

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Response to Rattapitoot et al., “Serum 7 α C4 as a pragmatic diagnostic tool for bile acid malabsorption in Mexico: Implications and future directions”



Respuesta a Rattapitoot et al., “El 7 α C4 sérico como herramienta diagnóstica pragmática para la malabsorción de ácidos bilíares en México: implicaciones y perspectivas futuras”

To the Editors,

We are grateful to Dr. Rattapitoot et al. for their thoughtful letter and interest in our study: “Real-world experience with the diagnosis of bile acid malabsorption (BAM) using serum 7-alpha-C4 and 48-h stool bile acids”.¹ Their remarks highlight several important considerations for the clinical

implementation and future research of BAM diagnosis in patients with functional diarrhea in Mexico.

First, we fully agree with the need to standardize serum 7 α C4 quantification at the local level to overcome the logistical and economic limitations associated with sending samples abroad. As noted by Camilleri and Vijayvargiyah, the clinical utility of this biomarker depends not only on its diagnostic performance but also on the availability of standardized, high-throughput mass spectrometry assays, which remains a challenge in many middle or low-income countries.²

Second, while our study reported a lower prevalence of BAM among patients with functional diarrhea compared with prior reports,^{3,4} we coincide that BAM must be actively investigated in all patients with chronic diarrhea of unknown origin. However, the cost of this approach remains a limitation in our setting. As outlined in our recently presented abstract during the 2025 Digestive Disease Week (DDW), we support a broader diagnostic strategy that includes evaluating other potential etiologies of chronic diarrhea, such as accelerated intestinal transit and post-infectious disorders,