

Title:

T3-T4 index as a prognostic marker of response to biological treatments in elderly patients with inflammatory bowel disease

Authors:

María Martínez Burgos, Raúl Vicente Olmedo Martín, Víctor Amo Trillo, Patricia Romero Cara, José Mostazo Torres, Miguel Jiménez Pérez

DOI: 10.17235/reed.2024.10445/2024 Link: <u>PubMed (Epub ahead of print)</u>

Please cite this article as:

Martínez Burgos María, Olmedo Martín Raúl Vicente, Amo Trillo Víctor, Romero Cara Patricia, Mostazo Torres José, Jiménez Pérez Miguel. T3-T4 index as a prognostic marker of response to biological treatments in elderly patients with inflammatory bowel disease. Rev Esp Enferm Dig 2024. doi: 10.17235/reed.2024.10445/2024.

This is a PDF file of an unedited manuscript that has been accepted for publication. As a service to our customers we are providing this early version of the manuscript. The manuscript will undergo copyediting, typesetting, and review of the resulting proof before it is published in its final form. Please note that during the production process errors may be discovered which could affect the content, and all legal disclaimers that apply to the journal pertain.

Title: T3-T4 index as a prognostic marker of response to biological treatments in elderly

patients with inflammatory bowel disease.

Authors: Martínez Burgos, María²; Olmedo Martín, Raúl Vicente¹; Amo Trillo, Victor¹;

Romero Cara, Patricia¹; Mostazo Torres, Jose¹; Jimenez Pérez, Miguel¹.

1: Hospital Regional Universitario de Málaga (Málaga)

2: Hospital Universitario Santa Lucía de Cartagena (Murcia)

Corresponding author email: mariamarbur@gmail.com (María Martínez Burgos)

Mr Editor:

Following the current trend in inflammatory bowel disease (IBD) of trying to individualise

treatment (1) and based on the work of Lorenzo Bertani (2) we have conducted a study

with the aim of reproducing that the T3-T4 index is a useful serological marker for

predicting response to biologic treatments in elderly patients with IBD.

We included all patients in the practice who were starting biological treatment for the

first time and were over 60 years of age, regardless of their comorbidity. Assessment of

response to biologic treatments was performed taking into account real clinical practice

parameters: steroid-free clinical remission (defined as Harvey Bradshaw Index (HBI) <4

in Crohn's disease or Clinical Activity Index <2 in ulcerative colitis) and biological

remission (defined as faecal calprotectin <250 µg/g and CRP <8 mg/dl). Measurement of

all these parameters was performed 12 weeks after treatment initiation.

A total of 68 patients were collected, 33 men and 35 women. The mean age was 68.47

± 6.28 years. The diagnosis was ulcerative colitis (UC) in 29.4 % and Crohn's disease (CD)

in 70.6 %. When analysing the mean T3/T4 index values between the steroid-free clinical

remission/no steroid-free clinical remission and biological remission/no biological

remission groups, no significant differences were found (Figure 1).

Discussion

There have been multiple observational clinical studies in which, in the absence of thyroid pathology, low T3 levels, or increased T4 relative to T3 (known as "low T3 syndrome"). These low levels have been associated with poor prognosis in various therapeutic scenarios and have even been proposed as a marker of frailty in the elderly patient. (3-5).

Recently, multiple studies on elderly IBD patients have been carried out (6-8), as this patient profile with comorbidities is becoming more and more frequent, and given its special complexity, the development of serological parameters such as the one proposed by Bertani could be very useful. With this letter we would like to encourage other groups to share their experience with this marker, as has been done in the past with HLA-DQA1*05 (5).

Regarding the uniqueness of the elderly patient, a tendency to low levels of activated thyroid hormone has been observed. Eighty percent of the conversion of thyroxine (T4) to the active hormone triiodothyronine (T3) is performed by deiodinases peripherally in the liver and kidney. Some studies suggest that chronic inflammatory states, low body mass index (BMI) and sarcopenia may be associated with increased protein catabolism and thus lower levels of activated thyroid hormone (9).

Regarding our work, it's undeniable that difference in the measurement technique of the thyroid hormone used in the Italian hospital compared to the one used in our centre makes it difficult to compare results, but this is a factor over which we cannot intervene.

Another limiting factor of the study could be that by measuring the 12-week response we might miss late responses compared to the original study, as well as the small sample size.

Given the extensive literature that supports the justification of the work and despite its negative results, with this communication we intend to stimulate the curiosity of colleagues on this topic and encourage them to share their results.

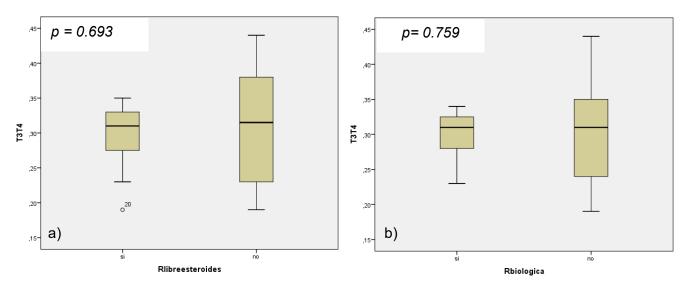


Figure 1. Box and whisker plots show that there are no statistically significant differences between group means according to T3/T4 index levels. a) Steroid-free remission, b) Biological remission.

Bibliography

- 1. Olmedo RV. Is individualization of biological treatment possible in inflammatory bowel disease? Revista Andaluza Patología Digestiva. Vol 45 Nº. 1, 2022, págs. 14-19
- 2. Bertani L, Tricòn D, Pugliese De et al. Serum triiodothyronine-to-thyroxine (T3/T4) ratio predicts therapeutic outcome to biological therapies in elderly IBD patients. Aliment Pharmacol Ther. 2021;53:273–280
- 3 -Mebis L, Van den Berghe G. Thyroid axis function and dysfunction in critical illness. Best Pract Res Clin Endocrinol Metab. 2011;25:745-757
- 4- Moura Neto A, Zantut-Wittmann DE. Abnormalities of thyroid hormone metabolism during systemic illness: the low t3 syndrome in different clinical settings. Int J Endocrinol. 2016;2016:2157583.
- 5-Pasqualetti G, Calsolaro V, Bernardini S, et al. Degree of peripheral thyroxin deiodination, frailty, and long-term survival in hospitalized older patients. J Clin Endocrinol Metabol. 2018;103:1867-1876

- 6. Adar T, Faleck D, Sasidharan S, et al. Comparative safety and effectiveness of tumor necrosis factor alpha antagonists and vedolizumab in elderly IBD patients: a multicentre study. Aliment Pharmacol Ther. 2019;49:873-879.
- 7. Ananthakrishnan AN, Donaldson T, Lasch K, et al. Management of inflammatory bowel disease in the elderly patient: challenges and opportunities. Inflamm Bowel Dis. 2017;23:882-893.
- 8. Shimoda F, Naito T, Kakuta Y, Kawai Y, Tokunaga K; NCBN Controls WGS Consortium; Shimoyama Y, Moroi R, Shiga H, Nagasaki M, Kinouchi Y, Masamune A. HLA-DQA1*05 and upstream variants of PPARGC1B are associated with infliximab persistence in Japanese Crohn's disease patients. Pharmacogenomics J. 2023 Nov;23(6):141-148. doi: 10.1038/s41397-023-00312-z. Epub 2023 Jul 17. PMID: 37460671.
- 9 -Adams DW, Gurwara S, Silver HJ, et al. Sarcopenia is common in overweight patients with inflammatory bowel disease and may predict need for surgery. Inflamm Bowel Dis. 2017;23:1182-1186