



EVALUATION OF LIVER FIBROSIS USING TRANSIENT ELASTOGRAPHY IN PATIENTS WITH PSORIASIS TREATED WITH METHOTREXATE

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BACKGROUND

Methotrexate is a widely used drug in the treatment of moderate to severe psoriasis and psoriatic arthritis. One of the most worrying adverse effects is hepatotoxicity, due to the possibility of inducing liver cirrhosis. There are some options for its monitoring, with liver biopsy (LB) being the gold standard for detecting liver fibrosis. Some consensuses indicate it routinely when the cumulative dose (CD) of 3.5 to 4 g is reached. Needle LB is an invasive method and has technical limitations. Therefore, transient elastography (TE) is currently being increasingly used to estimate the degree of liver fibrosis in a non-invasive way, with reliable results.

OBJECTIVES

To evaluate the presence of liver fibrosis using TE in patients with psoriasis treated with methotrexate; and to correlate the presence of changes in liver enzymes and the body mass index (BMI) value according to the TE result.

METHODS

Cross-sectional observational study, approved by the research ethics committee. From July 2018 to August 2019, patients with psoriasis, over 18 years old, of both genders, treated with methotrexate or topical therapy, who are not using immunobiologicals, were selected. All were referred for TE, at a single moment during the study, using Fibrosan®, with the Metavir score estimating the degree of liver fibrosis, ranging from F0 to F4. A result \geq F2 would be compatible with significant fibrosis. The results of liver biochemistry were also evaluated and the BMI was calculated. TE was performed in 90 patients: 35 using methotrexate with CD < 3.5 g; 28 using methotrexate with CD > 3.5 g; and 27 that weren't in systemic treatment and with a BMI greater \geq 25 kg/m², as a control group. The chi-square test (χ^2) was conducted to determine the significance between the groups.

RESULTS

There wasn't statistically significance between the CD of methotrexate and the the Metavir score ($\chi^2=2.57$; $p=0.86$), nor between the presence of changes in liver enzymes and the Metavir score ($\chi^2=5.04$; $p=0.168$). However, there was statistically significance between BMI and Metavir score ($\chi^2=16.8$; $p=0.01$).

CONCLUSION

We must take into consideration the presence of obesity as a risk factor for liver fibrosis, which is probably more important than CD of methotrexate. However, further research is needed so that we can better assess the risk factors related to the use of methotrexate.

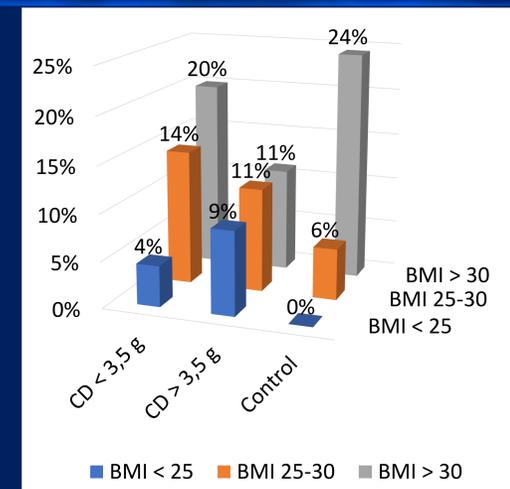


Figure 1: BMI

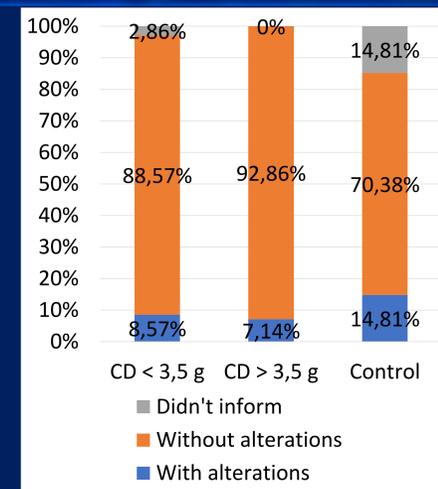


Figure 2: Liver enzymes

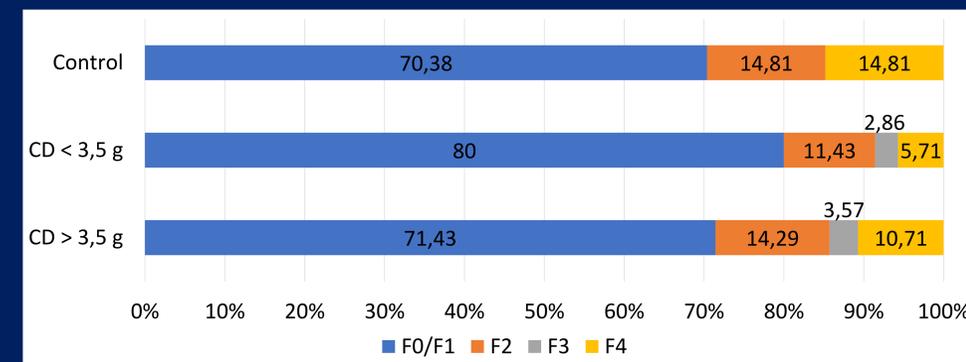


Figure 3: Metavir score

PARAMETERS	CD < 3,5 g	CD > 3,5 g	CONTROL	TOTAL
Patients (%)	35 (39%)	28 (31%)	27 (30%)	90 (100%)
Age (years)	56,52 ±11,8	56,43 ±11,56	56,87 ±11,21	56,44 ±11,77
Gender female / male (%)	22 (62,9%) / 13 (37,1%)	12 (42,9%) / 16 (57,1%)	13 (48,1%) / 14 (51,9%)	47 (52%) / 43 (48%)
CD (mg)	475 - 3446	3586 - 7231	0	475-7231
BMI (Kg/m ²)	30,97 ±5,66	30,95 ±5,67	31,21 ±5,66	31,11 ±5,61

Table 1: Baseline characteristics

REFERENCES

- Barker J, et al. Assessment and management of methotrexate hepatotoxicity in psoriasis patients: report from a consensus conference to evaluate current practice and identify key questions toward optimizing methotrexate use in the clinic. *J Eur Acad Dermatol Venereol.* 2011; 25:758–64.
- Thomas JA, Aithal GP. Monitoring liver function during methotrexate therapy for psoriasis: are routine biopsies really necessary? *Am J Clin Dermatol.* 2005; 6: 357–363
- Montaudie H, et al. Methotrexate in psoriasis: a systematic review of treatment modalities, incidence, risk factors and monitoring of liver toxicity. *J Eur Acad Dermatol Venereol.* 2011;25 Suppl 2:12–18.
- Poynard T, et al. Liver biopsy analysis has a low level of performance for diagnosis of intermediate stages of fibrosis. *Clin Gastroenterol Hepatol.* 2012; 10(6): 657-63.e7
- Friedrich-Rust M, et al. Comparison of ELF, FibroTest and FibroScan for the Non-Invasive Assessment of Liver Fibrosis. *BMC Gastroenterol.* 2010; 10:103.
- Berends MA, et al. Biochemical and biophysical assessment of MTX-induced liver fibrosis in psoriasis patients: Fibrotest predicts presence and Fibrosan predicts the absence of significant liver fibrosis. *Liver Int.* 2007; 27: 639-45.
- Bath RK, et al. A review of methotrexate-associated hepatotoxicity. *J Dig Dis.* 2014;15:517–524.
- Berends MA, et al. Liver injury in long-term methotrexate treatment in psoriasis is relatively infrequent. *Aliment Pharmacol Ther.* 2006;24(5):805-811.
- Lynch M, et al. The Use of Transient Elastography and FibroTest for Monitoring Hepatotoxicity in Patients Receiving Methotrexate for Psoriasis. *JAMA Dermatol.* 2014; 150: 856-62.