GUIDELINES FOR JUVENILE IDIOPATHIC ARTHRITIS MANAGEMENT: IS THERE A ROOM FOR COMBINED METHOTREXATE AND LEFLUNOMIDE THERAPY IN THE TREATMENT RECOMMENDATIONS

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Methods: The treatment guidelines were developed based on systematic review, local studies, formal consensus and feedback. Initiation of methotrexate or leflunomide was recommended for polyarticular JIA children who, after 3 month of treatment, did not respond to methotrexate or leflunomide monotherapy, or those whose DMARD dose could not be optimised; a new management step was introduced where a combination of both medications was administered. This was a multicentre study including 76 JIA patients who had been treated with the combination of methotrexate plus leflunomide. All patients were classified according to the International League of Associations for Rheumatology (ILAR) criteria. Recorded data included: demographics, JIA subtype, reason for starting combined treatment, treatment duration, withdrawals, causes of discontinuation, efficacy and safety. Patients were classified as “responders” or “non-responders.”

Results: Of the 76 children there were, oligoarthritis (34%), polyarthritis (31%), systemic JIA with spondylitis (>4 joints) (20%), and psoriatic arthritis (15%). Mean age at initiation of combined therapy 10.2±3.4 years, mean disease duration 9.4±6.8 months. The combined therapy was superior to methotrexate alone and did not significantly increase the rate of adverse events. ACR-Ped 30 was achieved in 64% at 3 months, 75% at 6 months. This was superior to methotrexate alone (37.3% and 53.8%; p<0.01 at 3- and 6-months respectively). At 1 year, 81% reached Ped 30, 74.5% reached ACR-Ped 50, 64% achieved ACR-Ped 70 whereas 51% met ACR-Ped 90 criteria. There were no serious adverse events. One of the two DMARDs was stopped in 51% of the children; of them: 25% were due to adverse events, clinical remission in 25% whereas 21% was switched to anti-TNF therapy according to guidelines due to ineffectiveness or loss of efficacy. All patients who had had uveitis responded well and achieved clinical remission.

Conclusions: For the children with polyarticular JIA who did not respond to monotherapy with methotrexate, combination of methotrexate and leflunomide treatment appeared to be efficacious and maintain a durable response. The current recommendation would be to use combined methotrexate and leflunomide in children with polyarticular JIA who are either intolerant to higher methotrexate doses or who did not have a satisfactory response to methotrexate. The combination should be considered prior to the use of a biologic agent.

Disclosure of Interest: None declared

IMPACT OF METHOTREXATE ON GROWTH IN CHILDREN WITH JUVENILE IDIOPATHIC ARTHRITIS

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Background: Juvenile idiopathic arthritis (JIA) is the most prevalent paediatric rheumatic disease. Long term complications include physical disability and a decreased quality of life. Since the introduction of anti-TNF drugs for JIA, its prognosis has improved significantly. Personalised medicine is the next step to improve treatment in JIA. Anti-TNF trough levels and demonstration of the presence of anti-drug antibodies (ADA) could help individualise treatment decisions in JIA patients, but evidence supporting this is missing.

Objectives: The objective of this study was to evaluate the effect of MTX on growth parameters in pre-pubertal children with JIA and to determine the factors affecting the growth velocity.

Methods: We assessed height and changes in the height standard deviation score (SDS) at disease onset, at the onset of MTX and at the last follow-up visit in a cross-sectional study of JIA children. All patients were pre-pubertal when MTX began and were followed for at least 6 months afterward. We compared growth parameters (height, growth rate, weight and body mass index (BMI)) in responders and non-responders to MTX. The growth rate was defined as the number of millimetres of height acquired during 1 year. Associations between changes in the height SDS and discrete variables were evaluated using chi-square or Fisher’s exact tests. The significance level was set at 0.05.

Results: We enrolled 36 pre-pubertal children with JIA (24 boys and 12 girls) who had been treated with MTX orally. Median patient age was 6.2 years at the onset of MTX and 8.4 years at the latest follow-up. The median disease duration was 2.7 years. Twenty-one patients (58.3%) had oligoarticular JIA, 2 patients (5.5%) had systemic JIA, 10 (27.7%) had polyarticular JIA and 3 (8.3%) had enthesitis-related arthritis. Nineteen patients (52.7%) had received corticosteroids during an average period of 1.7 years with a mean of 10 mg/day of prednisone or equivalent. The median duration of MTX at the latest follow-up was 3.1 years with a mean MTX dose of 10 mg/m2/week.

Twelve-eight patients responded to MTX treatment and 8 did not. There were no significant differences between the responders and non-responders for age at treatment initiation, disease duration and MTX dose. At MTX onset, no significant differences between the two groups in terms of height (p=0.52), growth rate (p=0.08), weight (p=0.74) and BMI (p=0.9) were found. One year after MTX therapy, mean height (0.2 versus −1.1; p=0.03), mean growth rate (0.5 versus −2.9 SDS; p<0.01), mean weight (0.4 versus −2.3 SDS; p<0.01) and mean BMI (0.6 versus −1.9; p=0.04) were significantly higher in the responder group than in non-responders, respectively. At the latest follow-up, this increase was significantly maintained for growth rate (p=0.001) and height (p=0.002) in the responder group. In the multivariate analysis, patients who required more than 10mg/m2/week of MTX, systemic JIA and patients with reliance on steroids had a significantly lower growth velocity after the onset of MTX (p<0.01, p=0.02, p=0.02 respectively).

Conclusions: In our study, the increase in growth parameters in pre-pubertal children with JIA was associated with a better control of the disease activity under MTX therapy.

Disclosure of Interest: None declared

THERAPEUTIC DRUG MONITORING OF BIOLOGICALS IN CHILDREN WITH JUVENILE IDIOPATHIC ARTHRITIS (JIA): AN OVERVIEW OF CURRENT PRACTICE IN ANTI-TNF THERAPY

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Background: Juvenile idiopathic arthritis (JIA) is the most prevalent paediatric rheumatic disease. Long term complications include physical disability and a decreased quality of life. Since the introduction of anti-TNF drugs for JIA, its prognosis has improved significantly. Personalised medicine is the next step to improve treatment in JIA. Anti-TNF trough levels and demonstration of the presence of anti-drug antibodies (ADA) could help individualise treatment decisions in JIA patients, but evidence supporting this is missing.