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## Methotrexate plus Medical Ozone combined therapy in Rheumatoid Arthritis. From experimental models to clinical trials [abstract]

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### ABSTRACT

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**Introduction.** The aim of this work has been to investigate the medical ozone effects on 2 experimental models: PG/PS induced RA and synovitis followed for a clinical trial of efficacy.

**Methods.** mRNA levels of TNF-alpha and IL-1 beta, injury and protective biomarker of oxidative stress, morphology studies, A1 adenosine receptor role, Anti-Cyclic Citrullinated (Anti-CCP) levels and clinical variables were determined.

**Results.** Medical ozone ameliorated the expression of proinflammatory cytokines, reduced oxidative damage while participation of A1 adenosine receptor in the antiinflammatory effects was demonstrated.

In RA patients (n=60) MTX + ozone reduced the activity of the disease while MTX merely showed a tendency to decrease it. MTX + ozone reduced Anti-CCP levels as well as increased antioxidant system, and decreased oxidative damage whereas MTX did not change.

**Conclusion.** Medical ozone was effective in the reduction of inflammatory response in 2 experimental studies. MTX + ozone increased the MTX clinical efficacy in RA patients. These results suggest that ozone can increase the efficacy of MTX probably because both share common therapeutic targets.