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Hypothesis generating study to identify the changes in synovial tissue early after initiation of infliximab therapy

Submission date
23/08/2007

Recruitment status
No longer recruiting

Registration date
23/08/2007

Overall study status
Completed

Last Edited
07/10/2021

Condition category
Musculoskeletal Diseases

[ ] Prospectively registered
[ ] Protocol
[ ] Statistical analysis plan
[X] Results
[ ] Individual participant data

Plain English summary of protocol
Not provided at time of registration

Contact information

Type(s)
Scientific

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Additional identifiers

EudraCT/CTIS number

IRAS number

ClinicalTrials.gov number

Secondary identifying numbers

N/A

Study information

Scientific Title
Hypothesis generating study to identify the changes in synovial tissue early after initiation of infliximab therapy

Study objectives
Exploratory study to investigate the effects of Tumour Necrotising Factor (TNF) targeted therapy with infliximab on the synovial cell infiltrate, and the induction of apoptosis.

Ethics approval required
Old ethics approval format

Ethics approval(s)
Ethics approval received from the Medical ethical committee of the Academic Medical Center /University of Amsterdam on the 1st October 2003 (ref: MEC 01/086).

Study design
Non-randomised, non-controlled, exploratory study

Primary study design
Interventional

Secondary study design
Non randomised controlled trial

Study setting(s)
Hospital

Study type(s)
Treatment

Participant information sheet

Health condition(s) or problem(s) studied
Rheumatoid arthritis

Interventions
Infliximab therapy (3 mg/kg intravenous [i.v.]) according to the normal regimen. At baseline and 1 hour (n = 5) or 24 hours (n = 5) after the first infliximab infusion synovial biopsies were obtained from an inflamed knee joint. Peripheral blood mononuclear cells were obtained before and 1 and 24 hours after infliximab infusion in 20 patients (10 only blood, 10 with paired synovial biopsies). Serum was drawn at similar timepoints.

Intervention Type
Drug

Phase

Not Specified

Drug/device/biological/vaccine name(s)
Infliximab

Primary outcome measure
1. Primary immunohistological outcome: detection of apoptosis in synovial tissue within 1 or 24 hours after initiation of treatment. Analysis by immunohistochemical staining and electron microscopy
2. Primary serological outcome: To determine whether TNF targeted therapy with infliximab results in apoptosis of peripheral blood mononuclear cells within 1 or 24 hours after initiation of treatment

Secondary outcome measures
To determine whether TNF targeted therapy with infliximab results in decreased synovial cellularity.

Overall study start date
01/10/2003

Completion date
01/09/2006

Eligibility

Key inclusion criteria
1. Rheumatoid Arthritis (RA) patients with active disease at baseline assessed by the Disease Activity Score (DAS-28)
2. Be more than or equal to 18 years of age
3. Use concurrent methotrexate treatment (7.5 - 30 mg/week; stable since greater than or equal to 28 days before initiation) during the study. Subjects may be taking non-steroidal anti-inflammatory drugs, provided the dose and frequency have been stable for at least 28 days. Subjects may be receiving prednisone therapy less than or equal to 10 mg/day provided that the dosage has been stable for at least a months prior to entry

Participant type(s)
Patient

Age group
Adult

Lower age limit
18 Years

Sex
Not Specified

Target number of participants
20

Key exclusion criteria
1. Pregnancy
2. Breastfeeding
3. A history of or acute inflammatory joint disease of different origin, e.g., mixed connective tissue disease, seronegative spondyloarthropathy, psoriatic arthritis, Reiter's syndrome, systemic lupus erythematosus or any arthritis with onset prior to age 16 years
4. Acute major trauma
5. Previous therapy at any time with:
5.1. TNF-directed monoclonal antibodies
5.2. p75 TNF receptor fusion protein
6. Therapy within the previous 45 days with:
6.1. Any experimental drug
6.2. Alkylating agents, e.g. cyclophosphamide, chlorambucil
6.3. Anti-metabolites
6.4. Monoclonal antibodies
6.5. Growth factors
6.6. Other cytokines
7. Therapy within the previous 28 days with:
7.1. Parenteral or intraarticular corticoid injections
7.2. Oral corticosteroid therapy exceeding a prednisone equivalent of 10 mg daily
7.3. Present use of Disease Modifying Anti-Rheumatic Drugs (DMARDs) other than methotrexate
8. Fever (orally measured greater than 38°C), chronic infections or infections requiring anti-microbial therapy
9. Manifest cardiac failure (stage III or IV according to New York Heart Association [NYHA] classification)
10. Progressive fatal disease/terminal illness
11. A hematopoietic disease
12. Body weight of less than 45 kg

Date of first enrolment
01/10/2003

Date of final enrolment
01/09/2006

Locations

Countries of recruitment
Netherlands

Study participating centre
Academic Medical Centre (AMC)
Amsterdam
Netherlands
1100 DD

Sponsor information

Organisation
Academic Medical Centre (AMC) (The Netherlands)

Sponsor details
Division of Clinical Immunology and Rheumatology
PO Box 22660
Amsterdam
Netherlands
1100 DD

Sponsor type
Hospital/treatment centre

Website
http://www.amc.uva.nl/

ROR
https://ror.org/03t4gr691

Funder(s)

Funder type
Industry

Funder Name
Centocor Inc. (USA)

Results and Publications

Publication and dissemination plan
Not provided at time of registration

Intention to publish date

Individual participant data (IPD) sharing plan

IPD sharing plan summary
Not provided at time of registration

Study outputs

Output type	Details	Date created	Date added	Peer reviewed?	Patient-facing?
Results article		30/10/2008	07/10/2021	Yes	No