

The lifespan of human neutrophils, eosinophils, basophils and monocytes under homeostatic conditions

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Ethical review	Approved WMO
Status	Recruitment stopped
Health condition type	Other condition
Study type	Observational invasive

Summary

ID

NL-OMON34718

Source

ToetsingOnline

Brief title

Leukocyte lifespan

Condition

- Other condition
- Allergic conditions

Synonym

Chronic immune diseases and basic knowledge on the human immune system

Health condition

fundamentele kennis mbt normale afweer

Research involving

Human

Sponsors and support

Primary sponsor: Universitair Medisch Centrum Utrecht

Source(s) of monetary or material Support: het Nederlands Astma Fonds

Intervention

Keyword: deuterated glucose, innate immunity, leukocyte, lifespan

Outcome measures

Primary outcome

Granulocyte and monocyte post-mitotic pool transit times and lifespans in normal healthy volunteers

Secondary outcome

None

Study description

Background summary

Remarkably little is known about the lifecycle of the cells of the innate immune system: monocytes, neutrophils, eosinophils and basophils. Knowledge about this lifecycle is important for fundamental insights in the immune system and also for the understanding of pathologies of inflammatory diseases. An important difficulty is the development of new medications is is the lack of knowledge on the basal characteristics of the cells that cause chronic inflammation: how fast they are produced, how long they remain in the blood and tissues and where they are cleared. Little is described in literature regarding these topics. In the 60's and 70's studies were performed on the lifespans of innate immune cells, but with inadequate techniques. These data probably underestimate the lifespans of these cells, but they are still mentioned in modern tekst books.

Study objective

The results of this study will lead to an improved insight in the lifecycle of these cells and our results will be of importance for more insight in chronic inflammatory diseases. For example, it will improve our ability to interpret the results from previous and future intervention studies that block survival

and production of leukocytes.

Study design

On day one the volunteers will go to the clinic on an empty stomach. First we will withdraw 20ml of blood for baseline measurements of glucose levels and DNA deuterium enrichment.

After that, the volunteers will be orally administered 1g of deuterated glucose per kilogram bodyweight in 12 doses over a period of 6 hours. Also, after 1, 3 and 6 hours after the first administration we will withdraw two drops of blood by skinprick to determine the amount of deuterated glucose in the blood of the volunteer.

During this day, the volunteer will receive low-carb breakfast and lunch.

At 5 more timepoints, the volunteer will come to the clinic to donate 20ml of blood. The exact days after intake of glucose differ for each volunteer but will not be in weekends or more than two days in a row. (A clear scheme of the withdrawals can be found in the "onderzoeksprotocol", paragraph 3.2.)

From the collected blood DNA will be isolated and white blood cell populations will be separated using high performance FACS sorting. DNA from these cells will be isolated and analysed for deuterium enrichment using a combination of gas chromatography and mass-spectrometry.

These data will be fed to a mathematical model, which can calculate the half-lives of the cells.

Study burden and risks

Risks:

The glucose used is considered safe, so risks are negligible

Burden:

Subject will pay five visits to the clinic for blood withdrawal. Each visit, 20ml of blood will be withdrawn, which can be easily missed by adults.

Besides, subjects will spend one day in hospital for one blood withdrawal and the intake of 1g/kg bodyweight of glucose

Taken together, we think that the risks for volunteers are minimal and the burden is low.

Contacts

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Trial sites

Listed location countries

Netherlands

Eligibility criteria

Age

Adults (18-64 years)

Elderly (65 years and older)

Inclusion criteria

18-40 years

Exclusion criteria

- Any infection (eg. HIV, Hepatitis, STDs)
- Smoking
- Asthma or COPD
- Hay fever or allergies
- Auto-immune diseases
- Use of medication, excluding: contraceptives and pain killers (if used less than once a week)
- exuberant alcohol consumption (for males > 36 glasses per week, for females >24 glasses per week)
- Drug use
- History of cancer

Study design

Design

Study type: Observational invasive

Masking: Open (masking not used)

Control: Uncontrolled

Primary purpose: Basic science

Recruitment

NL

Recruitment status: Recruitment stopped

Start date (anticipated): 30-11-2011

Enrollment: 6

Type: Actual

Ethics review

Approved WMO

Date: 20-05-2011

Application type: First submission

Review commission: METC Universitair Medisch Centrum Utrecht (Utrecht)

Study registrations

Followed up by the following (possibly more current) registration

No registrations found.

Other (possibly less up-to-date) registrations in this register

No registrations found.

In other registers

Register	ID
CCMO	NL31552.041.10