

SaDial: the adaptive immune response against *Staphylococcus aureus* in hemoDialysis patients

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| Submission date 24/09/2018 | Recruitment status No longer recruiting | <input type="checkbox"/> Prospectively registered |
| | | <input type="checkbox"/> Protocol |
| Registration date 02/10/2018 | Overall study status Completed | <input type="checkbox"/> Statistical analysis plan |
| | | <input checked="" type="checkbox"/> Results |
| Last Edited 08/05/2019 | Condition category Infections and Infestations | <input type="checkbox"/> Individual participant data |

Plain English summary of protocol

Background and study aims

Healthy human skin and mucosa is colonized by a variety of protective bacteria and microorganisms. A common component of this skin flora is the bacterium *Staphylococcus aureus* (*S. aureus*). Dialysis patients are chronically exposed to *S. aureus*, due to their frequent stays in dialysis centers, hospitals or rest homes, where this bacterium is common. In dialysis patients, the access point for the dialysis is a potential entry site for *S. aureus*, in particular when using a central venous catheter instead of an arteriovenous fistula. It has been shown that *S. aureus* carriers have a lower risk of sepsis in case of an endogenous infection (i.e. by their "own" *S. aureus* strain).

So far, it has not been possible to develop a vaccine that protects against an infection with *S. aureus*. The aim of this study is to collect information on the functionality of the immune system in dialysis patients, and a long-term reduction of serious clinical complications due to *S. aureus* infections.

Who can participate?

Hemodialysis patients in the KfH e.V. outpatient dialysis center in Greifswald, and 20 healthy control patients

What does the study involve?

We will follow a group of 86 hemodialysis patients from an outpatient dialysis center over a 30 month period. We will collect their demographic data and medical history, along with taking blood samples, nasal swabs and swabs from the hemodialysis access site every 6 months. These samples and swabs will be tested for *S. aureus*. We will then compare this to the results from healthy controls to reveal differences resulting from dialysis. We will also look at connections between demographic data and medical history and *S. aureus* infection.

What are the possible benefits and risks of participating?

The benefit of participating is that we will identify potential risk factors that make the occurrence of a bacterial infection more likely, especially in dialysis patients. We want to get an overview of the types of bacteria that are involved in infections, and we are also interested in the proportion of resistant *S. aureus* strains. With this knowledge, we aim to minimize the risk of infection for our patients. There are no known risks of participating in this study.

Where is the study run from?

1. University Medicine Greifswald (Germany)
2. Kuratorium für Dialyse und Nierentransplantation e.V., KfH-Nierenzentrum Greifswald (Germany)

When is the study starting and how long is it expected to run for?

January 2015 to December 2018

Who is funding the study?

1. DAMP Foundation (Germany)
2. University Medicine Greifswald (Germany)

Who is the main contact?

Prof. Dr. med. Sylvia Stracke
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Contact information

Type(s)

Scientific

Contact name

Prof Sylvia Stracke

Contact details

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Additional identifiers

EudraCT/CTIS number

IRAS number

ClinicalTrials.gov number

Secondary identifying numbers

BB O29/15

Study information

Scientific Title

Mortality and bloodstream infections in hemodialysis patients from an outpatient dialysis center (KfH e.V.) in Greifswald with respect to *S. aureus* carrier status, colonization density, *S. aureus* genotyping and host immune response (SaDial-study) compared to the general population of the same geographical region (SHIP-TREND-0)

Acronym

Study objectives

1. The *S. aureus* genotypes in hemodialysis patients differ from the general population due to frequent contact with medical environment
2. Nasal *S. aureus* colonisation protects hemodialysis patients from fatal outcome in case of bloodstream infection by *S. aureus*
3. The host immune response against *S. aureus* in hemodialysis patients predicts the course and the outcome of *S. aureus* sepsis

Ethics approval required

Old ethics approval format

Ethics approval(s)

University Medicine Greifswald, 17/03/2015, internal registration number: BB O29/15

Study design

Observational prospective cohort study

Primary study design

Observational

Secondary study design

Cohort study

Study setting(s)

Other

Study type(s)

Diagnostic

Participant information sheet

Not available in web format, please use contact details to request a participant information sheet

Health condition(s) or problem(s) studied

Staphylococcus aureus bloodstream infection and *S. aureus* carrier status in hemodialysis patients

Interventions

A cohort of 86 hemodialysis patients are followed over a 30 month period. Patient demographic data and medical history are collected, followed and statistically evaluated. Blood samples, nasal swabs and swabs from the hemodialysis access site are taken every 6 months for a period of 30 months and are tested for Staphylococcus aureus. The pathogens are cultured and further characterised by spa-PCR and DNA microarrays. Patient samples are analysed for *S. aureus*-specific antibodies using Luminex, and T-cell responses are analysed using Fluorospot. 20 healthy control patients will receive the same treatment as the cohort of hemodialysis patients; however, swabs and blood and serum samples are only take once in 2015 for this group. There is no follow-up period.

Intervention Type

Other

Primary outcome measure

1. Overall mortality rates, assessed using Kaplan-Meier analysis after the final sampling
2. Course and severity of bloodstream infection, determined by assessing for symptoms such as leukocytosis, fever and systemic inflammatory response syndrome, and a positive test for *S. aureus* in blood culture. This was assessed at the time of infection, and 7 and 14 days after

Secondary outcome measures

1. *S. aureus* genotype profiles over time, assessed after each 6 month sampling:
 - 1.1. *S. aureus* spa-types determined by spa-PCR and out coming sequences classified by Ridom software
 - 1.2. Clonal complexes (CC types) of *S. aureus* determined using *S. aureus* Genotyping Kit 2.0 from Alere Technologies (now Abbott)
2. *S. aureus* specific antibody response over time assessed using a multiplex assay and ELISA, with antibodies isolated every 6 months and stored and overall measurement taken once all samples are collected
3. T-cell response over time, assessed by a Fluorospot assay, with T-cells isolated every 6 months and stored and overall measurement taken once all samples are collected

Overall study start date

01/01/2015

Completion date

31/12/2018

Eligibility

Key inclusion criteria

Patients:

1. Hemodialysis patients of an outpatient dialysis centre (KfH e.V.) in Greifswald
2. Voluntary participation

Controls:

1. Voluntary participation
2. Aged 50-70
3. Renally healthy
4. In hospital for at least 1 week

Participant type(s)

Patient

Age group

Adult

Sex

Both

Target number of participants

86 hemodialysis patients, 20 renal healthy control patients

Total final enrolment

86

Key exclusion criteria

1. Refusal to participate
2. Moving to another town that is not Greifswald

Date of first enrolment

18/06/2015

Date of final enrolment

01/03/2018

Locations**Countries of recruitment**

Germany

Study participating centre

KfH-Nierenzentrum Greifswald

Ferdinand-Sauerbruch-Str.

Greifswald

Germany

17489

Sponsor information**Organisation**

Damp Stiftung

Sponsor details

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Sponsor type

University/education

Website

<http://www.damp-stiftung.de/index.php?id=stiftung>

Organisation

University Medicine Greifswald

Sponsor details

Universitätsmedizin Greifswald
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Sponsor type

University/education

Website

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Funder(s)**Funder type**

Not defined

Funder Name

Damp Stiftung

Funder Name

University Medicine Greifswald

Results and Publications**Publication and dissemination plan**

S. aureus carrier rates, genetic studies on *S. aureus* strains and comparison with the general population are currently being published in a peer-reviewed nephrological journal. Immunological work is planned to be published in the end of 2019 in a high-impact peer-reviewed journal.

Intention to publish date

31/12/2018

Individual participant data (IPD) sharing plan

The data could be made available on request after publication.

IPD sharing plan summary

Data sharing statement to be made available at a later date

Study outputs

| Output type | Details | Date created | Date added | Peer reviewed? | Patient-facing? |
|---------------------------------|---------|--------------|------------|----------------|-----------------|
| Results article | results | 06/05/2019 | 08/05/2019 | Yes | No |