

# Identification of potential pathogens responsible for influenza-like illness in elderly in The Netherlands. Evaluation of humoral and cellular immunity against identified microorganisms

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Primary: to determine the percentage of ILI attributable to influenza virus in elderly individuals \* 60 years of age Secondary: to determine the relative contribution of influenza viral subtypesSecondary: to determine humoral and cellular immune...

<b>Ethical review</b>	Approved WMO
<b>Status</b>	Recruitment stopped
<b>Health condition type</b>	Viral infectious disorders
<b>Study type</b>	Observational invasive

## Summary

### ID

NL-OMON40776

### Source

ToetsingOnline

### Brief title

ILI in elderly - GRIEP-3

### Condition

- Viral infectious disorders

### Synonym

flu, influenza

### Research involving

Human

## Sponsors and support

**Primary sponsor:** RIVM

**Source(s) of monetary or material Support:** Ministerie van OC&W

## Intervention

**Keyword:** elderly, humoral and cellular immunity, Influenza-Like-Illness (ILI), respiratory pathogens

## Outcome measures

### Primary outcome

Primary: presence of influenza virus in nasopharyngeal and oropharyngeal swab during ILI episodes

### Secondary outcome

Secondary: influenza virus subtypes in case of influenza infection

Secondary: antibody levels to influenza virus and activation of T cells after influenza infection (humoral and cellular immune responses)

Secondary: presence of viruses (other than influenza) in nasopharyngeal and oropharyngeal swabs collected in subjects reporting with ILI during ILI episodes, 2-3 weeks later and 7-9 weeks later, and additionally at two time points with a 2-3 week interval in a subset of participants without having had ILI

Secondary: presence of bacterial microorganisms in nasopharyngeal and oropharyngeal swabs collected in subjects reporting with ILI during ILI episodes, 2-3 weeks later and 7-9 weeks later after, and additionally at two time points with an interval of 2-3 weeks in a subset of participants without having had ILI

Secondary: humoral and cellular immune responses against potential pathogens

identified in PCR/culture data

Secondary: presence of *S. pneumoniae* in saliva

Secondary: a SF-36 (short-form health survey) questionnaire and vaccine

acceptance at the start of the season, i.e. after signing of Informed Consent

Exploratory: humoral and cellular immune responses against herpes viruses

Exploratory: profile of the intestinal microbiome

## Study description

### Background summary

The public is questioning the effectiveness of seasonal influenza vaccination in elderly as a result of the general impression that all influenza-like illness (ILI) is caused by an influenza virus infection. However, several pathogens, both viral and bacterial, can also cause ILI. A better understanding of the percentage of ILI caused by an influenza virus infection and the contribution of other respiratory viruses or involvement of bacteria will allow a better appreciation of seasonal influenza vaccines. In earlier studies in this population, the incidences of influenza and other microorganisms were determined in subjects with ILI, in seasons of varying influenza incidence, including the humoral response.

In the follow-up study additional information will be collected on the cellular immune responses induced by the different viruses, bacteria and influenza vaccination, the severity of respiratory symptoms during ILI, respiratory symptoms in the absence of ILI, the acceptance of influenza vaccination and the influence of the intestinal microbiome on the immune responses after infection or vaccination.

### Study objective

Primary: to determine the percentage of ILI attributable to influenza virus in elderly individuals \* 60 years of age

Secondary: to determine the relative contribution of influenza viral subtypes

Secondary: to determine humoral and cellular immune responses to influenza virus

Secondary: to identify which microorganisms (viral and bacterial) present in nasopharynx and oropharynx of elderly suffering from ILI are potential other causes for ILI

Secondary: to determine humoral and cellular immune responses towards the

potential pathogens identified in PCR/culture data

Secondary: to gain insight in the influence of viral presence on co-colonization of well-known respiratory bacteria like *S. pneumoniae*, *H. influenzae*, *M. catarrhalis*, *S. aureus* in elderly by comparing colonization during ILI, after recovery and without having had ILI (baseline)

Secondary: to compare the presence of *S. pneumoniae* in nasopharyngeal swab with saliva

Exploratory: to evaluate whether differences can be found in incidence of influenza virus infection between subjects who have, and those who have not, received the seasonal influenza vaccine in the year of study

Exploratory: to compare the incidences of the detected pathogens with other available age group cohorts or with other cohorts of the same age group, such as the previous GRIEP-1/-2 study

Exploratory: to evaluate the immune responses to different herpesviruses in the context of a possible role in immunosenescence

Exploratory: to analyze the intestinal microbiome in context of influenza vaccination response and identified microorganisms

Exploratory: to evaluate whether there is a difference in the general physical and mental health condition as assessed by SF-36 questionnaire and vaccine acceptance between the subjects that report with ILI and the whole study population

## **Study design**

Observational cross-sectional study without an investigational medicinal product but with invasive measurements.

## **Study burden and risks**

The burden associated with participation are reporting of ILI, and collection of a nasopharyngeal swab, an oropharyngeal swab, 6 tubes of blood (53 ml), some saliva and a small faecal sample in case a subject reports with ILI. This sample collection is repeated twice. We expect that at least 200 subjects will report with ILI during the season. In a subset of 150 subjects who have not (yet) experienced ILI the same sample sets are collected twice.

The potential risks are considered minimal. There are no benefits for the individual subjects who participate in this trial. The results may contribute to a better control of respiratory diseases on a population level in the future. The study population is the target of the yearly influenza vaccination campaign.

## **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

\* \* 60 years of age

\* Willing to present when ILI symptoms occur

\* Signed Informed Consent

### Exclusion criteria

none

## Study design

## Design

**Study type:** Observational invasive

Masking: Open (masking not used)

Control: Uncontrolled

Primary purpose: Diagnostic

## Recruitment

NL

Recruitment status: Recruitment stopped

Start date (anticipated): 15-07-2014

Enrollment: 2500

Type: Actual

## Ethics review

Approved WMO

Date: 04-07-2014

Application type: First submission

Review commission: METC Noord-Holland (Alkmaar)

Approved WMO

Date: 22-09-2014

Application type: Amendment

Review commission: METC Noord-Holland (Alkmaar)

Approved WMO

Date: 24-12-2014

Application type: Amendment

Review commission: METC Noord-Holland (Alkmaar)

## Study registrations

**Followed up by the following (possibly more current) registration**

No registrations found.

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

ID: 26253

Source: Nationaal Trial Register

Title:

## In other registers

Register	ID
CCMO	NL49128.094.14
OMON	NL-OMON26253