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Prospective 24-week, double-blind, randomised, placebo-controlled, multicentre study evaluating safety and change in efficacy-related surrogate parameters in patients with dementia of the Alzheimers type under treatment with increasing dosages of intravenous immunoglobulin (Octagam® 10%)

Submission date
23/01/2009

Registration date
28/01/2009

Last Edited
22/03/2016

Recruitment status
No longer recruiting

Overall study status
Completed

Condition category
Nervous System Diseases

[X] 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
NCT00812565

Secondary identifying numbers
GAM10-04

Study information

Scientific Title
Prospective 24-week, double-blind, randomised, placebo-controlled, multicentre study evaluating safety and change in efficacy-related surrogate parameters in patients with dementia of the Alzheimers type under treatment with increasing dosages of intravenous immunoglobulin (Octagam® 10%)

Study objectives
Comparison of different dosages and intervals of intravenous immunoglobulin (IVIG) treatment on surrogate parameters for Alzheimer's disease progression.

Ethics approval required
Old ethics approval format

Ethics approval(s)
1. Central Ethics Committee EC Marburg (Germany)
2. Local Institutional Review Boards (IRBs) of three Unites States of America (USA) sites; approval of protocol amendment no. 2 by IUPUI/Clarian IRB (Indianapolis, USA), 08/01/2009, ref: 0811-07

Study design
Prospective multicentre double-blind randomised placebo-controlled phase II study

Primary study design
Interventional

Secondary study design
Randomised controlled trial

Study setting(s)
Hospital

Study type(s)
Treatment

Participant information sheet
Not available in web format, please use the contact details below to request a patient information sheet

Health condition(s) or problem(s) studied
Alzheimer's disease (mild to moderate)

Interventions
Octagam 10% (12 infusions of 0.1 g/kg, 0.25 g/kg or 0.4 g/kg every 2 weeks or 6 infusions of 0.2 g /kg, 0.5 g/kg or 0.8 g/kg every 4 weeks) or matching placebo.

Blood samples will be drawn before each infusion and at day 1, 4, 7, 14, 21 and 28 (the latter two only for patients on 4-week interval) after last infusion. Lumbar puncture will be performed at baseline and 1 day after last infusion, MRI at screening, week 12 and 24 and 18-fluoro-2-deoxy-glucose-positron emission tomography (FDG-PET) scans at baseline and week 24.

Intervention Type
Drug

Phase
Phase II

Drug/device/biological/vaccine name(s)
Octagam®

Primary outcome measure
Evaluation of the decrease of total amyloid beta in the central nervous system (CNS) and the increase in blood plasma after 24 weeks (area under curve [AUC] of total amyloid beta [Abeta] concentration in plasma).

Secondary outcome measures
1. Further characterisation of the decrease of amyloid beta in the cerebrospinal fluid (CSF) and the increase in blood plasma by measuring an additional surrogate parameter (biomarker, Ab1-42), by assessing the changes in the biomarker proteins Tau and phosphorylated Tau (pTau 181) after 6 months of treatment and of the anti-Ab autoantibodies during the 6-month treatment period
2. Change in Alzheimer Disease Assessment Scale-Cognitive (ADAS-Cog), MMSE, Alzheimer's Disease Cooperative Study-Clinical Global Impression of Change (ADCS-CGIC) and Clinical Dementia Rating Sum of Boxes (CDR-SOB) at week 12 and 24 compared to baseline
3. Change in whole brain and hippocampal volume on volumetric MRI at week 12 and 24 compared to screening
4. Change in cerebral glucose metabolism determined by FDG-PET at week 24 compared to baseline

Overall study start date
01/02/2009

Completion date
21/09/2010

Eligibility

Key inclusion criteria
1. Probable Alzheimer's disease (AD) according to National Institute of Neurological and Communicative Disorders and Stroke and the Alzheimer's Disease and Related Disorders Association (NINCDS-ADRDA) criteria

2. Written informed consent by patient or, for significantly cognitively impaired individuals, their legally authorised representative
3. Aged greater than or equal to 50 and less than or equal to 85 years, either sex
4. Mini-mental State Examination (MMSE) greater than or equal to 16 and less than or equal to 26
5. Only for Germany: the patient's capacity to consent has to be confirmed by dated signature on the informed consent form by a second independent investigator who is otherwise not involved in study GAM10-04
6. Modified Hachinski-Rosen Score less than 5
7. Magnetic resonance imaging (MRI) of the head consistent with the diagnosis of AD

Participant type(s)
Patient

Age group
Senior

Sex
Both

Target number of participants
56 (7 patients per arm)

Key exclusion criteria
1. Other causes of dementia (e.g. vascular dementia, Lewy Body dementia, fronto-temporal dementia, Creutzfeld-Jacob disease, Huntington's disease, Parkinson's disease)
2. History of or present significant other diseases of the central nervous system (e.g. brain tumour, normal pressure hydrocephalus, stroke, severe brain trauma, brain surgery, epilepsy, encephalitis)
3. Geriatric depression scale of greater than 7 (short form with scale from 0 to 15)
4. Present significant psychiatric disorder (e.g. major depression)
5. History of psychosis or hallucinations
6. Mental retardation
7. Unstable medical disease in the opinion of the investigator
8. Insulin dependent diabetes mellitus
9. Acute infectious disease
10. Uncontrolled hypertension (diastolic blood pressure [BP] greater than 90 mmHg or systolic BP greater than 160 mmHg; sitting)
11. Symptomatic stroke
12. Transient ischaemic attack (TIA) within preceding 2 years
13. Participation in other drug trial currently or within the previous 3 months before screening

Date of first enrolment
01/02/2009

Date of final enrolment
21/09/2010

Locations

Countries of recruitment

Austria

Germany

United States of America

Study participating centre
Oberlaaer Str. 235
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Austria
1100

Sponsor information

Organisation
Octapharma AG (Switzerland)

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Sponsor type
Industry

Website
http://www.octapharma.com

ROR
https://ror.org/002k5fe57

Funder(s)

Funder type
Industry

Funder Name
Octapharma AG (Switzerland)

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?
Basic results				No	No
Results article	results	01/03/2013		Yes	No