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Safety, tolerability and pharmacokinetics of Ginkgo biloba special extract EGb 761® in patients with hepatic dysfunction

Submission date 02/10/2009	**Recruitment status** No longer recruiting	[_] Prospectively registered [_] Protocol
Registration date 30/11/2009	**Overall study status** Completed	[_] Statistical analysis plan [_] Results
Last Edited 30/11/2009	**Condition category** Digestive System	[_] Individual participant data [_] Record updated in last year

Plain English summary of protocol
Not provided at time of registration

Contact information

Type(s)
Scientific

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Additional identifiers

Protocol serial number
523001.01.092

Study information

Scientific Title

Single-centre, open-label, parallel-group trial to study the in-vivo effects of impaired hepatic function on the safety, tolerability and pharmacokinetics of Ginkgo biloba special extract EGb 761® in man

Acronym
EGb 761®: Impaired Hepatic Function

Study objectives
To describe the pharmacokinetics, safety, and tolerability of single oral doses of EGb 761® in hepatic dysfunction relative to matched healthy controls.

Ethics approval required
Old ethics approval format

Ethics approval(s)
Ethics Committee MHAT "St. Ivan Rilski" approved on the 25th August 2009 (ref: 16/25.08.2009)

Study design
Single-centre controlled open-label parallel-group trial

Primary study design
Interventional

Study type(s)
Treatment

Health condition(s) or problem(s) studied
Liver cirrhosis

Interventions
Period 1: a single oral dose 120 mg Ginkgo biloba special extract EGb 761®
Period 2: a single oral dose 240 mg Ginkgo biloba special extract EGb 761®

The first period is preceded by a screening visit for eligibility assessment within 21 to 2 days before hospitalisation. For each period, the subjects are hospitalised in the study clinic from the evening before dosing until 24.00 hours after dosing. Periods are at least one week apart for wash-out. An end-of-trial safety follow-up visit is scheduled within one week after Period 2.

Intervention Type
Drug

Phase
Phase I

Drug/device/biological/vaccine name(s)
EGb 761®

Primary outcome(s)
Clinical pharmacology criteria:
1. Pharmacokinetics: single-dose pharmacokinetics of relevant marker terpenes (Ginkgolide A,

Ginkgolide B and Bilobalide) in plasma (optional: in urine)
2. Wellbeing and adverse events
3. Recumbent resting blood pressure and pulse rate
4. Physical examination
5. Clinical laboratory safety tests

Key secondary outcome(s))
No secondary outcome measures

Completion date
28/02/2010

Eligibility

Key inclusion criteria
All subjects:
1. Males or females (females of non-child-bearing potential or of child-bearing potential while taking medically appropriate contraception)
2. Caucasian
3. Aged 21 to 60 years of age
4. Body mass index (BMI) between 18 - 30 kg/m^2
5. Body weight between 45 - 100 kg
6. Willing and able to provide informed consent

Healthy control subjects (CON):
7. Healthy based on the pre-study examination

Patients with moderate hepatic dysfunction (CTP-class B) (HEP):
8. Stable compensated liver cirrhosis (cryptogenic, post-hepatitis, alcohol-related) with histological or macroscopic (e.g. laparascopy, biopsy, ultrasound sonography or other adequate imaging techniques) confirmation
9. Child-Turcotte-Pugh (CTP) class B (sum of CTP-scores: 7 - 9)
10. Liver Vascular Index by Doppler ultrasonography T 12 cm/sec

Participant type(s)
Patient

Healthy volunteers allowed
No

Age group
Adult

Lower age limit
21 years

Upper age limit
60 years

Sex

All

Key exclusion criteria

General: all subjects -
1. Previous participation in the trial
2. Participant in any other trial during the last 90 days
3. Donation of blood during the last 60 days or a history of blood loss exceeding 300 ml within the last 3 months
4. History of any clinically relevant allergy (including hypersensitivity to the trial medications)
5. Presence of acute or chronic infection (HEP: other than related to the primary diagnosis - chronic hepatitis or chronic pancreatitis are no reason for exclusion)
6. Uncontrolled diabetes mellitus
7. Resting systolic blood pressure greater than 160 or less than 90 mmHg, diastolic blood pressure greater than 95 or less than 50 mmHg
8. Clinically relevant electrocardiogram (ECG)-abnormalities, prolonged QTc with greater than 450 msec in males and greater than 460 msec in females in particular
9. Positive human immunodeficiency virus (HIV) test
10. Positive alcohol or urine drug test on recruitment
11. Daily alcohol use of greater than 30 g alcohol
12. Smoking more than 10 cigarettes/day or equivalent of other tobacco products
13. Use of prohibited medication
14. Suspicion or evidence that the subject is not trustworthy and reliable
15. Suspicion or evidence that the subject is not able to make a free consent or to understand the information in this regard

General: all females -
16. Positive pregnancy test
17. Lactating
18. Not using appropriate contraception in premenopausal women

All healthy subjects:
19. Presence or history of any relevant co-morbidity
20. Presence of any clinically relevant abnormality in the laboratory safety tests, especially low haemoglobin, increased liver enzymes, increased serum creatinine
21. Positive serology for hepatitis B surface antigen (HBsAg), hepatitis B core antigen (anti-HBc) and hepatitis C virus antigen (anti-HCV)
22. History of alcohol and/or drug abuse

Patients with hepatic disease:
23. Biliary liver cirrhosis
24. Liver impairment due to space-occupying processes (e.g. carcinoma)
25. State after liver transplantation or patient scheduled for liver transplantation
26. Fluctuating or rapidly deteriorating hepatic function
27. Significant bleeding diathesis
28. Oesophageal bleeding within the last 8 weeks before study entry
29. More than moderate ascites on abdominal ultrasound (US)
30. Presence or history of any relevant co-morbidity other than hepatic disease
31. Clinically relevant abnormal laboratory values other than those associated or sufficiently explained by the existing liver disease, the cut-off level of serum haemoglobin for exclusion: 100 g/l
32. History of drug or alcohol abuse within 2 months prior to dosing

Date of first enrolment
01/11/2009

Date of final enrolment
28/02/2010

Locations

Countries of recruitment
Bulgaria

Study participating centre
Deptartment of Gastroenterology
Sofia
Bulgaria
1431

Sponsor information

Organisation
Dr. Willmar Schwabe GmbH & Co. KG (Germany)

ROR
https://ror.org/043rrkc78

Funder(s)

Funder type
Industry

Funder Name
Dr. Willmar Schwabe GmbH & Co. KG (Germany)

Results and Publications

Individual participant data (IPD) sharing plan

IPD sharing plan summary
Not provided at time of registration