GOOD MORNING
AND
WELCOME

The International Liver Congress
Berlin, April 2011
INTRODUCTION AND AIMS
PROTOCOL OVERVIEW, CRF AND "OTHER FORMS"
PROTOCOL OVERVIEW:

INCLUSION CRITERIA

- HBsAg +
- antiHDV +
- ≥ 18 years old (?)
- signed informed consent

IDEALLY

→ 1000 PATS. IN THE FIRST 2 YEARS
→ 300 PATS./YEAR THEREAFTER
• Any cause of relevant liver disease other than HDV.
PROTOCOL OVERVIEW: PROCEDURES

✓ ALL INCLUSION CRITERIA
✓ NO EXCLUSION CRITERIA
PROTOCOL OVERVIEW: PROCEDURES

- ALL INCLUSION CRITERIA
- NO EXCLUSION CRITERIA
PROTOCOL OVERVIEW: PROCEDURES

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✓ NO EXCLUSION CRITERIA
PROTOCOL OVERVIEW: PROCEDURES

- ALL INCLUSION CRITERIA
- NO EXCLUSION CRITERIA

INFORMED CONSENT

CF (centre form)
P (patient form)

INFORMED CONSENT

[Diagram of medical procedures]
PROTOCOL OVERVIEW: **PROCEDURES**

- ALL INCLUSION CRITERIA
- NO EXCLUSION CRITERIA
PROTOCOL OVERVIEW: PROCEDURES

- ALL INCLUSION CRITERIA
- NO EXCLUSION CRITERIA
PROTOCOL OVERVIEW: PROCEDURES

- Informed Consent
- CF (centre form)
- PF (patient form)

Flow diagram showing the process of obtaining informed consent and the roles of the centre and patient forms in the protocol overview.
PROTOCOL OVERVIEW: PROCEDURES

INFORMED CONSENT

CF (centre form)

PF (patient form)

Contact us
PROTOCOL OVERVIEW: PROCEDURES
PROTOCOL OVERVIEW: PROCEDURES
PROTOCOL OVERVIEW: PROCEDURES

HBV vaccination!

HDV?

INFORMED CONSENT

INFORMED CONSENT

CF (centre form)

PF (patient form)

HBV vaccination!

[Diagram with healthcare professionals and medical equipment]
PROTOCOL OVERVIEW: PROCEDURES

INFORMED CONSENT

CF (centre form)
PF (patient form)

HBsAg? antiHDV? HBV vaccination!
1. Information for the patients

- Updated medical history
- What is hepatitis delta and where to find more information
- “Healthy liver tips”
PATIENT FORM

1. Information for the patients
   • Updated medical history
   • What is hepatitis delta and where to find more information
   • “Healthy liver tips”

2. Information for other health professionals
   ▪ What is hepatitis delta?
   ▪ Special measures they should take
   ▪ Where to find more information about HDV
PATIENT FORM

1. Information for the patients
   • Updated medical history
   • What is hepatitis delta and where to find more information
   • “Healthy liver tips”

2. Information for other health professionals
   ▪ What is hepatitis delta?
   ▪ Special measures they should take
   ▪ Where to find more information about HDV

3. Double check the patient’s data
1. Overview of all the patients they have included

2. Identification of typos and inconsistencies
<table>
<thead>
<tr>
<th>SCREENING FORM</th>
<th>PATIENT DATA</th>
<th>CONCOMITANT CONDITIONS</th>
</tr>
</thead>
<tbody>
<tr>
<td>Patient number</td>
<td>Sex</td>
<td>Cirrhosis?</td>
</tr>
<tr>
<td>Date</td>
<td>Height</td>
<td>Previous Decompensation: date, type (ascites, liver encephalopathy, esophageal bleeding) grade</td>
</tr>
<tr>
<td>Diagnosis of chronic Hepatitis D: month-year /estimation</td>
<td>Weight</td>
<td>Diabetes?</td>
</tr>
<tr>
<td>Diagnosis of chronic Hepatitis B: month-year /estimation</td>
<td>Country of birth</td>
<td>Dialysis?</td>
</tr>
<tr>
<td>Informed consent given, discussed and signed; date</td>
<td>Are there any HDV infected relatives?</td>
<td>Alcohol intake?</td>
</tr>
<tr>
<td>Presence of a medical condition associated with chronic liver disease</td>
<td></td>
<td>Pregnant?</td>
</tr>
<tr>
<td>Organ transplantation with an existing functional graft</td>
<td></td>
<td>Other diagnosis, date, intensity</td>
</tr>
<tr>
<td></td>
<td></td>
<td>Concomitant medications: Name; dosis, start date</td>
</tr>
</tbody>
</table>

**PREVIOUS ANTIVIRAL THERAPIES**

| Total number | Therapy 1: IFN / pegIFN / nucleo(t)sides / others | Therapy 1: dates | Therapy 1 : Side effects |
CASE REPORT FORM: CRF

HEMATOLOGY
- Hemoglobin
- Leucocytes
- Neutrophils
- Thrombocytes

COAGULATION: INR

BIOCHEMISTRY
- ALT / GPT
- AST / GOT
- GGT
- Bilirubin total
- Alkaline phosphatase
- Protein total
- Albumin
- BUN
- Creatinine
- Sodium
- Cholinesterin
- Creatinine clearance

VIROLOGY
- antiHAV IgG
- antiHAV IgM
- antiHCV
- HCV-RNA
- antiHIV
- HBsAg qual. and quant.
- antiHBs
- HBeAg
- antiHBe
- HBV-DNA qual. and quant.
- antiHDV
- HDV RNA qual. and quant.

OTHER
- AFP
- AMA, ANA
- PBMCs available?
# CASE REPORT FORM: CRF

## LIVER BIOPSY
- Date
- Score (Ishak)
- Grading and staging
- Frozen liver tissue available?

## TRANSIENT ELASTOGRAPHY
- Date
- Elasticity
- Variability

## USG
- Date
- Spleen length
- Spleen diameter
- Liver steatosis
- Ascites
- Liver nodules

## ARFI: date; result

## ENDOSCOPY: date; esophageal varices grade
GOALS AND DEADLINES

• Unlimited

  * Duration of follow up
  * Number of recruited patients

• Interim analysis:
  • 500 patients enrolled/ 1 year after day 1
  • Once yearly thereafter
Management issues

• Ethics
• Data privacy
• Database; Hannover Clinical Trial Center (HCTC)
PATIENT INFORMATION AND INFORMED CONSENT (i)

Ethics and data privacy

1) Ethics: depending on the local ethic guidelines → we will soon have an approval from MHH’s ethics commission
PATIENT INFORMATION AND INFORMED CONSENT (i)

Ethics and data privacy

1) Ethics: depending on the local ethic guidelines
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Herewith I confirm, that our patient information and informed consent are translated properly

Yes, I have sent back the translated documents
In general: documents should be in accordance with the local standard.
PATIENT INFORMATION AND INFORMED CONSENT (ii)

In general: documents should be in accordance with the local standard.

Data privacy: depending on the local data protection law.

Location of the database is Germany = Datenschutzgesetz.

Ethics: also dependend on the local ethic rules; we will have a ethic vote from the mainly responsible ethics commission.
PATIENT INFORMATION AND INFORMED CONSENT (ii)

In general: documents should be in accordance with the local standard

Data privacy: depending on the local data protection law

location of the database is Germany = Datenschutzgesetz

Ethics: also dependend on the local ethic rules;
we will have an approvation from the mainly responsible ethics commission

→ You are in charge of getting in touch with the local ethics commission

→ Send back the translated form + a retranslation in English
BLINDING: PATIENT CODE

country

centre

patient
BLINDING: PATIENT CODE

e.g. Patient number 5 in Medizinische Hochschule Hannover (Germany)

0 0 1

Germany
BLINDING: PATIENT CODE

e.g. Patient number 5 in Medizinische Hochschule Hannover (Germany)

Germany

MHH
BLINDING: PATIENT CODE

e.g. Patient number 5 in Medizinische Hochschule Hannover (Germany)

Germany 0 0 1
MHH 0 1
Patient number 5 0 0 0 5
BLINDING: PATIENT CODE

e.g. Patient number 5 in Medizinische Hochschule Hannover (Germany)

001-01-0005
HCTC: Central Management Organisation for Clinical Trials:

- cost-effective academic CRO cooperating with medical facility Hannover Medical School (MHH)

- suited in the middle of a complex network of top level partners
HCTC: Central Management Organisation for Clinical Trials:

**Study set-up:**
- Scientific advice
- Regulatory support
- Clinical trial application

**Operational management:**
- Project management
- Monitoring

**Data management and collection**

**Biometric evaluation:**
in cooperation with Div. of Biometry at MHH

**Pharmacovigilance:**
in cooperation with Clinical Pharmacology at MHH
MARVIN
Integrated eclinical application

integrates numerous functions from the areas of
• EDC (Electronic Data Capture)
• DDE (Double Data Entry)
• CDM (Clinical Data Management)
• CTM (Clinical Trial Management)
• Patient Reported Outcome (ePro)
into one data base

Features:
• intuitive smooth data entry and query management
• No specific soft- and hardware requirements (except DSL/ISDN)
• Online reports, pdf print-outs
## eCRF: Marvin

### Laboratory Data

<table>
<thead>
<tr>
<th>Test</th>
<th>Lower Limit</th>
<th>Measured Value</th>
<th>Upper Limit</th>
<th>Unit</th>
<th>Finding</th>
</tr>
</thead>
<tbody>
<tr>
<td>Hemoglobin</td>
<td>12.2</td>
<td>11.2</td>
<td>18.1</td>
<td>g/dL</td>
<td>normal, not significant</td>
</tr>
<tr>
<td>Leucocytes</td>
<td>4.9</td>
<td>7.5</td>
<td>19.2</td>
<td>10^3/µL = Tedj/µL</td>
<td>normal, not significant</td>
</tr>
<tr>
<td>Monocytes</td>
<td>10.0</td>
<td>41.9</td>
<td>50.0</td>
<td>%</td>
<td>normal, not significant</td>
</tr>
<tr>
<td>Basophils</td>
<td>0.0</td>
<td>2.64</td>
<td>7.0</td>
<td>%</td>
<td>normal, not significant</td>
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<tr>
<td>Eosinophils</td>
<td>0.0</td>
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<td>37.7</td>
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<td>53.7</td>
<td>%</td>
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Medical information for patients and their families

Artikel vom 12. November 2011

Lorem ipsum dolor sit amet, consetetur sadipscing elitr, sed diam nonumy eirmod tempor invidunt ut labore et dolore magna aliquyam erat, sed diam voluptua. At vero eos et accusam et justo duo dolores et ea rebum. Stet clita kasd gubergren, no sea takimata sanctus est Lorem ipsum dolor sit amet.
Potential collaboration projects
Thank you!