Montage et conduite de projets de recherche clinique multinationaux Européens
F-CRIN 31 mai 2018

Focus sur les infrastructures de recherche Européennes et les règlements

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31 mai 2018
International cooperation: industry-sponsored vs. academic trials

ECRIN model: distributed infrastructure

Coordinating services provided by national partners

- **National partner:**
  network of clinical trial units (CTUs) able to manage trials in the country

- **National hub**

- **European Correspondent**
  hosted in national hub (ECRIN staff)
ECRIN and its national scientific partners

Core Team, European Correspondents, national hub, CTU networks

Framework contracts with national scientific partners

✓ provision of services
  -> “linked third party”
✓ non-profit cost
✓ hosting the European Correspondent
ECRIN SUPPORT SERVICES

1. PREPARATION: ADVICE & INFORMATION
   - Trial design and methodology
   - Funding sources and costs
   - Investigation sites and patient recruitment
   - Task distribution for multinational trial management
   - Funding applications
   - Regulatory, ethical and insurance requirements

2. REVIEW: PROTOCOL & FEASIBILITY
   - Scientific and methodological evaluation of the protocol
   - Assessment of project implementation plans

3. IMPLEMENTATION: TRIAL MANAGEMENT
   - Project management and trial coordination
   - Clinical study authorisations (regulatory, ethical) and follow-up
   - Monitoring
   - Vigilance
   - Data management
   - Health product and biosample management
ECRIN trial portfolio

Trials Portfolio (Nov 2017)

- German
- France
- Italy
- Spain
- United Kingdom
- Netherlands
- Belgium
- Denmark
- Sweden
- Czech Republic
- Poland
- Norway
- Portugal
- Switzerland
- Finland
- Hungary
- Austria
- Ireland
- Romania
- Turkey
- Iceland
- Bulgaria
- Croatia
- Greece
- Serbia
- Estonia
- Israel
- Latvia
- Lithuania
- Luxembourg
- Slovakia
Funding for multinational trials
### ERIC Participation in submitted proposals

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ERIC Participation in selected proposals

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Tools for multinational trials

ECRIN tools to facilitate multinational trials (www.ecrin.org)

- Quality management
- Data centre certification
- Regulatory and ethical database
- Methodology guidelines

- Outcome measure database
- Risk-based monitoring toolbox
- Mapping of investigation sites
- Training
- Communication
Central resource covering 22+ European countries and multiple study types.

**Use to:** Locate country-specific competent authorities and ethics committees

Consult summary of requirements in each country

Browse related documents
Risk based monitoring toolbox

- Enables researchers to create appropriate risk-based strategies

### Training and Other On-Site Activities

As stated by Baigent et al (Baigent et al. 2008), "on-site monitoring should be ... regarded as "mentoring", providing opportunities for training and supporting study staff". In this section, two papers describing on-site activities beyond SDV are reviewed.

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<th>Topic</th>
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<tr>
<td>Site Monitoring Process Using Peer Reviewers to Improve Staff Training, Site Performance, Data Collection and GCP Compliance</td>
<td>Implementation in a large multicentre trial is described, demonstrating a decrease of findings with ongoing site visits.</td>
<td>Lane et al. 2011</td>
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<td>Procedure for annual site visits performed by an experienced team, with detailed description of proposed on-site activities</td>
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<td>Clinical Trial Educator (CTE) Program – to accelerate enrolment</td>
<td>Non-randomised evaluation of the programme in a large-scale trial showed significantly better recruitment rates in sites visited by a CTE,</td>
<td>Kendall et al. 2012)</td>
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<td>Program involving regular site visits by specifically trained personnel in order to train and educate investigators and site staff with respect to recruitment challenges.</td>
<td></td>
<td>Central Monitoring</td>
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### Central Monitoring
Methodology and design

Review Article

Evidence-based clinical practice: Overview of threats to the validity of evidence and how to minimise them

Silvio Garattini a, Janus C. Jakobsen b,c, Jørn Wetterslev b, Vittorio Bertelé a, Rita Banzi a, Ana Rath d, Edmund A.M. Neugebauer e, Martine Laville f, Yvonne Masson f, Virginie Hivert f, Michaela Eikermann g, Burc Aydin h, Sandra Ngwabiy t d, Cecilia Martinho l, Chiara Gerardi a, Cezary A. Szmigielski j, Jacques Demotes-Mainard k, Christian Gluud b,*

Barriers to the conduct of randomised clinical trials within all disease areas

Snezana Djurisic d, Ana Rath 2, Sabrina Gaber 3, Silvio Garattini 4, Vittorio Bertelé 4, Sandra-Nadia Ngwabiy t 2, Virginie Hivert 5, Edmund A.M. Neugebauer 6, Martine Laville 7, Michael Hiesmayr 8, Jacques Demotes-Mainard 3, Christine Kubiak 3, Janus C. Jakobsen 1,9 and Christian Gluud 1,*
Methodology and design

A systematic literature review of evidence-based clinical practice for rare diseases: what are the perceived and real barriers for improving the evidence and how can they be overcome?

Specific barriers to the conduct of randomised clinical trials on medical devices

Evidence-based practice within nutrition: what are the barriers for improving the evidence and how can they be dealt with?
Experience of ECRIN data center certification, and perspectives

Raising standards in clinical research — The impact of the ECRIN data centre certification programme, 2011–2016

C. Ohmann a,*, S. Canham b, J. Demotes c, G. Chêne d, J. Lauritsen e, H. Martins f, R.V. Mendes g, E.B. Nicolis h, A. Svobodnik i, F. Torres j

https://authors.elsevier.com/sd/article/S2451865416300825
SHARING PATIENT-LEVEL CLINICAL TRIAL DATA

✅ Group Nominal Process
✅ data protection
✅ informed consent
✅ anonymized / pseudonymized
✅ access (open vs. controlled)
✅ standard data format
✅ security
✅ GDPR compliance
✅ type of repositories

**BMJ Open**

Sharing and reuse of individual participant data from clinical trials: principles and recommendations


*Ohmann et al., BMJ Open 2017;7:e018647*

[http://bmjopen.bmj.com/cgi/content/full/bmjopen-2017-018647?ijkey=79SivGTa9igpfbN&keytype=ref](http://bmjopen.bmj.com/cgi/content/full/bmjopen-2017-018647?ijkey=79SivGTa9igpfbN&keytype=ref)
ESFRI-roadmap Biological and Medical Science Research Infrastructures

BMS RIs
BIOLOGICAL AND MEDICAL SCIENCES RESEARCH INFRASTRUCTURES

ERINHA

CORBEL

ELIXIR

INSTRUCT

EMBRC

TARGET IDENTIFICATION

DRUG DISCOVERY/DEVELOPMENT

BIOMARKERS

TRANSLATIONAL RESEARCH

CLINICAL RESEARCH

ESFRI ROADMAP BIOLOGICAL AND MEDICAL SCIENCES RESEARCH INFRASTRUCTURES

European Research Infrastructures for High Pathogenic Agents

BioBanking and BioMolecular resources Research Infrastructure

European infrastructure for translational medicine

Chemical Keys for Life's Locks

Infrastructure for Systems Biology

European Marine Biological Resource Centre

European Clinical Research Infrastructure Network

European Research Infrastructure for Translational Medicine

European Infrastructure for Systems Biology
Medical Infrastructures / Users Forum
EOSC-Life: a European approach to the digitalization of clinical research

- INFRAEOSC-4 application
  - Secure, GDPR compliant cloud environment
  - Share research data and samples
  - Multimodal data analysis and AI algorithms for patient stratification
  - Reuse of health data for research purposes: EHR, EDC, national databases