



BIH QUEST CENTER FOR TRANSFORMING BIOMEDICAL RESEARCH

QUALITY | ETHICS | OPEN SCIENCE | TRANSLATION

Paving the way to robust and reproducible research

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Universität Ulm, 28.3.2019

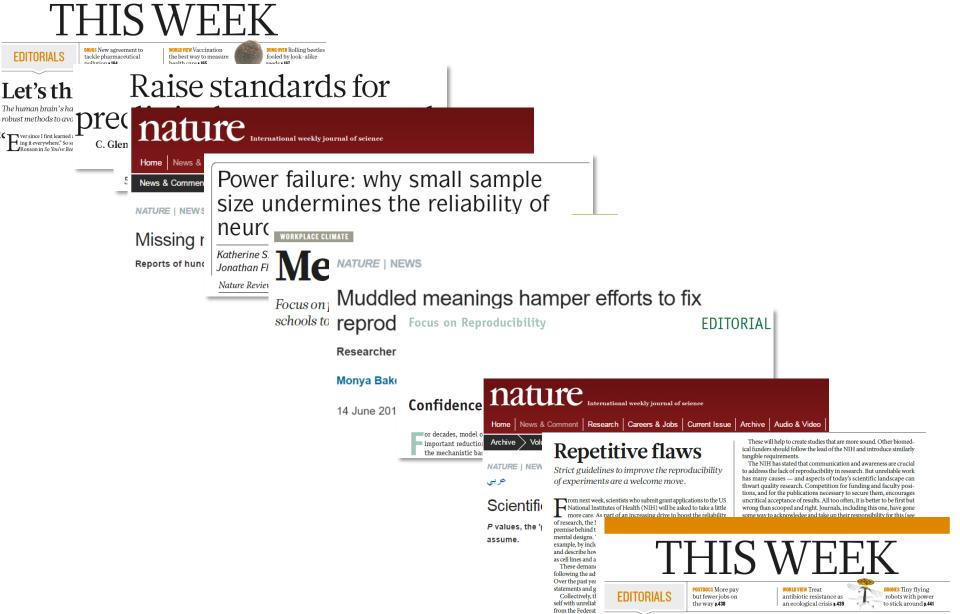


QUEST Mission





.... to increase the value of biomedical research at the Berlin Institute of Health (i.e. Charité and Max Delbrück Center) and beyond.



Reality check on reproducibility

A survey of Nature readers revealed a high level of concern about the problem of irreproducible results. Researchers, funders and journals need to work together to make research more reliable.

Te there a reproducibility crisis in science? Yes, according to the readers of *Nature*. As we report on page 452, two-thirds of researchers who responded to a survey by this journal said that current levels of reproducibility are a major problem.

The ability to reproduce experiments is at the heart of science, yet

for example, call practices. The Soc

of everything from

person ready to question whether a data point or a sample should really be excluded from analysis can help to cut down on cherrypicking, conscious or not. A couple of senior scientists have set up workflows that avoid having a single researcher in charge of preparing images or collecting results. Dozens of respondents reported steps to







BMJ 2018;360:j5845 doi: 10.1136/bmj.j5845 (Published 10 January 2018)

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FEATURE

INVESTIGATION

Oxford vaccine study highlights pick and mix approach to preclinical research

Researchers were disappointed when a clinical trial of a new tuberculosis vaccine failed to show benefit, but should it have gone ahead when animal studies had already raised doubts and what does it mean for future research? **Deborah Cohen** investigates

Deborah Cohen associate editor. The BMJ

"Our meta-analysis suggests an apparent lack of evidence of efficacy in animals in data collected before the start of the recent phase 2b trial in South African children that enrolled children between 15 July 2009 and 4 May 2011," the authors wrote.

"SR of 8 studies showed no efficacy of the vaccine..."

QUEST Approaches





Open Science



Incentives



Education



Stakeholder Engagement



Quality Assurance



Meta Research

QUEST Structure



QUEST-Center

Founding Director: Prof. Ulrich Dirnagl

Research

Office

Transforming biomedical research

Bioethics of translation

Infrastructures & Services

Meta-research

Visiting fellows

Projects

Services & Initiatives





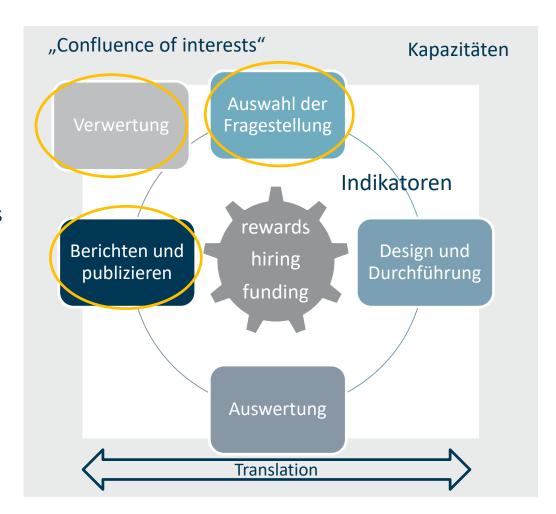
Incentives and indicators-Research assessment





Indicators:

JIF, RCR
3rd party funding
number of
publications
number of patents
author position



MERIT — Mechanisms of robust and reproducible research in translational medicine





Science-based, iterative **policy implementation** project that aims

- to incentivize robust research mechanism by introducing
- 1. Responsible indicators (QUEST criteria)
- 2. Robust evaluations

into the institutional research assessment for funding, hiring and promotion at BIH/Charité

-> to increase patient-oriented translation

MERIT – areas of implementation



1. Intramural funding schemes

- 2. Hiring of professors
- 3. Performance-based funding (LOM)
- 4. Doctoral programs

MERIT — Mechanisms of robust and reproducible research in translational medicine





Two basic assumptions

- Robustness and reproducibility precede patient-oriented translation
- Robustness and reproducibility are multifactorial

Responsible indicators – robust evaluations





QUEST criteria

- different sets i.e. for animal studies, clinical studies
- relating to the mechanisms of robust research in 4 dimensions:
- a) Priority setting /Scientific Premise
- b) Strategies for establishing scientific rigor
- c) Transparency and dissemination of results
- d) Stakeholder engagement

Good evaluation practice guidelines

- WHAT research researcher, environment?
- **WHY** to detect robust and reproducible research
- HOW
- transparent, multidimensional criteria
- anonymous evaluation
 - 1) independent reviewers
 - 2) consensus
- transparent feedback

strong

medium

poor

CURRENT STATE: feasibility phase/development phase

QUEST criteria animal studies 3R plus robustness, registration and reporting





Priority setting /Scientific Premise

Systematic review of the evidence/literature

SYRCLE

Database search: registries

Replacement

Strategies for establishing scientific rigor

e.g. Strategies to reduce risk of bias

sex /gender consideration

Reduction

Transparency and dissemination of results

(Pre)registration

Reporting guidelines (ARRIVE)

Timeline reporting of all results

Layperson comprehensible version

Stakeholder engagement

CURRENT STATE: feasibility phase/development phase

1. MERIT - Intramural funding schemes (BIH)





- Proof of concept QUEST criteria (2/2018-01/2019):
 - introduction to all major funding schemes
 - over 120 applications, 9 board meetings
 - approval of the BIH board
- Analysis, adaption
 - -> status quo of current practices related to QUEST criteria
 - →relevant for selection for funding
 - → relevant for concept for educational measures
 - -> different sets of QUEST criteria (resp. study design, research question)
- Development good evaluation practice guidelines

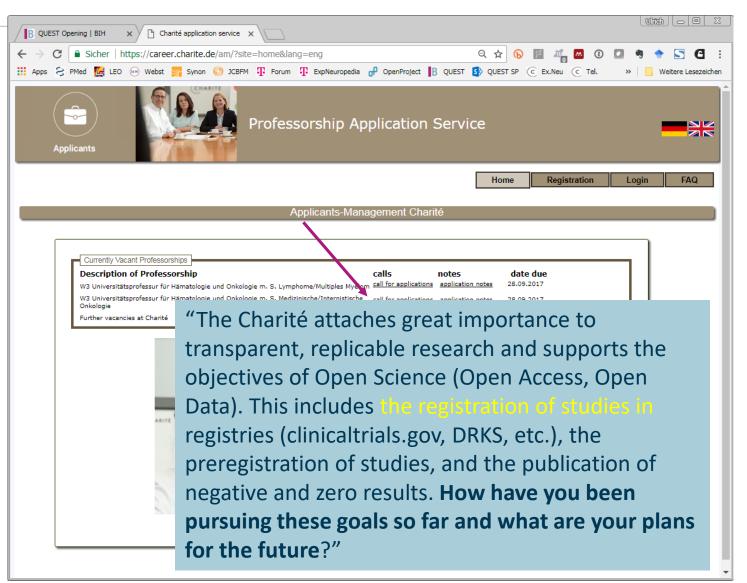
MERIT – hiring commissions professorships (BIH/Charité)











3. MERIT – performance-based funding Open data LOM 2019





- Successful implementation of new indicator "open data" in LOM system Charité (and equivalent for MDC researchers)
- First medical faculty in Germany
 - Development of ODD Algorithm (Riedel, Bobrov, Kip)
 - Retrospective identification of publications with Open Data in institutional publications record (PURE and FACT Science)
 - ca. 3% of all journal articles (2015-2017) with open data (Charité)
- Next Step: Preregistration in LOM by 2021

Awards





The QUEST 1,000 € Preregistration Award

Preregister your preclinical studies challenge - Increase the credibility of your results and get early credit for your ideas!

The QUEST Center supports the preregistration of planned preclinical studies. Preregistration increases the credibility of results, the conclusions based on them, and secures the originality of ideas and hypotheses before experiments have been performed.

QUEST is offering **15 awards of 1,000 €** to first/last/corresponding BIH, MDC or Charité authors of preclinical research papers containing preregistered (basic or translational, not clinical) studies.



The QUEST 1,000 € NULL results and Replication study Award

Publish your NULL results - Fight the negative publication bias!

Publish your Replication Study - Fight the replication crises!

The QUEST Center supports the publication of results which do not support the initial hypothesis, so called NULL results, sometimes unfairly labelled as 'negative studies'. We also want to promote the publication of studies with the explicit attempt to replicate own results or the results of others.

QUEST is offering **15** awards of **1,000** € each to first/last/corresponding BIH, MDC or Charité authors of research papers in which the main result is a NULL or 'negative', or in which the replication of own results or the results of others is attempted. Basic, translational as well as clinical studies are eligible.



THE QUEST 1,000 € OPEN DATA REUSE AWARD

Make use of Open Data!

Open data sharing to increase robustness and value in biomedical research is becoming increasingly common. It is requested by many funders and journals, and both the Charité and the MDC have also signed the Berlin Declaration, which endorses the open sharing of research data. The rapidly growing amount of openly available data has also spurred the development of dataset search engines, which allow the location of relevant datasets. Corresponding to this





QUEST Protected writing time



in cooperation with GB Forschung



- identifies potential candidates for up to 3 months protected time to finalize a first DFGapplication
- criteria (selection):
 - research output (minimum of three 1st author publications in the past 5 years)
 - AND minimum of 2 QUEST criteria shown in at least one of the publications
 - women men ratio 1:1

Institutional publication record (PURE)



 Set up of a comprehensive, continuously updated monitoring instrument of the institutional research output



- Reporting of status quo and trends regarding, e.g.
 - Open science (open data, open access)
 - Preregistration
 - Null results and replication
 - Gender
 - Internationality





Science base MERIT

Theory validation

Development QUEST criteria and Good evaluation practice guidelines

- systematic review of the literature, relevant documents (incl. grey literature, policy documents, reviewer guidelines etc..), expert opinions
- iterative testing and validation
- Implementation
 Introduction to the field
 Data collection and analyses (applications, publication records, field notes, interviews etc..)
- Capacity building
 Teaching measures, tools, services
- Program evaluation and analysis
 Infrastructure, process and outcome evaluation





Policy implementation plan MERIT

- Need and field assessment of the institutional incentive structures
 - identification of relevant areas
 - mapping of the areas (stakeholders,
 - legal or other requirements (approvals, data protection etc...)
 - attending relevant board meetings, committees, trust building
- Agenda and strategy setting
 - engagement of relevant stakeholders
 - development action plans
 - trust building
- Introduction to the field
 - to funding schemes, hiring commissions etc...
 - data collection, document analyses, field notes, interviews
 - development and introduction of teaching measures
- Adjustment







- Wissenschaftsrat 2018
 QUEST's Development of new criteria for the assessment of research "best practice"
- DORA: MERIT hiring commission project among top 10 most promising projects in 2018





THANK YOU!

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