

Enhancing clinical trials

The use of Electronic Medical Records and pre-existing data integration



Collaboration for translational research in dementia

Alzheimer's
Research UK Oxford
Drug
Development
Institute



Wellcome Trust Strategic awards:

Target Enabling Packages

Neuroinflammation



Dementia Discovery cohorts (n=2m)

Dementia UK BioBank cohort (n=10k)

Deep and Frequent Phenotyping cohort



IMI-EPAD

European
Prevention of
Alzheimer's
disease

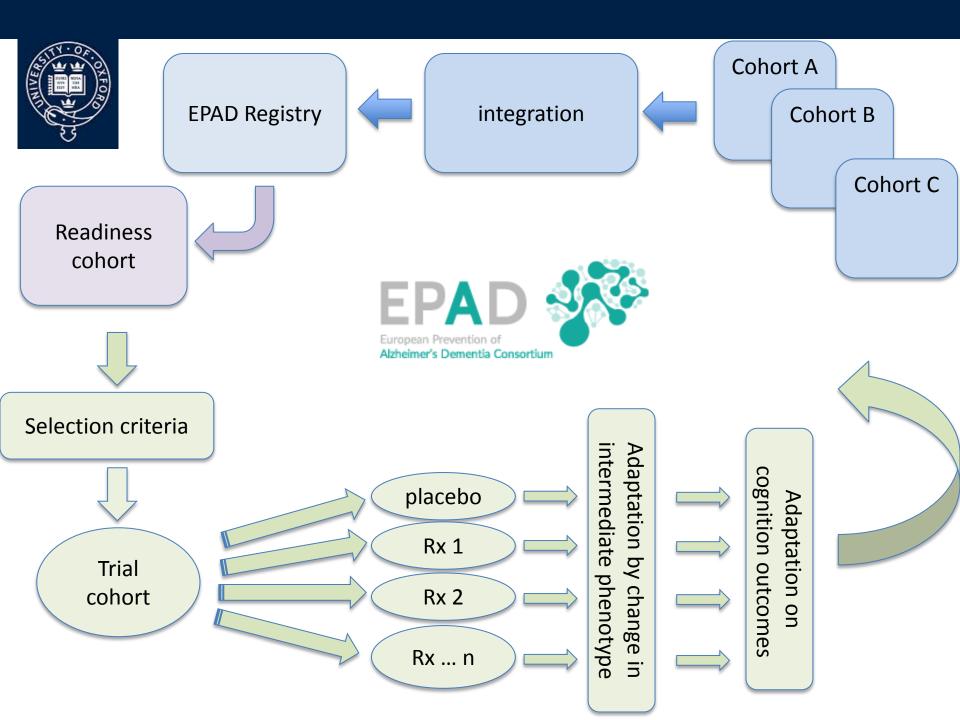
Readiness register, cohort and Early Phase Trials

Pre-competitive data driven drug development

Pre-competitive cohort repurposing for experimental medicine

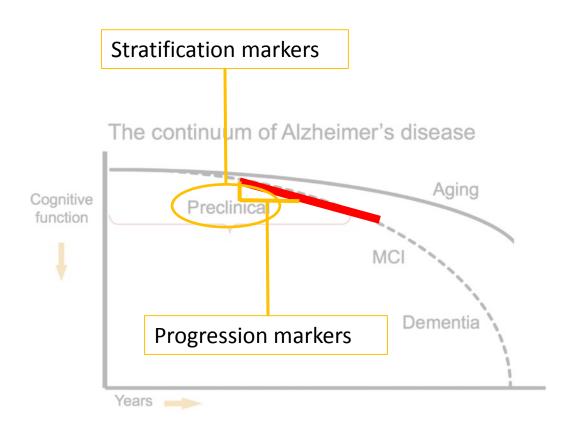
Pre-competitive reutilisation of research and real-world data

Pre-competitive data driven PoC/PhII



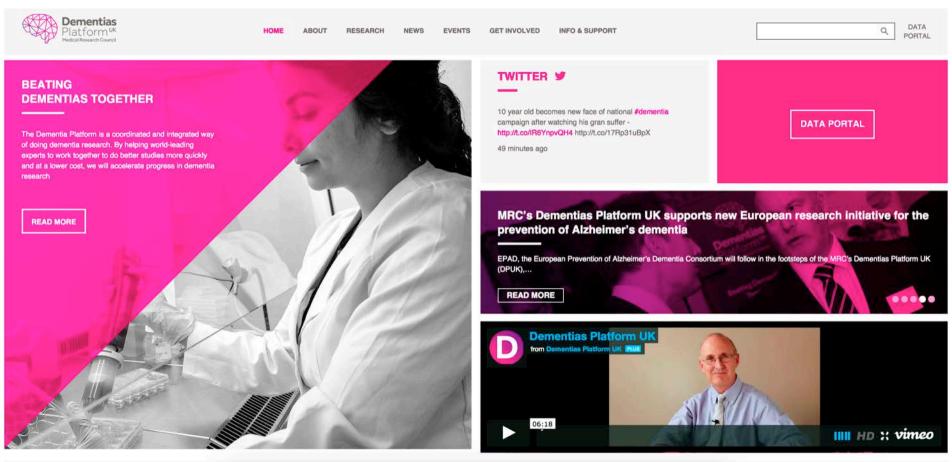


Biomarkers for secondary prevention





Dementias Platform UK



http://www.dementiasplatform.uk/



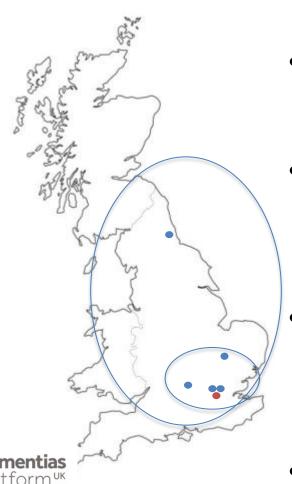
UK Biobank for Dementia Research

- 500,000 participants; baseline assessments 2005-8
- Web-based questionnaires for additional exposures and outcomes (cognition, mental health, occupation..)
- Multimodal imaging in 100,000 brain, cardiac and body fat MRI; bone & joint DEXA; 3D carotid ultrasound
- Repeat Neuroimaging, cognition and sampling in 20,000
- Genotyping of all participants (820,000 SNPs)
- Connectivity to EMRs for mental health





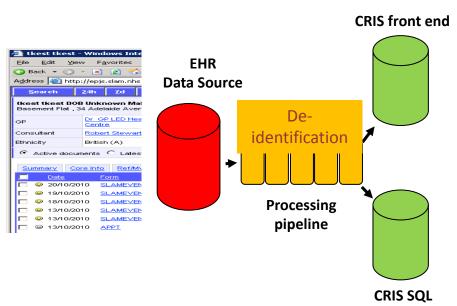
Use of EMRs for research

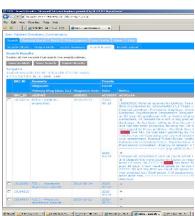


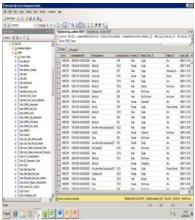
- SLaM CRIS
 - South London and Maudsley NHS BRC implementation
- D-CRIS
 - Cambridge & Peterborough, Oxford Health,
 West London, Camden and Islington
 - 1 million plus patients
- UK-CRIS
 - 10 site extension
 - Connectivity to UK BioBank
- Mike Denis and Simon Lovestone



CRIS – core functionality

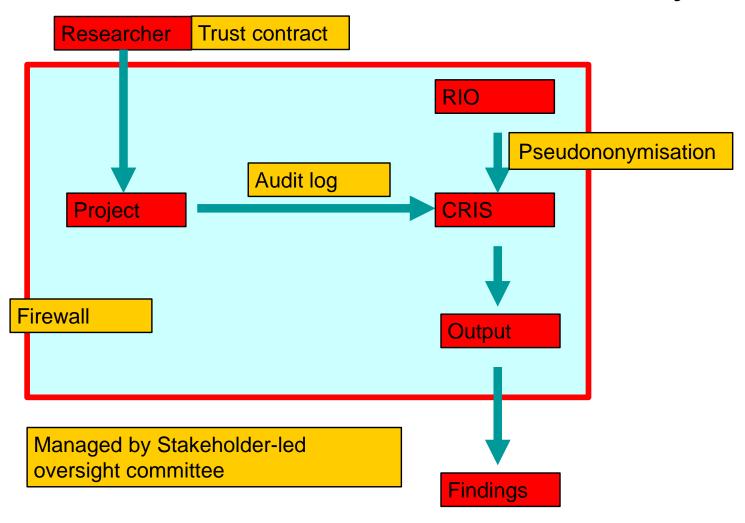








CRIS Security





GATE – Text Parsing

FAST enables information retrieval, i.e. search and retrieval by matching against user defined strings;

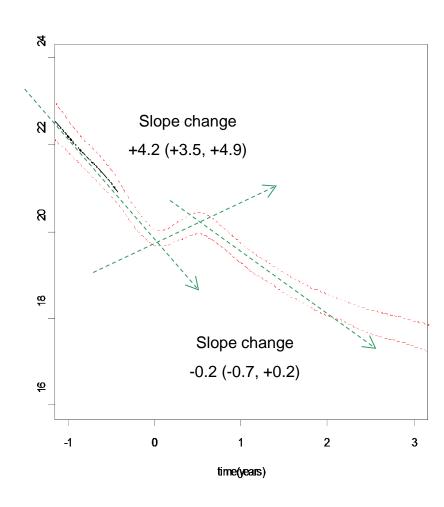
GATE enables information extraction, i.e. extracts 'meaning' (structure) from free text context



EHR-data re-use for research

(Cholinesterase inhibitors and Alzheimer's disease)

- Phase IV of ACHel
 - > 2500 patient years of therapy
 - > 8 fold dataset compared to Cochrane
- Costs and effectiveness
 precompetitive collaboration with pharma
 Text mining derivation of service utilisation and costs
- Predictors of response
 Biomarkers and clinical



Data from Robert Stewart, KCL



European Medical Information Framework







Data available through consortium

Large variety in "types" of data



Primary care data sets



Hospital data



Administrative data



Regional record-linkage systems



Registries and cohorts (broad and disease specific)



Biobanks

 Data is available from more than 53 million subjects from seven EU countries, including

> >40,000 subjects in AD cohorts



>90,000 subjects in metabolic cohorts



EMIF Catalogue





Research cohorts on AD or aging

- Type of cohorts:
 - Population-based cohorts
 - Clinical cohorts
 - National multicenter studies
 - European multicenter studies
 - Clinical trial cohorts





Data available through cohorts in catalogue

Number of subjects	Controls	SCI	MCI
All	31219	3173	11976
Plasma	3537	1764	3961
DNA	8763	3163	5731
RNA	1637	484	2128
CSF	846	1336	3757
Urine	2307	1116	2349
MRI	3449	1035	5586
FDG-PET	551	308	1942
Amyloid PET	335	143	492
EEG	301	773	1173

Either access to results from analysis or access to samples/scans

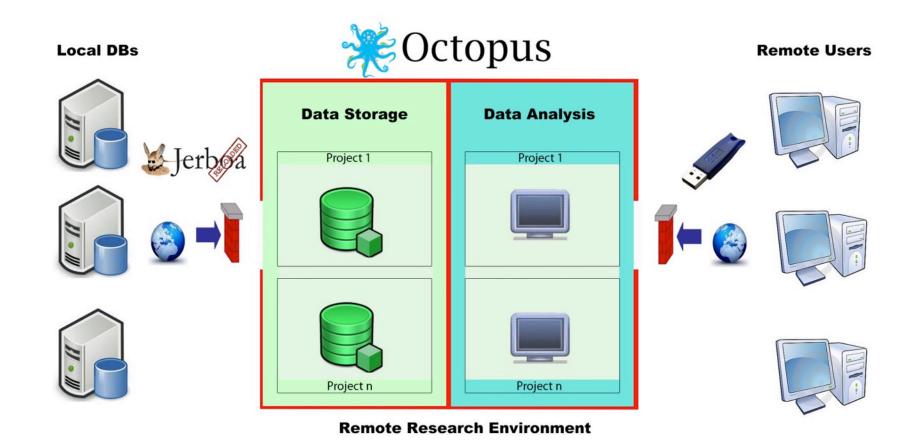


Available EMR data sources – EMIF-Platform

Database name	total (cumulative) # of subjects	# of active patients in January 2012
Aarhus (DK)	2.3 million	1.8 million
Regional DB Tuscany (IT)	4.8 million	3.8 millions
GePaRD (DE)	17 million	12.655.616 (Jan 2007)
THIN (UK)	> 11 million	3.8 million (Sept 2012)
IPCI (NL)	2.9 million	1.6 million
Health Search (IT)	> 2 million	930,000
Pedianet (IT) (0 to 14 years)	400,000	150,000
Pharmo (NL)	> 7 million	5 million
Birth Cohort (UK)	1,184	960
Cardiovasc. Interv. Registry (UK)	460,000	3,000
Salford Diabetes registry (UK) TBC	12,000	12,000
IMASIS (ES)		
EGCUT (ES)	52,000	NA
TOTAL	~ 48 million	~ 17 million

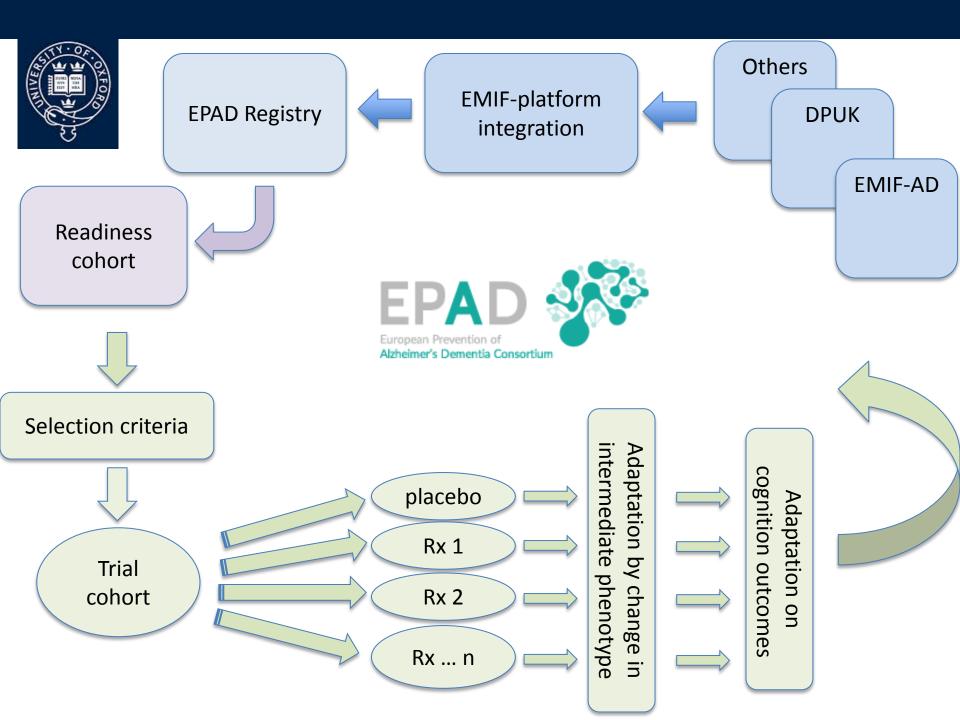


Private Remote Research Environments



Readiness register Modeling cohort Standing PoC adaptive trial







With thanks to....

- > 50 executive members
- > 150 principal investigators
- > 500 scientists
- >1500 collaborators
- > 50,000 volunteers
- > 50,000,000 individuals contributing data
- > 200,000,000 euros from funders





Mike Denis