Co-operating with industry in biomarker studies

Tim Bowen

Department of Nephrology
Wales Kidney Research Unit
Cardiff University









Diabetic nephropathy (DN): A major UK health problem with no effective prognostic biomarkers



Diabetes: Most NHS costs wasteful, says Diabetic Medicine

The majority of NHS spending on diabetes is avoidable, says a report in the journal Diabetic Medicine.

It suggests that 80% of the NHS's £9.8bn annual UK diabetes bill goes on the cost of treating complications.

Experts say much of this is preventable with health checks and better education - something the Department of Health says it is tackling.



Type 2 diabetes is linked to obesity





Kidney care 'needs to improve'

By Adam Brimelow

Health Correspondent, BBC News

An NHS report has called for better detection and earlier treatment to help tackle chronic kidney disease.

The study commissioned by NHS Kidney Care says it causes up to 45,000 premature deaths each year, more than lung and breast cancer combined.

An improved approach could help many people to lead better lives and save the NHS money, it adds

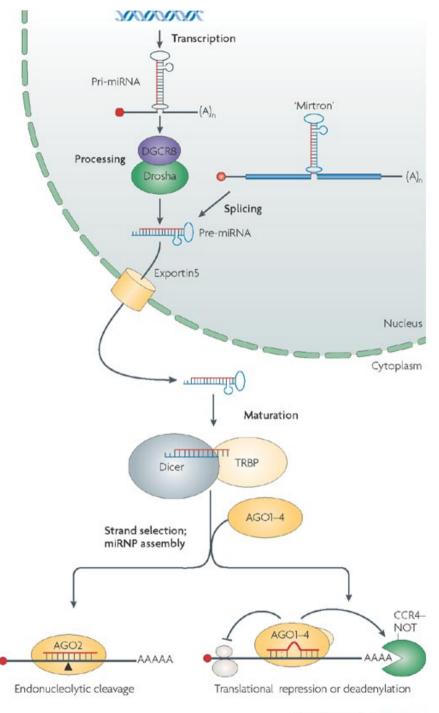


Half of NHS spending on the disease funds dialysis

Investigating the utility of urinary microRNAs as diabetic nephropathy biomarkers

microRNAs bind to specific mRNA targets and inhibit protein synthesis

- 22-25 nucleotide RNA regulators of gene expression
- Imperfect base pairing to target sites in the 3'-untranslated region (UTR)
 - repress translation
 - enhance mRNA degradation
- Sequence-specific target recognition
 - seed region
 - multiple binding sites in 3'-UTR



Knowledge Economy Skills Scholarship (KESS) **European Social Fund & BBI Group**

Invention for Innovation (i4i) early-stage product development grant NISCHR & BBI Group

Industrial CASE 4-year PhD studentship BBSRC & Gwent Group

Knowledge Economy Skills Scholarship (KESS) European Social Fund & BBI Group

Identification of microRNA biomarkers of chronic kidney disease and development of minimally-invasive methods of molecular detection

MicroRNAs: global regulators of gene expression with conserved function

- Accessible, non-invasive sample type
 Alternative to renal biopsy
 Analysis by RT-qPCR

Evaluate feasibility of miRNA test for diabetic kidney disease

Knowledge Economy Skills Scholarship (KESS) **European Social Fund & BBI Group**

Invention for Innovation (i4i) early-stage product development grant NISCHR & BBI Group



Industrial CASE 4-year PhD studentship
BBSRC & Gwent Group

Innovation for Invention (i4i) early-stage product development grant **NISCHR & BBI Group**

Development of urinary microRNAs as biomarkers for diabetic kidney disease

RT-qPCR validation and development of test platform

- Evaluate microRNA utility across disease stages Evaluate microRNA detection in independent patient cohorts Optimise sample preparation / develop test

Prototype microRNA biomarker test / Intellectual property

Knowledge Economy Skills Scholarship (KESS) **European Social Fund & BBI Group**

 \downarrow

Invention for Innovation (i4i) early-stage product development grant NISCHR & BBI Group



Industrial CASE 4-year PhD studentship BBSRC & Gwent Group

Probe development for electrochemical miRNA detection

- Synthesis of electrode-tethered microRNA probe Preliminary data from synthetic urine analysis promising Aim to develop a microRNA "dipstick" probe

In collaboration with Dr James Redman, CU School of Chemistry

Knowledge Economy Skills Scholarship (KESS) **European Social Fund & BBI Group**

Invention for Innovation (i4i) early-stage product development grant

NISCHR & BBI Group

iCASE 4-year PhD studentship BBSRC & Gwent Group

Responsive Funding

Kidney Research UK

Analysis of
urinary microRNAs
as biomarkers
for acute kidney injury

Biomedical Catalyst Confidence in Concept (CiC)

MRC

Development of urinary microRNAs as biomarkers following kidney transplantation

Responsive Funding

IP

Wellcome Trust

The role of microRNAs
in the regulation
and response
to insulin signalling
in the podocyte

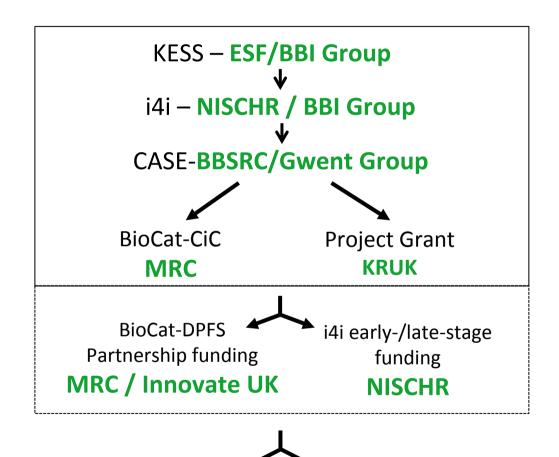
Biomedical Catalyst Confidence in Concept (CiC) MRC

Development of urinary microRNAs as biomarkers following kidney transplantation

MiRNA data from human kidney transplant patient urine samples

- Urine samples from post-transplant patients
- Comparison with IRI/IPC rat models
 Analysis of perfusate solution underway

Biomarker miRNAs to predict transplantation outcomes



IP

Test development

and manufacture

Acknowledgements

WKRU, Cardiff University

Cristina Beltrami

Kate Simpson

Usman Khalid

Robert Jenkins

Chris Carrington

Donald Fraser

Cardiff University

Aled Clayton

James Redman

Ian Weeks

BBI Group Pete Corish

The Gwent Group Guido Drago

Dept Renal Medicine, QEH, Birmingham

Paul Cockwell et al.

Academic Renal Unit, Bristol University Simon Satchell, Richard Coward et al.

All Wales Kidney Research Network

























Contact details

Tim Bowen PhD

Senior Lecturer in Matrix and Molecular Biology
Department of Nephrology
School of Medicine
College of Biomedical and Life Sciences
Cardiff University
Heath Park
Cardiff CF14 4XN UK

Tel direct: +44-(0)29-2074-8389

Tel office: +44-(0)29-2074-8467

Fax office: +44-(0)29-2074-8470

Web home: http://medicine.cf.ac.uk/person/dr-timothy-bowen/