

UK therapeutic

cancer prevention network

collaborating to prevent cancer

Farhat Din

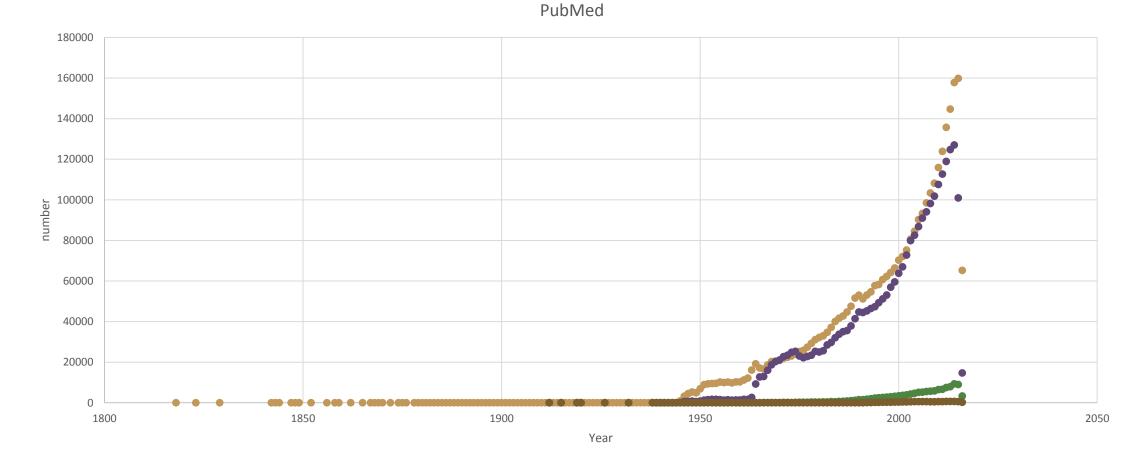
University of Edinburgh



The Experimental Cancer Medicine Centre Initiative is jointly funded by Cancer Research UK, the National Institute for Health Research in England and the Health Departments for Scotland, Wales and Northern Ireland.

"...cannot treat our way out of cancer"

Chris Wild, IARC

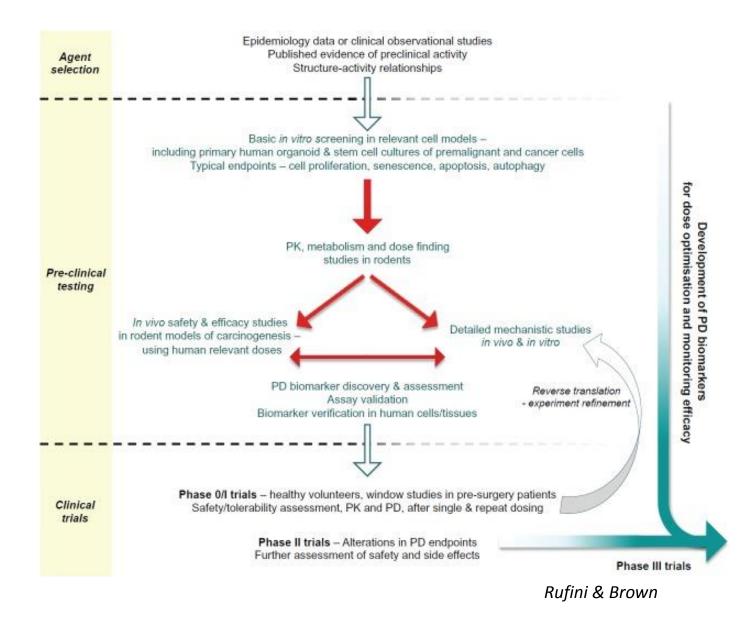


● cancer ● chemotherapy ● prevention ● chemoprevention



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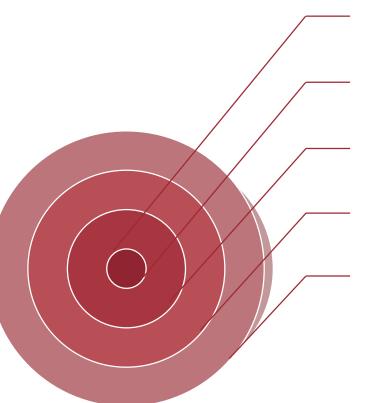
Gap in studies to accelerate cancer prevention field





UK therapeutic cancer prevention network collaborating to prevent cancer

Vision



Cohesive multidisciplinary team of experts working together

Scientific vigour – state of the art well-characterised preclinical models

Robust evidence of activity, identification of mechanisms of action & rational choice of dose range for clinical trials

Reverse translation - preclinical practices informed by clinical experience

Utilise expertise in biomarker discovery and application for the identification of high risk patient groups and monitoring efficacy

Identify, develop and accelerate the translation of therapeutic cancer preventive agents and lifestyle interventions into clinical trials and routine clinical practice

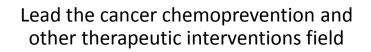


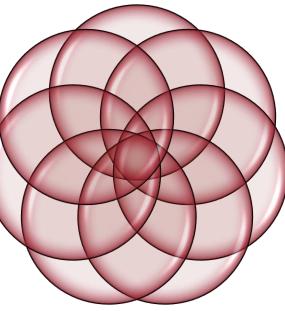
UKTCPN Objectives

Establish network of clinical &non-clinical researchers Diverse complementary backgrounds

Promote collaboration-joint grant applications harnessing complementary strengths across the network

Champion for increased research funding in the field of therapeutic cancer prevention





Identify areas of strength among research groups across the UK and disseminate this information to facilitate collaboration

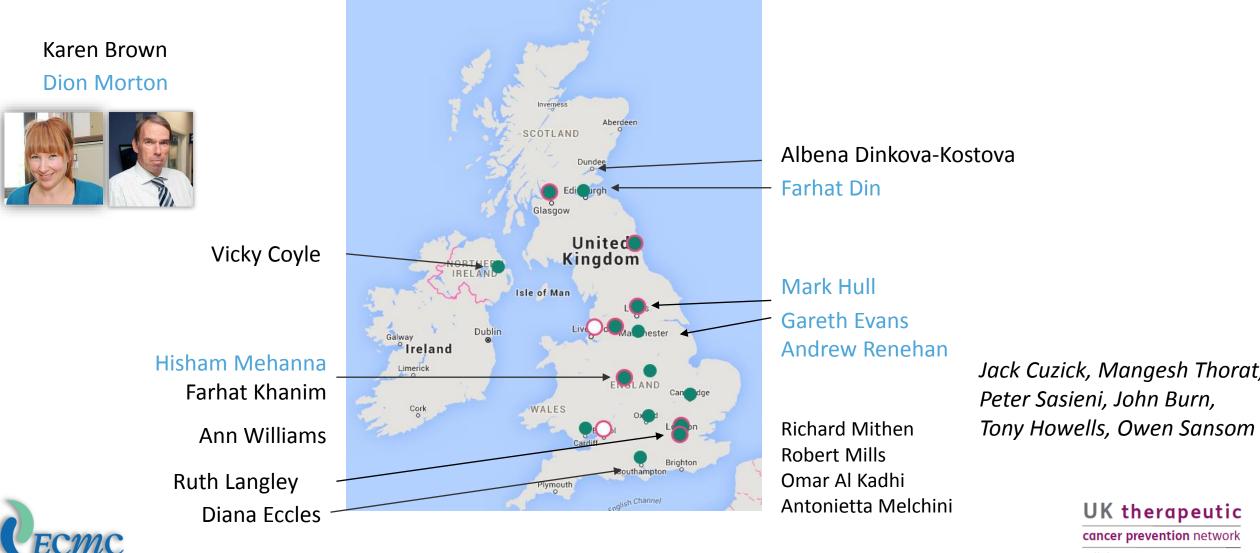
Provide infrastructure -organise scientific meetings focussing on therapeutic interventions cancer prevention

Increase awareness of the field amongst the cancer research community, government & funding bodies

> **UK therapeutic** cancer prevention network collaborating to prevent cancer



Steering committee



collaborating to prevent cancer

Comprehensive anatomical & therapeutic coverage

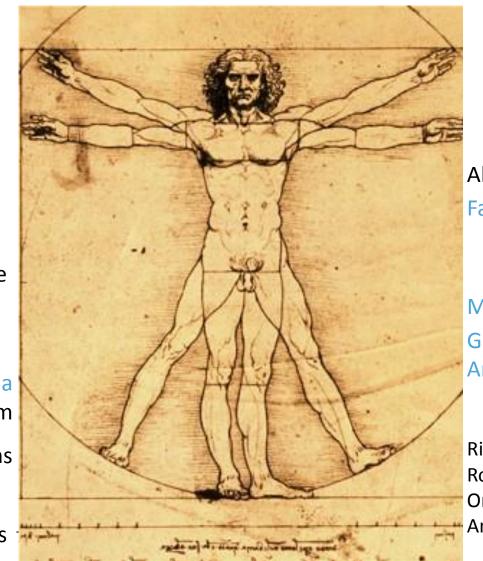
Karen Brown

Dion Morton



Vicky Coyle

Hisham Mehanna Farhat Khanim Ann Williams Ruth Langley Diana Eccles



Albena Dinkova-Kostova Farhat Din

Mark Hull Gareth Evans Andrew Renehan

Richard Mithen Robert Mills Omar Al Kadhi Antonietta Melchini Jack Cuzick, Mangesh Thorat, Peter Sasieni, John Burn, Tony Howells, Owen Sansom



UK therapeutic cancer prevention network

Patient representation is vital



Home Centres Groups

Sylvia Bailey

Maired MacKenzie







Role in early phase trials

Integration of efforts on prevention & treatment is necessary to control cancer epidemic Increased preneoplastic diagnosis → increase the burden of population requiring prevention

Therapeutic cancer prevention expanding role in cancer management

• secondary prevention.

Stratification of at risk populations using predictive biomarkers & precision medicine Evaluation of these therapies

 same skills
 MDTs as advanced disease CRUK & UKTCPN ECMC key role to conduct biomarker studies and develop the tools to target therapies for these patients



Laboratory Clinical Trials Biomarkers Biomarkers Basic Science Discovery Preclinical Phase I Phase II Phase III

Core areas of ECMC activity

Dose escalation of dietaryderived agent in volunteers to assess safety, tolerability, pharmacokinetics & metabolism Pre-surgical window trial cancer patients- target-tissue concentration & drug pharmacodynamic effects with repurposing potential

Dose-finding trial- effect of therapy on aberrant crypt foci, precursor to CRC Randomised trial - effect of a repurposed drug on CTdetected lung nodules, represent potential pre- or early invasive neoplasia Randomised biomarker trial of a new preventive combination in subjects with precursor lesions, such as breast intraepithelial neoplasia

2015-2016 UKTCPN meetings

Edinburgh Feb 2015

Vitamin D-Gene-Environment interactions in CRC- Malcolm Dunlop

Bristol Sept 2015

Inflammatory microenvironment in zebrafish models – Paul Martin Cancer epidemiology integrated research- Richard Mithen

Leeds Jan 2016

Off-label/licensing- Claire Potter, Medicines, Pharmacy & Industry group PPI strategy-Mark Hull







Prevention is better than cure – cancer chemoprevention in 2015 Tue 03-Nov-2015 15:40-17:40 Room 3B Host: Jack Cuzick

This session will cover several topical areas in chemoprevention research in 2015.

It will draw on examples spanning from 'bench to bedside' to highlight emerging areas, as well as a reflecting on lessons learnt and past achievements using the more traditional approaches to chemoprevention studies.

This session will delve into what advances in basic science, stratification and mouse-models can bring to the therapeutic cancer prevention studies of the future. It will address how epidemiological approaches to prevention studies can be joined up with new opportunities for sequencing cancer genomes and improved preclinical models, to identify candidate agents and biomarkers for chemoprevention, including repurposed drugs.

It will also examine how new strategies for personalised prevention can give rise to more cost-effective and focused approaches to research in chemoprevention that will ultimately bring benefits to patients faster.



Speakers

Andrew Chan, Massachusetts General Hospital, USA

Jack Cuzick, Queen Mary University of London, UK

Karen Brown, Leicester, UK

Panel

cer prevention network

Anthony Howell, University of Manchester, UK

ECMC

Farhat Khanim, University of Birmingham, UK

Ruth Langley, MRC Clinical Trials Unit at UCL, UK



Ilaria Mirabile @1ila · 3 Nov 2015 Panel discussion with speakers and members of #ECMC #UKTCPN #chemoprevetion #NCRI2015 ecmcnetwork.org.uk/network-groups...





Ilaria Mirabile @1ila · 3 Nov 2015 .@ECMC_UK at stand 57 for more info on the #UKTCPN ecmcnetwork.org.uk /network-groups... #NCRI2015

BCI @QMBCI

Excellent talk on setting up studies that fully consider physiology & pharmacokinetics in #prevention #NCRI2015 twitter.com/LeadingEdgeSci...

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Ilaria Mirabile @1ila · 3 Nov 2015 Can we exploit mechanisms to personalise #chemoprevention? Andrew Chan @MassGeneral #UKTCPN #NCRI2015

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Drug re-purposing

Co-ordinated by Karen Brown, Dion Morton, Farhat Khanim, Jack Cuzick, Vicky Coyle

• Screening of drugs available to the UK/EU market with the aim

of testing and re-purposing for a prevention setting

- Exploit expertise in drug repurposing and epidemiology/biostatistics
- Combined expertise in preclinical test systems and conducting window trials, using mechanism-based molecular endpoints as surrogate endpoints
- Across multiple cancer types
- Aim \rightarrow to advance drugs to <u>tertiary</u> prevention trials

Comprehensive ambitious programme covering several cancer types – difficult to conduct in other countries

UK therapeutic cancer prevention network



Drug re-purposing

Off Patent Drugs Bill

debated twice by House of Commons & yet to progress beyond Second Reading

- 12 January 2016
- Joint Assoc. of Medical Research Charities & Department of Health Roundtable event
- -develop a better shared understanding of issues raised by new uses for existing drugs
- UKTCPN representation (Karen Brown; Tony Howell)
- **Output prepared currently:**

Position paper -restricted access to off-patent drugs & therapeutic cancer prevention





Collaborative proposals in progress

Mechanisms of action aspirin grant proposal

-submitted to EME -Langely & Din

Small grants

James Thorne-Breast Cancer UK

Mangesh Thorat- CRUK Cancer Prevention Innovation Workshop Farhat Din- CRUK Cancer Prevention Innovation Workshop





2016-2017 Aims

Raise profile & standing of the network to a position where we are the first call nationally for advice/opinions on therapeutic prevention to support policy makers

Raise the profile of the UKTCPN and wider ECMC internationally by building links and collaborations with European researchers

To maximise public involvement in therapeutic cancer prevention discussion

Organise scientific conferences, symposia and workshops

Provide collaboration among investigators and develop joint grant applications





BACR-ECMC CONFERENCE



Deadline for abstract submission : June 1th Deadline for early bird registration : June 8th

http://www.bacr.org.uk/events/

BACR Chemoprevention Conference in Bristol. Great speakers packing out a fascinating day. #UKTCPN #chemoprevention

the way forward

July 18th – 19th 2016 University of Bristol School of Chemistry



OVERVIEW:

This multidisciplinary meeting will address the enormous opportunities and some of the challenges faced by researchers in the field of therapeutic cancer prevention. It will draw on examples spanning from 'bench to bedside' and beyond, to highlight what advances in basic and translational research can bring to the prevention trials of the future.

SESSIONS:

- · Dietary interventions for cancer prevention
- · Aspirin what else do we need to know?
- · Overcoming barriers to prevention and the way ahead
- Drug repurposing what next?
- New approaches and recent advances

SPEAKERS:

Bernardo Bonanni Farhat Khanim Milan, Italy John Burn Newcastle, UK Ruth Langley Rob Coleman Sheffield, UK **Richard Mithen** Jack Cuzick London, UK Andrea De Censi Genova, Italy Carlo Patrono Michael Pollack Farhat Din Edinburgh, UK Edinburgh, UK Malcolm Dunlop Peter Rothwell Owen Sansom Michelle Harvie Manchester, UK Mark Hull Leeds, UK

 Farhat Khanim
 Birmingham, UK

 Ruth Langley
 London, UK

 Richard Mithen
 Norwich, UK

 Chris Paraskeva
 Bristol, UK

 Carlo Patrono
 Rome, Italy

 Michael Pollack
 Montreal, Canada

 Peter Rothwell
 Oxford, UK

 Owen Sansom
 Glasgow, UK

Unique multidisciplinary team with National Coverage

Brings together scientists involved in research to **developing therapies** for **prevention / delay** development of malignancy

