PORTFOLIO NEWSLETTER

FEBRUARY 2021

NIHR University College London Hospital Clinical Research Facility Cancer Trials

INSIDE THIS ISSUE:

COVID-19 and Early Phase Cancer Trials at UCLH

During this difficult time we are trying to keep the early phase cancer trials program open to enrolment. The CRF is a geographically distinct unit to the main hospital and patients are treated in individual rooms as much as possible to minimize infection risk.

We therefore continue to welcome referrals; however due to the pressures at the front line, we are running at limited capacity and may not be able to take your patient immediately. Please liaise with the PI of the study you are considering referring to.

This newsletter provides you with details of the current ongoing cancer trials.

Dr. Rakesh Popat, Consultant Haematologist

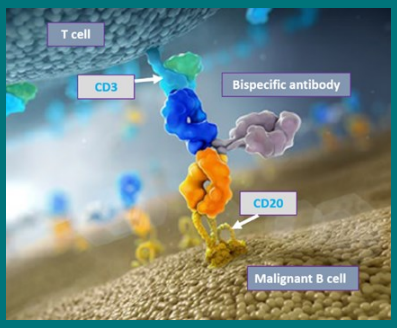
Cancer Lead NIHR UCLH Clinical Research Facility

Spotlight on Research

Consultant Haematologist & Honorary Senior Lecturer UCL

This month we will highlight research conducted by Dr. William Townsend.
William Townsend is a consultant haematologist specialising in lymphoma. He leads a programme of early phase clinical trials for patients with all types of lymphoma. Over the last 2 years he has focussed on bringing a new class of treatments called bi-specific antibodies to the CRF.

Bi-specific antibodies are a type of immunotherapy that enhance the immune-response against cancer by bringing T-cells of the patient's immune system into close contact with the malignant cells so leading to tumour cell death. It is proving a successful approach to treat lymphomas but with a range of side effects that need careful monitoring and expert management. Patients treated at the UCLH CRF were amongst the first in the country to receive bi-specific antibodies for their lymphoma. We currently have 5 trials using these drugs open with others in set-up:



NP40126: In this trial the CD3-20 bispecific antibody Glofitamab is combined with standard immunochemotherapy (R-CHOP). We are one of the leading recruiters for this trial globally. In part 1, this novel combination was tested in patients with relapsed follicular lymphoma. In part 2, patients with previously untreated diffuse large B cell lymphoma (DLBCL) will be treated.

NP39488: The CD3-20 bispecific antibody Glofitamab is combined with a PD1 inhibitor in this trial. The principle is to try to augment the immune response by combining these immunotherapy agents. Recruitment to this trial is currently paused but it will re-open soon with a novel imaging component in which we will study how T-cells traffic through the body in response to treatment. It recruits patients with relapsed or refractory B-NHL.

BP41072: This trial combines the CD3-20 bispecific antibody Glofitamab or an anti-CD20 antibody (obinutuzumab) with another novel drug that is designed to activate T cells (RO7227166, a CD-19 directed 41BB ligand). This exciting trial opened recently, we are one of only 2 centres in the UK recruiting to this protocol. This trial is open to patients with relapsed or refractory B-NHL.

CO41942: Opened in February 2021, this trial combines the CD3-CD20 bispecific antibody Mosunetuzumab with Lenalidomide in patients with relapsed follicular lymphoma. The aim is to identify if the two drugs act together to mount an effective immune response against the tumour.

NVG111: Due to open in February 2021, this first-in-human trial of a novel T-cell engaging drug that targets CD3 on T-cells and ROR1 on the surface of malignant cells is a success of the UCL-UCLH bench-to-bedside translational research programme. The study will recruit patients with CLL or Mantle Cell Lymphoma who have had an incomplete response to prior therapy.

It has taken a huge team effort to bring these trials to the CRF. I would like to thank the entire team who have helped to deliver this comprehensive portfolio of trials for our patients with lymphoma here at UCLH.
I am happy to be contacted directly to discuss potential patients for these or other trials.

Latest News

62nd American Society of Haematology (ASH) Annual Meeting and Exposition

Our Haematology PIs have co-authored a number of early phase clinical trial abstracts presented at the 62nd ASH annual congress in December 2020. This reflects excellent recruitment numbers and close working relationships with the trial sponsors. Dr Rakesh Popat has been involved in abstracts for the myeloma trials CC220, DREAMM-2 and DREAMM6 and Dr Willian Townsend has co-authored the BP41072 trial presentation for non-Hodgkin's lymphoma. We thank you for your on-going collaboration with referrals and shared care pathways.
Abstracts can be viewed at https://www.hematology.org/meetings/annual-meeting/abstracts.

How to refer a patient

By Post:
NIHR UCLH Clinical Research Facility,
University College London Hospitals NHS Foundation Trust, 4th Floor, 170 Tottenham Court Road,
London, W1T 7HA

By Telephone:
Reception: 020344 72929/72930
Reception Fax: 020344 72994

By Email*:
*When referring patients, ensure you use nhs.net emails. Patient identifiable data *should not* be sent to/via non-NHS email accounts.

Principal Investigator

Prof. John Bridgewater

Dr. Martin Forster

Prof. Daniel Hochhauser

Prof. Sam Janes

Dr. Mark Linch

Prof. Tim Meyer

Dr. Rowan Miller

Dr. Jenny O'Nions

Dr. Dionysis Papadatos-Pastos

Dr. Rakesh Popat

Dr. Rebecca Roylance

Dr. Beatrice Seddon

Dr. Heather Shaw

Dr. Sandra Strauss

Dr. William Townsend

Contents

As it appears in alphabetical order

UCLH Find a Study

To find more information on UCLH CRF Early Phase Cancer Clinical Trials visit the UCLH Find a Study database:

https://findastudy.uclh.nhs.uk/#/trial

Using the 'Study Name' and/or 'Local Project Reference' (LRP).

NIHR UCLH Clinical Research Facility
Cancer Trials Open to Recruitment

For more information on a trial, including eligibility criteria click the hyperlinked **Local Project Reference (LRP)** ID.

Breast

Study Acronym/ Full Title	LPR	PI	Drug Class/ Tumour Target
B-PRECISE-01 (MEN1611-01) Open-label, Multicentre, Phase Ib Dose-escalation Study of MENI6I 1, a P13K Inhibitor Combined with Trastuzumab ± Fulvestrant, in Subjects with PIK3CA Mutated HER2-positive Locally Recurrent Unresectable (advanced) or Metastatic Breast Cancer Progressed to Anti-HER2 Based Therapy.	120910	Dr. Rebecca Roylance	**P13K Inhibitor** PIK3CA Mutated HER2-positive Locally Recurrent Unresectable (advanced) or Metastatic Breast Cancer

Gynaecological

Study Acronym/ Full Title	LPR	PI	Drug Class/ Tumour Target
GCT1015-05 ENGOT innovaTV 205: A Phase 1b/2 Open-Label Trial of Tisotumab Vedotin (HuMax-TF-ADC) in Combination with Other Agents in Subjects with Recurrent or Stage IVB Cervical Cancer.	113569	Dr. Rowan Miller	**Antibody-Drug Conjugate (ADC) Targeting Tissue Factor (TF)** Recurrent or Stage IVB Cervical Cancer

Haematology

Study Acronym/ Full Title	LRP	PI	Drug Class/ Tumour Target
ASTX660 Phase 1-2 Study of the Safety, Pharmacokinetics, and Preliminary Activity of ASTX660 in Subjects with Advanced Solid Tumors and Lymphomas.	106548	Dr. William Townsend	**Dual Antagonist of Cellular Inhibitor of Apoptosis cIAP & XIAP** Relapsed T-Cell Lymphomas
BP41072 An Open-Label, Phase I Study To Evaluate The Safety, Pharmacokinetics And Preliminary Antitumor Activity Of Ro7227166 (A Cd19 Targeted 4-1bb Ligand) In Combination With Obinutuzumab And Incombination With Ro7082859 (Cd20-Tcb) Following A Pre-Treatment Dose Of Obinutuzumab Administered In Participants With Relapsed/ Refractory B-Cell Non-Hodgkin's Lymphoma.	129333	Dr. William Townsend	**Ro7227166 (CD19-directed 41BB ligand) + Obinutuzumab (anti-CD20 Monoclonal Antibody) or Glofitamab (CD3-CD20 T-Cell Bispecific Antibody)** Relapsed/Refractory B-Cell Non-Hodgkin's Lymphoma

Haematology (II)

Study Acronym/ Full Title	LRP	PI	Drug Class/ Tumour Target
CC-220-MM-001 A Phase 1B/2A Multicenter, Open-Label, Dose-Escalation Study to Determine the Maximum Tolerated Dose, Assess the Safety and Tolerability, Pharmacokinetics and Preliminary Efficacy of CC-220 Monotherapy and in Combination with Dexamethasone in Subjects with Relapsed and Refractory Multiple Myeloma.	16/0336	Dr. Rakesh Popat	**Cereblon E3 Ligase Modulator** Relapsed and Refractory Multiple Myeloma
CC-92480-MM-001 A Phase 1 Multicenter, Open-Label Study to Assess The Safety, Pharmacokinetics And Preliminary Efficacy of CC-92480 in Combination With Dexamethasone in Subjects With Relapsed And Refractory Multiple Myeloma.	18/0040	Dr. Rakesh Popat	**Cereblon E3 Ligase Modulator** Relapsed and Refractory Multiple Myeloma
CO41942 Ph1b study of bispecifics with lenalidomide in relapsed/ refractory FL.	130416	Dr. William Townsend	**CD3-CD20 Bispecific Antibody** Relapsed or Refractory Follicular Lymphoma
DREAMM-6 A Phase I/II, Open-label, Dose Escalation and Expansion Study to Evaluate Safety, Tolerability, and Clinical Activity of the Antibody-Drug Conjugate GSK2857916 Administered in Combination with Lenalidomide Plus Dexamethasone (Treatment Arm A), or Bortezomib Plus Dexamethasone (Treatment Arm B) in Participants with Relapsed or Refractory Multiple Myeloma.	18/0571	Dr. Rakesh Popat	**Humanised Monoclonal Antibody (anti-BCMA)** Relapsed or Refractory Multiple Myeloma
Teclistamab in Relapsed or Refractory Multiple Myeloma A Phase 1/2b, First-in-Human, Open-Label, Dose Escalation Study of Teclistamab, a Humanised BCMA x CD3 Bispecific Antibody, in Subjects with Relapsed or Refractory Multiple Myeloma.	134949	Dr. Rakesh Popat	**Humanised Bispecific Antibody (BCMA & CD3)** Relapsed or Refractory Multiple Myeloma

Hepatobiliary

Study Acronym/ Full Title	LPR	PI	Drug Class/ Target
ADP-0033-001 A Phase I Open Label Clinical Trial Evaluating the Safety and Anti-Tumor Activity of Autologous T Cells Expressing Enhanced TCRs Specific for Alpha Fetoprotein ($AFP^{c332}T$) in HLA -A2 Positive Subjects With Advanced Hepatocellular Carcinoma (HCC) or Other AFP Expressing Tumor Types.	17/0093	Prof. Tim Meyer	**T-Cell Therapy** Advanced Hepatocellular Carcinoma (HCC) or Other AFP Expressing Tumour Types

Lung

Study Acronym/ Full Title	LPR	PI	Drug Class/ Target
TACTICAL Targeted Stromal Cells Expressing TRAIL as a therapy for lung cancer.	14/0453	Prof. Sam Janes	**MSC-TRAIL with chemo-immunotherapy as first line therapy** Advanced Lung Cancer

Sarcoma

Study Acronym/ Full Title	LPR	PI	Drug Class/ Target
Afatinib in Chordoma A phase 2, single arm, European multi-center trial evaluating the efficacy of afatinib as first-line or later-line treatment in advanced Chordoma.	17/0146	Dr. Sandra Strauss	**EGFR Small Molecule Inhibitor** Advanced Chordoma
DCC-3014 A Multicenter Phase 1, Open-Label Study of DCC-3014 to Assess the Safety, Tolerability, Pharmacokinetics, and Pharmacodynamics in Patients with Advanced Tumors.	128373	Dr. Beatrice Seddon	**Tyrosine Kinase Inhibitor.** Cohort B only (Expansion Phase) — Tenosynovial Giant Cell Tumour (DTGCT)

Solid Tumours

Study Acronym/ Full Title	LPR	PI	Drug Class/ Target
AGI-134 A phase I/IIa, multicentre, open label study designed to evaluate the safety and tolerability of AGI-134 as monotherapy and in combination with pembrolizumab, in unresectable metastatic solid tumours.	104916	Dr. Martin Forster	**Anti-αGal** Unresectable Superficial Metastatic Melanoma or Squamous Cell Cancers
BLU-667 A Phase 1 Study of the Highly-selective RET Inhibitor, BLU-667, in Patients with Thyroid Cancer, Non-Small Cell Lung Cancer (NSCLC) and Other Advanced Solid Tumors.	17/0783	Dr. Martin Forster	**RET Inhibitor** Thyroid Cancer, NSCLC, Other RET-driven Advanced Solid Tumours
Cancer Peptide Vaccine Open-label, phase 1 study of S-488210/S-488211 to evaluate the safety and tolerability in patients with unresectable recurrent and/or metastatic solid tumor.	119614	Dr. Mark Linch	**5-peptide cancer vaccine** Unresectable recurrent and/or Metastatic Solid Tumour
D9170C00001 A Phase I/IIa, Open-Label Study to Assess the Safety, Tolerability, Pharmacokinetics and Preliminary Efficacy of Ascending Doses of AZD7648 Monotherapy or in Combination with either Cytotoxic Chemotherapies or Novel Anti-Cancer Agents in Patients with Advanced Malignancies	18/0580	Dr. Rowan Miller	**DNA-PK Inhibitor** Advanced Malignancies

Solid Tumours (II)

Study Acronym/ Full Title	LRP	PI	Drug Class/ Target
FIGHT-207 A Phase 2, Open-Label, Single-Arm, Multicenter Study to Evaluate the Efficacy and Safety of Pemigatinib in Participants With Previously Treated Locally Advanced/Metastatic or Surgically Unresectable Solid Tumor Malignancies Harboring Activating FGFR Mutations or Translocations.	124183	Prof. John Bridgewater	**FGFR Inhibitor** Advanced/Metastatic or Surgically Unrespectable Solid Tumour Malignancies Harbouring Activating FGFR Mutations or Translocations
Garnet A Phase 1 Dose Escalation and Cohort Expansion Study of TSR-042, an anti-PD-1 Monoclonal Antibody, in Patients with Advanced Solid Tumors.	16/0355	Dr. Rowan Miller	**Anti PD-L1 Monoclonal Antibody** Cohorts Open: *Part 2B: Cohort A1* dMMR/MSI-H endometrial cancer *Part 2B: Cohort F* non-endometrial dMMR/MSI-H & POLE-Mut cancers
LOXO-TRK-15002 (Navigate) LOXO-TRK-15002: A Phase II Basket Study of the Oral TRK Inhibitor LOXO-101 in Subjects with NTRK Fusion-Positive Tumors.	16/0077	Dr. Martin Forster	**TRK Inhibitor** NTRK Fusion-Positive Tumours
MOv18 A Cancer Research UK Phase I study of MOv18 IgE, a first in class chimeric IgE antibody against folate receptor-α, in patients with advanced solid tumours.	17/0121	Dr. Rowan Miller	**IgE Antibody against Folate Receptor-α** Advanced solid tumours
Starpharma CTX A phase 1/2 dose-escalation study to evaluate the safety, tolerability, pharmacokinetics and preliminary efficacy of CTX-SPL9111 (a cabazitaxel (CTX)-dendrimer conjugate) in patients with advanced solid tumours.	18/0016	Dr. Martin Forster	**Drug-Dendrimer Conjugate** Advanced solid tumours
Starpharma DTX A phase 1/2 dose-escalation study to evaluate the safety, tolerability, pharmacokinetics and preliminary efficacy of DTX-SPL8783 (a docetaxel (DTX)-dendrimer conjugate) as monotherapy in patients with advanced solid tumours or in combination with nintedanib in patients with non-small cell lung cancer (NSCLC).	17/0585	Dr. Martin Forster	**Drug-Dendrimer Conjugate (Tubulin Polymerase Inhibitor)** Advanced solid tumours Or non-small cell lung cancer (NSCLC)
TACTI-002 (Two ACTive Immunotherapeutics): A multicenter, open label, phase II study in patients with previously untreated unresectable or metastatic non-small cell lung cancer (NSCLC), or recurrent PD-X refractory NSCLC or with recurrent or metastatic squamous head and neck cancer (HNSCC) receiving the soluble LAG-3 fusion protein eftilagimod alpha (IMP321) in combination with pembrolizumab (PD-1 antagonist).	18/0560	Dr. Martin Forster	**APC Activator & Anti-PD-1 Monoclonal Antibody** Previously untreated unresectable or metastatic non-small cell lung cancer (NSCLC), Or recurrent PD-X refractory NSCLC Or with recurrent or metastatic squamous head and neck cancer (HNSCC)

Urology

Study Acronym/ Full Title	LRP	PI	Drug Class/ Target
BXCL701 A Phase 1b/2 Study of BXCL701, a Small Molecule Inhibitor of Dipeptidyl Peptidases (DPP), Administered in Combination with the Anti-Programmed Cell Death 1(PD-1) Monoclonal Antibody Pembrolizumab (PEMBRO; Keytruda®) in Patients with Small Cell Neuroendocrine Prostate Cancer (SCNC; NEPC).	113103	Dr. Mark Linch	**Dipeptidyl Peptidases (DPP) Inhibit & Anti-PD-1 Monoclonal Antibody** Small Cell Neuroendocrine Prostate Cancer (SCNC; NEPC)
CORT125281 Phase 1/2 Dose-Escalation and Expansion Study to Evaluate the Safety, Tolerability, and Pharmacokinetics of COR-T125281 with Enzalutamide in Patients with Metastatic Castration-Resistant Prostate Cancer.	18/0135	Dr. Mark Linch	**Glucocorticoid Receptor (GR) Antagonist** Metastatic Castration-Resistant Prostate Cancer
FIDES-02 An open-label multi-cohort Phase 1/2 study of derazantinib and atezolizumab in patients with urothelial cancer expressing activating molecular FGFR aberrations (FIDES-02)	124615	Dr. Mark Linch	**FGFR Inhibitor & Anti PD-L1 Monoclonal Antibody** Advanced Urothelial cancer expressing FGFR genetic aberrations. *Substudy 4:* Urothelial progression - FGFR-inhibitor resistant (>12 weeks) & previous chemo and CPI
PRO-MERIT (Prostate Cancer Messenger RNA Immunotherapy): A first-in-human, dose titration and expansion trial to evaluate safety, immunogenicity and preliminary efficacy of W_pro1 in patients with metastatic castration resistant prostate cancer and W_pro1 in combination with cemiplimab and/or goserelin acetate in patients with high-risk, localized prostate cancer.	122815	Dr. Mark Linch	**W_pro1 mRNA Cancer Vaccine** *Group 1:* mCRPC that have exhausted conventional treatment *Group 2:* High-risk, Localized Prostate Cancer prior to Prostatectomy