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De-ESCALaTE HPV: Determination of Epidermal growth factor receptor-inhibitor (cetuximab) versus Standard Chemotherapy (cisplatin) early And Late Toxicity Events in Human Papillomavirus-positive oropharyngeal squamous cell carcinoma

Submission date 20/02/2012	**Recruitment status** No longer recruiting	[X] Prospectively registered [ ] Protocol
Registration date 20/02/2012	**Overall study status** Completed	[ ] Statistical analysis plan [X] Results
Last Edited 29/10/2021	**Condition category** Cancer	[ ] Individual participant data

Plain English summary of protocol
http://www.cancerresearchuk.org/cancer-help/trials/a-trial-looking-side-effects-treatment-throat-cancer-de-escalate-hpv

Study website
http://www.warwick.ac.uk/go/deescalate

Contact information

Type(s)
Scientific

Contact name
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Additional identifiers

EudraCT/CTIS number
2011-005165-21

IRAS number

ClinicalTrials.gov number
NCT01874171

Secondary identifying numbers
11723

Study information

Scientific Title
De-ESCALaTE HPV: Determination of Epidermal growth factor receptor-inhibitor (cetuximab) versus Standard Chemotherapy (cisplatin) early And Late Toxicity Events in Human Papillomavirus-positive oropharyngeal squamous cell carcinoma: a randomised controlled trial

Acronym
De-ESCALaTE HPV

Study objectives
Oropharyngeal squamous cell carcinoma (OPSCC) incidence is increasing rapidly in the developed world. This has been attributed to a rise in Human Papillomavirus (HPV) infection. HPV+ OPSCC is considered a distinct disease entity, affecting younger patients and has a good prognosis following treatment. Subsequently, patients can live with the considerable side effects for several decades.

Radiotherapy and cetuximab have demonstrated similar efficacy to platin chemoradiotherapy in head and neck cancer, but is potentially less toxic.

Results of this trial will be used to determine the optimum treatment of this debilitating cancer, with the primary aim of decreasing toxicity and improving quality of life for HPV+OPSCC patients.

Ethics approval required
Old ethics approval format

Ethics approval(s)
NRES Committee West Midlands Coventry & Warwickshire, 29/11/2011, ref: 11/WM/0381

Study design
Both; Interventional; Design type: Treatment

Primary study design
Interventional

Secondary study design
Randomised controlled trial

Study setting(s)
Hospital

Study type(s)
Treatment

Participant information sheet
Not available in web format, please use the contact details below to request a patient information sheet

Health condition(s) or problem(s) studied
Topic: National Cancer Research Network; Subtopic: Head and Neck Cancer; Disease: Head and Neck

Interventions
HPV positive patients will be randomised to receive either cisplatin + radiotherapy (Arm A) or cetuximab + radiotherapy (Arm B) and will be followed up for two years. Patients that are HPV negative will enter into the Registration Cohort Study.

Cetuximab, Initial dose of 400mg/m2, administered intravenously, 1 week before start of radiotherapy followed by 7 weekly doses of 250mg/m2, administered intravenously. during radiotherapy.

Cisplatin, Three doses of cisplatin 100mg/m2, administered intravenously, on days 1, 22 and 43 of radiotherapy.

Intervention Type
Drug

Phase
Phase III

Drug/device/biological/vaccine name(s)
Cetuximab, cisplatin

Primary outcome measure
Current primary outcome as of 13/03/2012:
Severe toxicity (acute and late) (Grade 3-5); Timepoint(s): Two years from end of treatment

Previous primary outcome:
Severe toxicity (Grade 3-5); Timepoint(s): Two years from end of treatment

Secondary outcome measures
1. Acute severe toxicity; Timepoint(s): 3 months from end of treatment
2. Late severe toxicity; Timepoint(s): Two years from end of treatment

3. Quality of Life; Timepoint(s): Two years from end of treatment
4. Dysphagia; Timepoint(s): Two years from end of treatment
5. Cost effectiveness; Timepoint(s): Two years from end of treatment
6. Overall survival, recurrence and metastasis; Timepoint(s): Two years from end of treatment

Overall study start date
01/09/2011

Completion date
31/07/2019

Eligibility

Key inclusion criteria
Current inclusion criteria as of 13/03/2012:
1. Stage III-IVa oropharyngeal squamous cell tumours
2. Clinical multidisciplinary team decision to treat with primary curative chemoradiotherapy
3. No previous treatment for the primary tumour, including surgery, neck dissection or tracheostomy [except node biopsies or diagnostic tonsillectomy]
4. Medically fit Eastern Cooperative Oncology Group (ECOG) 0, 1 or 2
5. Adequate cardiovascular, haematological, renal and hepatic function
6. Age 18 years or over
7. Written informed consent given
8. Using adequate contraception [male and female participants]. Must take contraceptive measures during, and for at least three months after treatment.

Previous inclusion criteria:
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8. Using adequate contraception [male and female participants]

Participant type(s)
Patient

Age group
Adult

Lower age limit
18 Years

Sex
Both

Target number of participants

Planned Sample Size: 304; UK Sample Size: 304; Description: Depending on the number of participants recruited by international sites, the UK sample size may be smaller.

Total final enrolment
334

Key exclusion criteria
Current exclusion criteria as of 13/03/2012:
1. Distant metastasis (i.e. stage IVc disease)
2. Tumor, Node, Metastasis (TNM) Stage T1-2N0 disease
3. Treated with primary radical surgery to the primary site e.g. resection
4. Concurrent use of CYP3A4 inducers or inhibitors
5. Serious cardiac illness or other medical conditions precluding the use of cisplatin or cetuximab
6. HPV+ patients who have p16+ tumours who also have N2b, N2c or N3 nodal disease and who also smoke more than 10 pack years (i.e. have both risk factors)
7. Pregnant or lactating
8. Previous treatment for any other cancer with cytotoxics, radiotherapy or antiEGFR therapies
9. Inadequate renal, haematological or liver functions
10. Patients with clinically significant hearing impairment.
11. Life expectancy less than three months
12. Other malignancy within the past three years except basal cell skin cancer or preinvasive carcinoma of the cervix

Previous exclusion criteria:
1. Distant metastasis (i.e. stage IVc disease)
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9. Inadequate renal, haematological or liver functions
10. Life expectancy less than three months
11. Other malignancy within the past three years except basal cell skin cancer or preinvasive carcinoma of the cervix

Date of first enrolment
09/10/2012

Date of final enrolment
28/10/2016

Locations

Countries of recruitment
Belgium

England

Ireland

Italy

Netherlands

Scotland

United Kingdom

Wales

Study participating centre
Warwick Clinical Trials Unit
University of Warwick
Coventry
United Kingdom
CV4 7AL

Study participating centre
Aberdeen Royal Infirmary
Foresterhill Road
Aberdeen
United Kingdom
AB25 2ZN

Study participating centre
St Luke's Centre for Radiation at Beaumont Hospital
Beaumont Road
Dublin
Ireland
-

Study participating centre
Bradford Royal Infirmary
BIHR – Temple Bank House
Bradford Royal Infirmary
Bradford
United Kingdom
BD9 6RJ

Study participating centre

Bristol Haematology & Oncology Centre
Horfield Road
Bristol
United Kingdom
BS2 8ED

Study participating centre
Castle Hill Hospital
Castle Road
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HU16 5JQ

Study participating centre
Cheltenham General Hospital
Sandford Road
Cheltenham
United Kingdom
GL53 7AN

Study participating centre
Clatterbridge Cancer Centre
Clatterbridge Road
Bebington,
Wirral
United Kingdom
CH63 4JY

Study participating centre
Colchester General Hospital
Turner Road
Colchester
Essex
CO4 5JL
Colchester
United Kingdom
CO4 5JL

Study participating centre
Glan Clwyd Hospital
North Wales Cancer Treatment Centre

Glan Clwyd Hospital
Bodelwyddan
Denbighshire
Bodelwyddan
United Kingdom
LL18 5UJ

Study participating centre
James Cook University Hospital
Marton Road
Middlesbrough
United Kingdom
TS4 3BW

Study participating centre
Leicester Royal Infirmary
Infirmary Square
Leicester
United Kingdom
LE1 5WW

Study participating centre
Musgrove Park Hospital
The Beacon Centre
Taunton
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TA1 5DA

Study participating centre
New Cross Hospital
Deansley Centre
United Kingdom
WV10 9PQ

Study participating centre
Nottingham City Hospital
Hucknall Road
Nottingham
United Kingdom
NG5 1PB

Study participating centre
Queen Elizabeth Hospital Birmingham
Cancer Centre, Edgbaston
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B15 2TH

Study participating centre
Royal Derby Hospital
Uttoxeter Road
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DE22 3NE

Study participating centre
Royal Marsden Hospital (London)
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SW3 6JJ

Study participating centre
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Surrey
Sutton,
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Study participating centre
Royal Surrey County Hospital
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GU2 7XX

Study participating centre

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Combe Park
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Study participating centre
Singleton Hospital
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SA2 8QA

Study participating centre
St James's Institute of Oncology
Beckett Street
Leeds
United Kingdom
LS9 7TF

Study participating centre
St Luke’s Hospital
Highfield Road
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University Hospitals Coventry & Warwickshire
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Study participating centre
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Whitchurch

Cardiff
United Kingdom
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Study participating centre
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1081 HV

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Manor Hospital
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Western General Hospital
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EH4 2XU

Study participating centre
Weston Park Hospital
Whitham Road
Sheffield
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S10 2SJ

Study participating centre
Northampton General Hospital
Cliftonville
Northampton
United Kingdom
NN1 5BD

Study participating centre

Norfolk & Norwich University Hospital
Colney Lane
Norwich
United Kingdom
NR4 7UY

Sponsor information

Organisation
University of Warwick (UK)

Sponsor details
Warwick Medical School
Medical School Building
Coventry
England
United Kingdom
CV4 7AL

Sponsor type
University/education

ROR
https://ror.org/01a77tt86

Funder(s)

Funder type
Charity

Funder Name
Cancer Research UK (CRUK) (UK)

Alternative Name(s)
CR_UK, Cancer Research UK - London, CRUK

Funding Body Type
Private sector organisation

Funding Body Subtype
Other non-profit organizations

Location
United Kingdom

Results and Publications

Publication and dissemination plan
Planned publication in a high-impact peer reviewed journal.

Intention to publish date
31/12/2019

Individual participant data (IPD) sharing plan
The current data sharing plans for the current study are unknown and will be made available at a later date.

IPD sharing plan summary
Data sharing statement to be made available at a later date

Study outputs

Output type	Details	Date created	Date added	Peer reviewed?	Patient-facing?
Results article	results	05/01/2019		Yes	No
Plain English results		06/02/2019	29/10/2021	No	Yes
HRA research summary			28/06/2023	No	No