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Oral administration of S 78454 in combination with cisplatin in patients with advanced non-keratinising nasopharyngeal carcinoma

Submission date 19/04/2012	**Recruitment status** No longer recruiting	[_] Prospectively registered [_] Protocol [_] Statistical analysis plan [X] Results [_] Individual participant data
Registration date 08/06/2012	**Overall study status** Completed	
Last Edited 18/04/2018	**Condition category** Cancer	

Plain English Summary
Not provided at time of registration and not expected to be available in the future

Contact information

Type(s)
Scientific

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Additional identifiers

EudraCT/CTIS number

IRAS number

ClinicalTrials.gov number

Secondary identifying numbers
CL1-78454-009

Study information

Scientific Title
Phase I dose escalation study of oral administration of Pan-Histone Deacetylase (HDAC) inhibitor S 78454 given in combination with a fixed dose infusion of cisplatin in patients with advanced non-keratinising nasopharyngeal carcinoma.

Study hypothesis
Establish the safety and tolerability of S 78454 given in combination with a fixed dose infusion of cisplatin in patients with advanced non-keratinising nasopharyngeal carcinoma in terms of the maximum tolerated dose (MTD) and the dose-limiting toxicities (DLTs), and establish the recommended Phase II dose (RP2D).

Ethics approval required
Old ethics approval format

Ethics approval(s)
Ethics approval was obtained before recruitment of the first participants

Study design
Multicentre international non-randomised non-comparative open-label phase I study

Primary study design
Interventional

Secondary study design
Non randomised study

Study setting(s)
Hospital

Study type(s)
Screening

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

Condition
Advanced non-keratinising nasopharyngeal carcinoma

Interventions
S 78454 capsules 80 mg twice a day (b.i.d.) to 140mg b.i.d.

Cisplatin / 1 infusion per cycle / 75mg/m² maximum

1. At least 2 cycles of combination treatment.
2. Cisplatin limited to 6 cycles
3. S78454 as long as the disease does not progress and treatment sufficiently tolerated or consent withdrawal

Intervention Type
Drug

Phase
Phase I

Drug/device/biological/vaccine name(s)
Cisplatin

Primary outcome measure
1. MTD and DLTs of oral S 78454 capsules with a fixed dose infusion of cisplatin
2. Establish the recommended phase II dose

Secondary outcome measures
1. Safety profile (adverse events, laboratory tests, physical exam, ECOG, vital signs, ECG, clinical neurological examination, audiometric tests)
2. To determine pharmacokinetic (PK) profile
3. Measure Tumour response according to revised Response Evaluation Criteria In Solid Tumors (RECIST)_ and plasma Epstein-Barr Virus (EBV) DNA levels

Overall study start date
01/03/2012

Overall study end date
30/11/2013

Eligibility

Participant inclusion criteria
1. Male or female patients aged ≥21 (Singapore) ≥20 (Taiwan)
2. Histologically documented, measurable or evaluable advanced non-keratinising nasopharyngeal carcinoma, that has relapsed or is refractory to conventional, standard forms of therapy.
3. Ability to swallow oral capsule(s) without difficulty
4. Eastern Cooperative Oncology Group (ECOG) performance status ≤ 1
5. Estimated life expectancy > 12 weeks
6. Adequate haematological, renal and hepatic functions-Serum albumin 30 g/L
7. Written informed consent

Participant type(s)
Patient

Age group
Adult

Sex
Both

Target number of participants

30 patients in the escalation part + 10 patients in the confirmatory part = 40 patients

Participant exclusion criteria
1. Pregnant or breastfeeding women, women of childbearing potential or men without effective contraception
2. Involvement in another clinical trial at the same time or within 4 weeks prior to inclusion, or patient already enrolled in the study
3. Major surgery within previous 4 weeks
4. Chemotherapy within previous 3 weeks (6 weeks in case of nitroso-ureas)
5. Biologic/target therapy or immunologic agents within previous 3 weeks
6. Radiotherapy within previous 4 weeks (except for palliative radiotherapy at localised lesions)
7. Abnormal thyroid function (defined as thyroid-stimulating hormone or free T4) except for patients with hypothyroidism diagnosed prior to study entry and stable on thyroid replacement
8. Concurrent therapeutic anticoagulation by anti-vitamin K (AVK)
9. Uncontrolled diabetes mellitus
10. Concomitant uncontrolled infection or severe systemic disease
11. Symptomatic or progressive brain metastasis
12. Patients with pre-existing gastrointestinal disorders
13. Patient with impaired cardiac function
14. Prior exposure to any Histone deacetylase inhibitors (HDACi)
15. Known organ dysfunction
16. Peripheral neuropathy > grade 1
17. Hearing impairment/tinnitus > grade 2
18. Known hypersensitivity to cisplatin

Recruitment start date
01/03/2012

Recruitment end date
30/11/2013

Locations

Countries of recruitment
Singapore

Taiwan

Study participating centre
National University Cancer Institute
-
Singapore
119074

Sponsor information

Organisation
Institut de Recherches Internationales Servier (France)

Sponsor details
50 rue Carnot
Suresnes
France
92284

Sponsor type
Industry

Website
http://www.servier.com/

ROR
https://ror.org/034e7c066

Funder(s)

Funder type
Industry

Funder Name
Institut de Recherches Internationales Servier (France)

Results and Publications

Publication and dissemination plan
Publication plan:
Summary results are published in https://clinicaltrials.servier.com.

Intention to publish date

Individual participant data (IPD) sharing plan
The datasets generated during and/or analysed during the current study will be available upon request from https://clinicaltrials.servier.com if a Marketing Authorisation has been granted after 1st January 2014.

IPD sharing plan summary
Available on request

Study outputs

Output type	Details	Date created	Date added	Peer reviewed?	Patient-facing?
Basic results				No	No