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An open-label, multicenter, randomized Phase II Study to compare the effects of Paclitaxel /Carboplatin and Lonafarnib to those of Paclitaxel/Carboplatin for 1st line Treatment of patients with epithelial ovarian cancer International Federation of Gynecology and Obstetrics (FIGO) Stages IIB-IV

Submission date
12/07/2005

Recruitment status
No longer recruiting

Registration date
23/08/2005

Overall study status
Completed

Last Edited
02/07/2012

Condition category
Cancer

[X] Prospectively registered
[_] Protocol
[_] Statistical analysis plan
[X] Results
[_] Individual participant data

Plain English summary of protocol
Not provided at time of registration

Study website
http://www.ago-ovar.de

Contact information

Type(s)
Scientific

Contact name
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Ludwig-Erhard-Str. 100
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65199

Additional identifiers

EudraCT/CTIS number

IRAS number

ClinicalTrials.gov number

Secondary identifying numbers
AGO-OVAR 15

Study information

Scientific Title

Acronym
AGO-OVAR 15

Study objectives
Standard chemotherapy for ovarian cancer patients after primary cytoreductive surgery is paclitaxel in combination with carboplatin. Several phase III studies are evaluating the efficacy of a third drug within this standard trial either as a combined or as a consolidation therapy. The final results of these studies have not yet been published. The addition of farnesyltransferase (FT) inhibitors or epidermal growth factor inhibitors to primary chemotherapy are very promising approaches to optimize primary therapy. Lonafarnib is a FT inhibitor that is active against a broad spectrum of tumor cell lines in vitro and tumor xenografts in nude mice. Lonafarnib has single agent antitumor activity as well as enhanced activity in combination with taxanes in a number of tumor cell lines and mice models. Based upon positive results from clinical studies demonstrating enhanced activity when combining taxanes with lonafarnib, combination therapy of paclitaxel and carboplatin with lonafarnib is expected to have greater efficacy than standard therapy or FTI therapy alone in primary ovarian cancer patients.

Ethics approval required
Old ethics approval format

Ethics approval(s)
Not provided at time of registration

Study design
Randomised controlled trial

Primary study design
Interventional

Secondary study design
Multi-centre

Study setting(s)
Not specified

Study type(s)

Treatment

Participant information sheet

Health condition(s) or problem(s) studied
Epithelial Ovarian Cancer, First-Line Treatment

Interventions
Paclitaxel/Carboplatin +/- Lonafarnib

The previous sponsor for this trial (until November 2009) was:
MedServ. GmbH (Germany)
Ludwig-Erhard-Str. 100,
65199 Wiesbaden
Germany

Intervention Type
Drug

Phase
Phase II

Drug/device/biological/vaccine name(s)
Paclitaxel/Carboplatin and Lonafarnib

Primary outcome measure
Comparison of the effect (progression-free survival [PFS]) of paclitaxel/carboplatin and lonafarnib to that of paclitaxel/carboplatin in patients with previously untreated epithelial cancer of the ovary FIGO stages IIBIV. The primary purpose of this study is to determine whether the additional effect of lonafarnib is sufficient to conduct a phase III study.

Secondary outcome measures
The secondary objectives are to evaluate response to treatment and overall survival, and to assess the safety in both treatment arms and to assess exposure (PK) and PD of lonafarnib.

Overall study start date
01/09/2005

Completion date
30/09/2006

Eligibility

Key inclusion criteria
1. Previously untreated patients with a histologically confirmed diagnosis of cancer of the ovary or the fallopian tube or extra-ovarian papillary serous tumors
FIGO stage IIBIV regardless of measurable or non-measurable disease
2. Age ≥18 years
3. Eastern Cooperative Oncology Group (ECOG) performance status ≤2
4. Life-expectancy of at least 6 months

5. Adequate bone marrow, renal and hepatic function defined as
white blood cell count (WBC) >3.0/nl, Neutrophils (ANC) ≥1.5/nl, Platelets ≥100/nl, Hemoglobin >6 mmol/l (>10.0 g/dl), Bilirubin ≤1 x upper limit of normal range
6. Aspartate aminotransferase (AST) and alanine aminotransferase (ALT) <1.5 x upper limit of normal range
7. Alkaline phosphatase <2.5 x upper limit of normal range
8. Estimated glomerular filtration rate GFR ≥50 ml/min according to Jelliffe or Cockroft-Gault formula
9. Patients who have given their signed and written informed consent to participate in the trial after fully understanding the implication and constraints of the protocol
10. Patients must be geographically accessible for treatment and follow-up
11. Time between definitive surgery and randomization ≥6 weeks

Participant type(s)
Patient

Age group
Adult

Lower age limit
18 Years

Sex
Both

Target number of participants
100

Key exclusion criteria
1. Ovarian tumors of low malignant potential (borderline tumors
2. Non-epithelial ovarian or mixed epithelial/non epithelial tumors (e.g. mixed Mullerian tumors)
3. Patients who have received previous chemotherapy or radiotherapy
4. Prior treatment with FT inhibitors
5. Patients with a prior diagnosis of any malignancy not cured by surgery alone less than 5 years before study entry (except in situ carcinoma of the cervix or adequately treated basal cell carcinoma of the skin)
6. Complete bowel obstruction or the presence of symptomatic brain metastases
7. Concurrent severe medical problems unrelated to malignancy which would significantly limit full compliance with the study or expose the patient to extreme risk or decreased life expectancy
8. Patients with a history of seizure disorder or central nervous system disorders
9. Pre-existing motor or sensory neurologic pathology or symptoms >National Cancer Institute - Common Toxicity Criteria (NCI-CTC) grade 1
10. History of congestive heart failure (New York Heart Association [NYHA] Classification >2), even if medically controlled
11. History of clinical and electrocardiographically documented myocardial infarction within the last 6 months
12. History of atrial or ventricular arrhythmias (≥LOWN II)
13. Patients with significant Fridericia QTc (QTcF) prolongation at Baseline (i.e. QTcF >470 msec)
14. Patients with severe active infection
15. Patients with a history of severe hypersensitivity reactions to products containing Cremophor EL (cyclosporin or vitamin K) and/or patients with known hypersensitivity to

compounds chemically related to Carboplatin and Paclitaxel
16. Fertile women not using adequate contraceptive methods
17. Women who are pregnant or breast feeding
18. Administration of other anticancer therapy or simultaneous chemotherapeutic and/or hormonal drugs, or radiotherapy during the study treatment period (except: hormonal replacement therapy and/or steroid antiemetics)
19. Patients who have used any investigational drugs within 30 days of study entry
20. Patients who are participating in any other clinical study
21. Dementia or significantly altered mental status that would prohibit the understanding and giving of informed consent

Date of first enrolment
01/09/2005

Date of final enrolment
30/09/2006

Locations

Countries of recruitment
Germany

Study participating centre
Ludwig-Erhard-Str. 100
Wiesbaden
Germany
65199

Sponsor information

Organisation
AGO Research GmbH (Germany)

Sponsor details
Kaiser-Friedrich-Ring 71
Wiesbaden
Germany
65185

Sponsor type
Industry

Website
http://www.ago-ovar.de

ROR
https://ror.org/01jdhsq12

Funder(s)

Funder type
Charity

Funder Name
AGO Ovarian Cancer Study Group (AGO-OVAR)

Results and Publications

Publication and dissemination plan
Not provided at time of registration

Intention to publish date

Individual participant data (IPD) sharing plan

IPD sharing plan summary
Not provided at time of registration

Study outputs

Output type	Details	Date created	Date added	Peer reviewed?	Patient-facing?
Results article	results	01/08/2012		Yes	No