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Three arm randomized parallel phase II/III study evaluating the efficacy and safety of the combinations Epirubicin and Taxotere® (ET), Taxotere® and Navelbine® (TN) and Navelbine® and Epirubicin (EN) as first line therapy in patients with metastatic breast cancer

Submission date 27/01/2006	**Recruitment status** No longer recruiting	[_] Prospectively registered [_] Protocol
Registration date 27/01/2006	**Overall study status** Completed	[_] Statistical analysis plan [_] Results
Last Edited 03/07/2009	**Condition category** Cancer	[_] Individual participant data [_] Record updated in last year

Plain English summary of protocol
Not provided at time of registration

Contact information

Type(s)
Scientific

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Additional identifiers

EudraCT/CTIS number

IRAS number

ClinicalTrials.gov number

Secondary identifying numbers
NTR472

Study information

Scientific Title

Acronym
ETN study

Study objectives
Primary objectives: To assess the efficacy in terms of response rate of the combinations Epirubicin and Taxotere (ET), Taxotere and Navelbine (TN) and Navelbine and Epiribicin (EN).
Secondary objectives: To determine progression free survival and toxicity profiles.

Ethics approval required
Old ethics approval format

Ethics approval(s)
Received from the local media ethics committee

Study design
multicentre, randomised, active controlled, parallel group study

Primary study design
Interventional

Secondary study design
Randomised controlled trial

Study setting(s)
Hospital

Study type(s)
Treatment

Participant information sheet

Health condition(s) or problem(s) studied
Breast cancer

Interventions

Arm A: Epirubicin 75 mg/m2, day 1 and docetaxel 60 mg/m2 day 1
Arm B: Vinorelbine 20 mg/m2 day 1 + 8 and docetaxel 60 mg/m2 day 8 (closed January 2003)
Arm C: Epirubicin 75 mg/m2 day 1 and vinorelbine 25 mg/m2 days 1 and 8.
One course consists of 21 days. Cycle is repeated every 3 weeks, for a maximum of 6 cycles.

Intervention Type
Drug

Phase
Phase II/III

Drug/device/biological/vaccine name(s)
epirubicin, docetaxel (Taxotere®), vinorelbine (Navelbine®)

Primary outcome measure
1. Time to progression
2. Response rate

Secondary outcome measures
1. Toxicity profile
2. Feasibility

Overall study start date
01/04/2001

Completion date
01/12/2005

Eligibility

Key inclusion criteria
1. Histologically proven breast cancer at first diagnosis. At study entry histological or cytological proof of metastasis is required in case of a single metastatic target lesion.
2. Female metastatic breast cancer patients
3. Measurable disease or evaluable disease (bone metastases only allowed)
4. Previous chemotherapy: Adjuvant: Patients may have had adjuvant and/or neoadjuvant chemotherapy but no more than 240 mg/m2 cumulative dose of prior doxorubicin or no more than 450 mg/m2 of epirubicin. Taxanes in adjuvant setting are allowed. However, there must be at least 12 months interval between the end of (neo-)adjuvant chemotherapy and protocol entry. This interval is not required for patients who received non-anthracycline/non-taxane adjuvant and/or neoadjuvant chemotherapy. No previous chemotherapy for metastatic breast cancer is allowed.
5. Previous hormonal treatment: Previous hormonal treatment is allowed provided discontinuation >4 weeks before start of study treatment
6. Previous radiation: Previous radiation therapy may have been given provided it is not the only site to assess response
7. Age >18 and <70 years
8. WHO performance status 0, 1 or 2
9. Laboratory requirements:

a. Hematology: White blood cell count >3.0 x 10^9/l (if WBC <3.0 x 10^9/l, Neutrophils should be >1.5 x 10^9/l), Platelets >100 x 10^9/l, Hemoglobin >10 g/dl (>6.2 mmol/l)
b. Hepatic function: Total bilirubin <1.00 times the upper-normal limits (UNL) of the institutional normal values. ASAT (SGOT) and/or ALAT (SGPT) <2.5 UNL, alkaline phosphatase <5 UNL (unless bone metastasis are present in the absence of any liver disorders). NB: Patients with ASAT and /or ALAT >1.5 UNL associated with alkaline phosphatase >2.5 UNL are not eligible for study.
c. Renal function: Serum creatinine <80 umol/l. If serum creatinine >80 umol/l, calculated creatinine clearance (Cockroft Gould) should be >60 ml/min
10. Normal left ventricular ejection fraction (LVEF) or superior to the lower limits of the institution (determined by either MUGA scan or ultrasound methods)
11. Patients must be accessible for treatment and follow-up
12. Measurability of the disease and evaluation of response according to RECIST criteria
13. Complete initial work-up within 3 weeks prior to first infusion
14. Written informed consent

Participant type(s)
Patient

Age group
Adult

Lower age limit
18 Years

Sex
Female

Target number of participants
111

Key exclusion criteria
1. Prior chemotherapy for metastatic disease
2. Locally advanced inoperable breast cancer (Stage III B) as only manifestation of the disease
3. Non-measurable disease
4. Pregnant or lactating women or women of childbearing potential not using adequate contraception
5. History of prior malignancies (other than non melanoma skin cancer or excised cervical carcinoma in situ)
6. Clinical evidence of cerebral metastasis
7. Symptomatic peripheral neuropathy > grade 2 according to the NCI Common Toxicity Criteria
8. WHO PS >2
9. Concurrent treatment with other experimental drugs
10. Participation in another clinical trial with any investigational drug within 30 days prior to study screening
11. Concurrent treatment with any other anti-cancer therapy except for concomitant treatment with bisphosphonates, provided that bone metastases are not the only evaluable lesions for response to therapy (see measurability of disease and evaluation of response)

Date of first enrolment
01/04/2001

Date of final enrolment
01/12/2005

Locations

Countries of recruitment
Netherlands

Study participating centre
VU University Medical Center
Amsterdam
Netherlands
1007 MB

Sponsor information

Organisation
VU University Medical Center (Netherlands)

Sponsor details
Van der Boechorststraat 7
Amsterdam
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1081 BT

Sponsor type
Not defined

ROR
https://ror.org/00q6h8f30

Funder(s)

Funder type
Industry

Funder Name
Amgen (Netherlands)

Alternative Name(s)
Amgen Inc., Applied Molecular Genetics Inc.

Funding Body Type
Government organisation

Funding Body Subtype
For-profit companies (industry)

Location
United States of America

Funder Name
Pfizer (Netherlands)

Alternative Name(s)
Pfizer Inc., Pfizer Consumer Healthcare, Davis, Charles Pfizer & Company, Warner-Lambert, King Pharmaceuticals, Wyeth Pharmaceuticals, Seagen

Funding Body Type
Government organisation

Funding Body Subtype
For-profit companies (industry)

Location
United States of America

Funder Name
Pierre Fabre (France)

Funder Name
Sanofi-Aventis (France)

Funder Name
VU University Medical Center (Netherlands)

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