

**GBG**

<b>Country:</b>	Germany
<b>Group:</b>	German Breast Group (GBG)
<b>Chairs:</b>	<p>Professor Dr med. G. von Minckwitz J.W. Goethe University Frankfurt Department of Obstetrics and Gynaecology Theodor-Stern-Kai 7 60590 Frankfurt GERMANY Tel: +49 69 6301 7024 Fax: +49 69 6301 7938 Email: minckwitz@em.uni-frankfurt.de</p> <p>Professor Dr med. M. Kaufmann J.W. Goethe University Frankfurt Department of Obstetrics and Gynaecology Theodor-Stern-Kai 7 D-60590 Frankfurt GERMANY Tel: +49 69 6301 5115 Fax: +49 69 6301 4717 Email: kaufmann@em.uni-frankfurt.de</p>
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**Title:** A multi-center randomized Phase III study evaluating 4 cycles of docetaxel, doxorubicin and cyclophosphamide (TAC) *versus* 4 cycles of vinorelbine and capecitabine (NX) in patients not sufficiently responding to 2 cycles of TAC and 4 cycles of TAC *versus* 6 cycles of TAC in patients sufficiently responding to 2 cycles of TAC as preoperative treatment of locally advanced (T4 a-d, N0-3,M0) or operable ( $T \geq 2$  cm, N0-2,M0) primary breast cancer/GBG 24.

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**Summary:** *Design:*

Prospective, randomized Phase III trial including an internal Phase II trial opened in September 2001, Phase III trial opened in July 2002

*Study Population:*

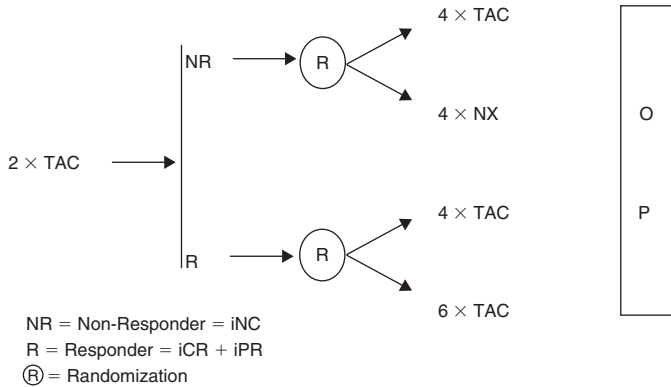
Locally advanced (T4 a-d, N0-3,M0) or operable ( $T \geq 2$  cm, N0-2,M0) primary breast cancer

**Primary Objectives:**

- To determine the pCR rate of 4 cycles of TAC and of 4 cycles of NX (TAC *versus* NX) as a salvage treatment in patients not sufficiently responding (i.e. cNC) to 2 cycles of TAC as preoperative treatment of operable ( $T \geq 2$  cm, N0-2, M0) primary breast cancer.
- To determine the pCR rate of 6 cycles *versus* 8 cycles of docetaxel, doxorubicin and cyclophosphamide (TAC  $\times$  6 *versus* TAC  $\times$  8) in patients sufficiently responding (i.e. cPR or cCR) to the first 2 cycles of TAC as preoperative treatment of operable ( $T \geq 2$  cm, N0-2, M0) primary breast cancer.

**Scheme:**

Gepartrio Design



**Update:**

- 2104 patients enrolled until enrolment ended in May 2005.

**Related Publications:**

von Minckwitz G, Blohmer JU, Raab G, *et al.* German Breast Group. *In vivo* chemosensitivity-adapted preoperative chemotherapy in patients with early-stage breast cancer: the GEPARTRIO pilot study. *Ann Oncol* 2005; 16(1): 56–63.

**Topics:**

- Anthracyclines
- Capecitabine
- Locally advanced breast cancer
- Taxanes
- Vinorelbine

**Keywords:**

Preoperative chemotherapy, breast cancer

**Title:** A multi-center randomized Phase III study to compare capecitabine alone or in combination with trastuzumab in patients with HER-2 positive metastatic breast cancer and progression after previous treatment with trastuzumab (Treatment Beyond Progression, TBP).  
**BIG 3-05 – GBG 26**

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**Summary:** *Design:*

Prospective, randomized Phase III trial

*Study Population:*

Patients with HER-2 positive metastatic breast cancer and progression during or after previous chemotherapy and trastuzumab treatment as follows:

- Taxanes + trastuzumab given as adjuvant therapy
- Taxanes + trastuzumab given as first line therapy for palliation
- Trastuzumab given as first line therapy for palliation alone or in combination with chemotherapeutic agents other than capecitabine or taxanes

- Trastuzumab has to be given previously for at least 12 weeks, treatment-free interval of trastuzumab for a maximum of 6 weeks. No more than one chemotherapy for palliation

*Aims:*

The primary aim is to compare the time to disease progression in patients with HER 2 positive metastatic breast cancer and progression after previous treatment with trastuzumab randomized to capecitabine alone or in combination with trastuzumab.

*Secondary Aims/Endpoints:*

- To compare the objective response rate between the two arms.
- To compare the duration of response.
- To compare the clinical benefit defined as CR, PR, or stable disease >24 weeks between the two arms.
- To evaluate the safety of the capecitabine + trastuzumab combination.
- To compare overall survival between the two arms.

*Treatment:*

- Capecitabine 2500 mg/m<sup>2</sup> orally day 1–14 q day 22 until progression or unacceptable toxicity, patient's request or withdrawal from study, and discontinuation of Trastuzumab.
- Capecitabine and Trastuzumab Capecitabine 2500 mg/m<sup>2</sup> orally day 1–14 q day 22 until progression or unacceptable toxicity, patient's request or withdrawal from study.
- Trastuzumab 6 mg/kg body weight every 3 weeks intravenous (i.v.) as a 90 minutes infusion until progression or unacceptable toxicity, patient's request or withdrawal from study.

**Scheme:** None available

**Update:**

- 135 enrolled as of September 2006.

**Related Publications:** None available

**Topics:**

- Capecitabine
- HER-2 positive patients
- Metastatic breast cancer
- Trastuzumab

**Keywords:** Trastuzumab beyond progression, HER-2 positive breast cancer

**Title:** A Phase III multi-centre double blind randomized trial of celecoxib *versus* placebo in primary breast cancer patients.  
**BIG 1-03 – ICCG / C/20/01 – GBG 27**  
 (see also description under ICCG)

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**Coordinator(s):** *Management:*

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Dr P. Hupperets, Maastricht (ICCG)

Professor Dr G. von Minckwitz, Frankfurt (GBG)

International Collaborative Cancer Group (ICCG) and the German Breast Group (GBG) Intergroup Study

**Summary:**

- A multi-centre, Phase III, placebo controlled randomized trial. Patients are randomized between 2 years celecoxib and placebo in a 2:1 ratio in favour of celecoxib

*Chemotherapy:*

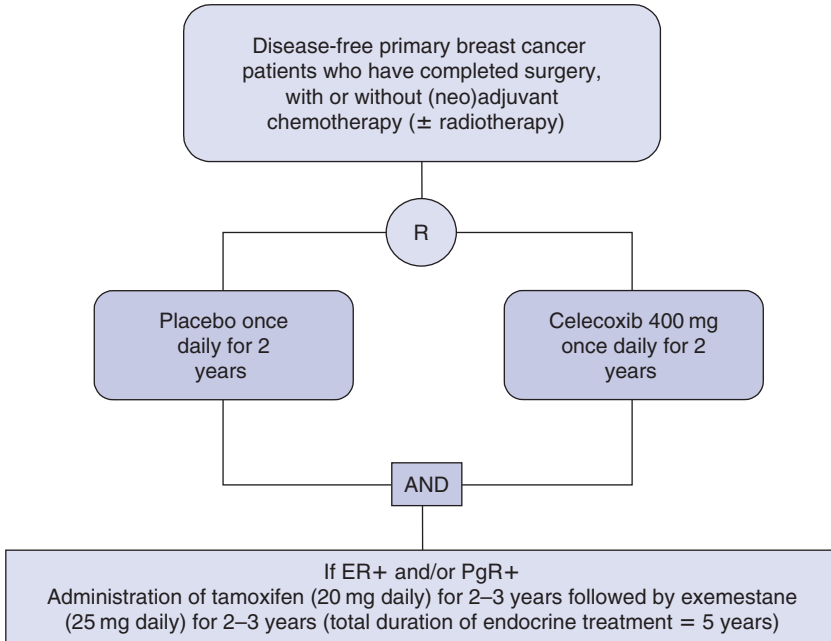
Prior to randomisation, patients may have received chemotherapy (all HR negative patients **MUST** have received chemotherapy). All patients who have received chemotherapy should have finished their treatment, which will have consisted of at least 4 cycles. The schedule of preference is FEC 3-weekly for 6 courses, however, other dose schedules of FEC or FAC plus combinations that contain EC/AC followed by a taxane, or Epirubicin/Doxorubicin plus a taxane are permitted. CMF may be substituted for patients where Epirubicin is contraindicated.

*Aims:*

The primary aim is to assess the disease-free survival (DFS) benefit of 2 years adjuvant therapy with the COX-2 inhibitor celecoxib compared with placebo in primary breast cancer patients.

*Secondary Aims/Endpoints:*

- To compare overall survival.
- To define the safety of adjuvant therapy with celecoxib in this patient population.
- To compare the incidence of second primary cancers.
- In postmenopausal hormone receptor (HR) positive patients, to assess tolerability of celecoxib with tamoxifen.
- To assess DFS benefit of 2 years adjuvant celecoxib compared with placebo in HR positive (i.e. ER positive and/or PR positive) and in HR negative (i.e. ER negative/PR negative) disease.

**Scheme:**

**Update:**

- Planned study start: November 2006.

**Related Publications:** None available

**Topics:**

- Aromatase inhibitors
- Celecoxib
- Tamoxifen

**Keywords:** Breast cancer, aromatase inhibitors, celecoxib, tamoxifen



**Title:** Prospective Register Study of the German Adjuvant Breast Cancer Study Group (GABG) or Diagnosis and Treatment of Breast Cancer in Pregnancy / BIG 2–03, GBG 29.

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Funding guaranteed by the BANSS-Foundation.

**Summary:**

- Breast cancer complicating pregnancy is a rare coexistence. Breast cancer is after the age of 25 the most common malignancy in pregnancy. Little is known about the incidence in Germany and Western Europe. We therefore want to initiate a trial collecting prospectively as much data as possible for the diagnostic, treatment, and maternal and foetal outcome during pregnancy. The primary endpoint is the foetal outcome 4 weeks after delivery. Secondary endpoints will include maternal outcome, pregnancy outcome, diagnostic procedures used and the biology of the tumour. A flow sheet for the treatment is given and the acceptance of these guidelines will be evaluated.

*Primary Endpoint:*

- Foetal outcome 4 weeks after delivery.

*Secondary Endpoints:*

- Maternal outcome of pregnancy.
- Stage of and biological characteristics of breast cancer.

- Breast cancer therapy (treatment, response to chemotherapy, type of surgery).
- Sensitivity and specificity of diagnostic procedures (palpation, US, mammogram).
- Outcome of the newborn after 5 years of therapy.
- Outcome of breast cancer 5 years after diagnosis.

**Scheme:** Prospective and retrospective registration of patients who have been diagnosed with breast cancer during pregnancy

**Update:** • Accrual as of March 2006: 61.

**Related Publications:** Loibl, *et al.* Breast cancer during pregnancy – recommendations from an international expert panel. *Cancer* 2006; 106: 237–246.

**Topics:** • Premenopausal patients

**Keywords:** Breast cancer, pregnancy, foetal outcome

**Title:** Ibandronate with or without capecitabine in elderly patients with early breast cancer – ICE – Study / BIG 4-04, GBG 32. (see also description under WSG)

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**Summary:**

- Prospective, multi-center, controlled, open-label, randomized Phase III

*Study Population:*

Age  $\geq 65$  years, N+ or N– (high risk: T  $\geq 2$  cm or G II/III or receptor negative)

*Primary Objective:*

- To compare the event-free survival in elderly patients after local treatment for primary breast cancer treated with either ibandronate alone or ibandronate and capecitabine as adjuvant treatment.

*Secondary Objectives:*

- To compare the overall survival between the two arms.
- To determine the compliance in both arms.

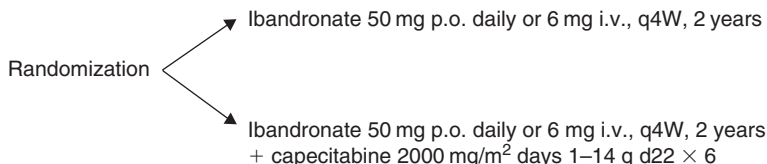
- To determine the toxicity in both arms.
- To determine the rate of bone-related events in hormone-sensitive and insensitive disease (with or without anastrozole).
- To determine the preference to oral or intravenous application of ibandronate.
- To assess quality of life.
- To compare a geriatric assessment by Charlson *versus* VES 13 score.

*Tertiary Objectives:*

- To determine prognostic factors on tumour tissue collected from primary surgery and to correlate them with study treatment effect.
- To evaluate the prognostic impact of age, serum albumin, hemoglobin level, creatinine clearance, Charlson score, VES-score in a multivariate analysis for the prediction of treatment associated adverse events and limited life time expectancy.

**Scheme:**

Age  $\geq$  65 years, N+ or N- (high risk: T  $\geq$  2 cm or G II/III or receptor negative)



If ER and/or PR  $\oplus$ : Anastrozole 1 mg p.o. daily 5 years (in sequence to capecitabine)

**Update:**

- Planned patient No: 1397
- Accrual as of March 2006: 580.

**Related Publications:**

O'Shaughnessy J, Miles D, Vukelja S, *et al.* Superior survival with capecitabine plus docetaxel combination therapy in anthracycline-pretreated patients with advanced breast cancer: phase III trial results. *J Clin Oncol* 2002; 20: 2812–2823.

O'Shaughnessy JA, Blum J, Moiseyenko V, *et al.* Randomized, open-label, phase II trial of oral capecitabine (Xeloda) *versus* a reference arm of intravenous CMF (Cyclophosphamide, methotrexate and 5-fluorouracil) *Ann Oncol* 2001; 12: 1247–1254.

**Topics:**

- Bisphosphonates
- Capecitabine
- Elderly patients

**Keywords:**

Elderly patients, chemotherapy, bisphosphonates, breast cancer

**Title:** GAIN: German Adjuvant Intergroup Node-positive Study.  
A Phase III trial to compare ETC *versus* EC-TX and ibandronate *versus* observation in patients with node-positive primary breast cancer / GBG 33.

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**Summary:**

- A German Intergroup Study of the Arbeitsgemeinschaft Gynäkologische Onkologie (AGO) German Breast Group (GBG) and Nordostdeutsche Gesellschaft für Gynäkologische Onkologie (NOGGO)

*Design:*

Prospective, multi-center, controlled, non-blinded, randomized Phase III with a 2 × 2 factorial design

*Primary Objectives:*

- To compare the disease-free survival after adjuvant chemotherapy with “ETC” (Arm A1) or “EC-TX” (Arm A2) in patients with primary node-positive breast cancer.
- To compare the disease-free survival with (Arm B1) or without ibandronate (Arm B2) treatment for 2 years in patients with primary node-positive breast cancer.

*Secondary Objectives:*

- To compare overall survival between Arms A1 *versus* A2 and B1 *versus* B2.
- To evaluate the compliance in Arms A1 *versus* A2 and in B1.
- To compare the safety between Arms A1 *versus* A2 and B1 *versus* B2.
- To assess the rate of responders to erythropoiesis stimulating factors in Arms A1 and A2.
- To compare the incidence of secondary primaries between Arms A1 *versus* A2.
- To compare the event-free survival in subgroups of hormone sensitive and insensitive disease and in groups with 1–3, 4–9 or 10+ involved nodes between Arms A1 *versus* A2 and B1 *versus* B2.

*Tertiary Objective:*

- To determine prognostic factors like TS or TP and others on tumour tissue collected from primary surgery and to correlate them with study treatment effect.

**Related Publications:**

V.J. Möbus, M. Untch, A. du Bois, *et al.* Dose-dense sequential chemotherapy with epirubicin (E), paclitaxel (T) and cyclophosphamide (C) (ETC) is superior to conventional dosed chemotherapy in high-risk breast cancer patients ( $\geq 4$  + LN). First results of an AGO-trial. ASCO oral presentation, 5.6.2004.

**Topics:**

- Node positive disease
- Dose dense therapy
- Capecitabine
- Paclitaxel
- Bisphosphonate

**Keywords:**

Breast cancer, dose-dense, bisphosphonates

**Title:** A randomized, multi-center, open Phase III study comparing the postoperative use of zoledronic acid *versus* no treatment in patients with histological tumour residuals after preoperative anthracycline and taxane-containing chemotherapy for primary breast cancer (Neo-Adjuvant Trial Add-On) / GBG 36.

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**Summary:**

*Design:*

Prospective, randomized, open Phase III trial

*Study Population:*

Completely resected unilateral or bilateral primary carcinoma of the breast with histologically detectable tumour residuals (ypT2-4) and/or histologically confirmed involvement of axillary nodes (ypN1-3) after prior preoperative chemotherapy for at least 4 cycles, of which at least two must contain a taxane and an anthracycline.

*Primary Objective:*

To determine the event-free survival (EFS) after zoledronic acid for 5 years *versus* no postoperative treatment in patients with "chemo-insensitive" breast cancer (ypT2-4 and/or ypN1-3) after preoperative anthracycline/taxane-containing chemotherapy.

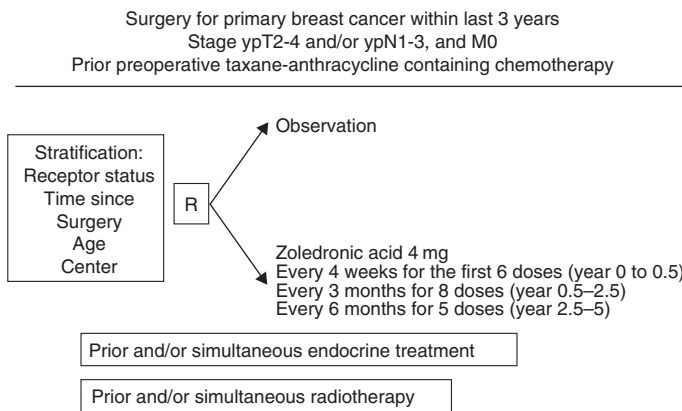
**Treatment:**

Zoledronic acid 4 mg (or adjusted dose based on renal function) will be given i.v.:

- Every 4 weeks for the first 6 doses (year 0–0.5)
- Every 3 months for 8 doses (year 0.5–2.5)
- Every 6 months for 5 doses (year 2.5–5)

Patients with hormone sensitive tumours (ER and/or PgR positive) should receive simultaneous endocrine treatment according to current treatment guidelines. In case an aromatase inhibitor is indicated for postmenopausal patients >50 years, letrozole should be used. Letrozole will be provided free of charge.

**Scheme:**



**Update:**

- Enrolment period: December 2004–December 2007.
- Planned patients: 904.
- Enrolled patients: 130.

**Related Publications:**

Diel IJ, Solomayer EF, Gollan C. Bisphosphonates in the reduction of metastases in breast cancer – extended follow-up results [Abstract 314]. *Proc Am Soc Clin Oncol* 2000.

Powles T, Paterson S, Kanis JA, McCloskey E, Ashley S, Tidy A, *et al.* Randomized, placebo-controlled trial of clodronate in patients with primary operable breast cancer. *J Clin Oncol* 2002; 20: 3219–3224.

**Topics:**

- Aromatase inhibitors
- Bisphosphonates
- Hormonal therapy

**Keywords:**

Breast cancer, adjuvant, bisphosphonates



**Title:** Prospective randomized multi-center study to prevent chemotherapy-induced ovarian failure with the GnRH-agonist goserelin in young hormone-insensitive breast cancer patients receiving anthracycline-containing (neo-)adjuvant chemotherapy / GBG 37, ZORO-Study

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**Summary:** *Design:*

Prospective, randomized, open Phase II trial

*Study Population:*

Age < 45 years, spontaneous and regular menstrual periods before study entry with FSH below 15 mIU/mL in follicular phase; histologically confirmed primary breast cancer with the need for anthracycline-based chemotherapy; steroid receptor (estrogen and progesterone) negative tumour (diagnosis according to hospital standard procedures).

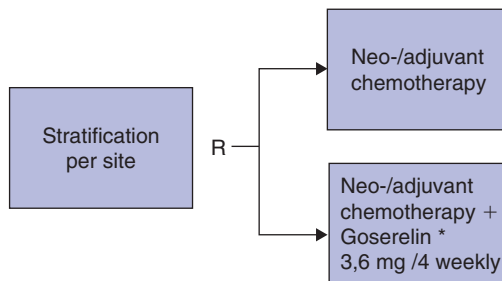
**Primary Objective:**

- To increase the percentage of patients with normal ovarian function at 6 months after application of (neo)adjuvant, anthracycline-containing polychemotherapy in parallel with goserelin compared to chemotherapy alone.

**Secondary Objectives:**

To compare the two treatment groups regarding:

- Compliance to treatment
- Toxicity
- Quality of life
- Menopausal symptoms score
- Ovarian function at 6, 12, 18 and 24 months
- Duration until recovery of regular menstrual period
- Pregnancy rate

**Scheme:**

\*first goserelin application >2 weeks prior to start of chemotherapy

**Update:**

- Planned Patient No.: 62.
- Patients recruited as of October 2006: 40.

**Related Publications:**

Recchia, *et al. Cancer* 2006; 106: 514–523.

**Topics:**

- Fertility and chemotherapy
- HR negative breast cancer
- Premenopausal patients

**Keywords:**

Ovarian function, goserelin, breast cancer

**Title:** A multi-centre Phase I–II study to investigate the combination of bendamustine with weekly paclitaxel as first or second line therapy in patients with anthracycline-pretreated metastatic breast cancer / GBG 38, Rita – Study.

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**Summary:** *Design:*

Prospective, multi-centre, sequential Phase I–II

*Primary Objectives:*

- To determine the maximum tolerated dose of the combination of bendamustine and weekly paclitaxel (Phase I part).
- To determine the objective response rate achievable with the recommended dose of this combination (Phase II part).

*Secondary Objectives:*

- To determine the time to progression (Phase II part).
- To determine the safety and tolerability of the combination.
- To determine the dose-limiting toxicity (DLT) (Grade IV haematological toxicities or Grade III non-haematological toxicities).

**Scheme:**

Dose level	Bendamustine (mg/m <sup>2</sup> )	Paclitaxel (mg/m <sup>2</sup> )
I	50	60
II	60	60
III	60	80
IV	70	80
V	70	90

Treatment will be given on day 1, 8, 15 and repeated on day 29. When the recommended dose has been determined a total of 47 patients will be treated at this dose level.

**Update:**

- Accrual October 2006 : 11 patients, Dosis-Level III filled, no DLTs.

**Related****Publications:**

None available

**Topics:**

- Metastatic breast cancer

**Keywords:**

Metastatic breast cancer, bendamustine, Phase I

**Title:** A multi-center Phase II study to determine the efficacy of capecitabine as first line monochemotherapy in patients with HER-2 negative metastatic breast cancer / GBG 39, Monica – Study.

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**Summary:** *Design:*

Prospective, open Phase II trial

*Primary Objective:*

- To determine the time to disease progression in patients with HER-2 negative metastatic breast cancer after 1st line monochemotherapy with capecitabine.

*Secondary Objectives:*

- To determine the objective response rate.
- To determine the duration of response.

- To determine the clinical benefit defined as CR, PR, or stable disease  $\geq 24$  weeks.
- To evaluate the safety and toxicity of capecitabine.
- To assess quality of life within 1 year after start of capecitabine treatment.
- To determine overall survival.
- To determine the objective response rate in male patients.
- To evaluate QoL the modified Brunner Score.

*Tertiary Objective:*

- To determine the DPD and proteomics in serum.

**Scheme:** Capecitabine 2000 mg/m<sup>2</sup> orally day 1–14 q day 22 until progression

**Update:**

- Planned patients: 200.
- Accrual October 2006: 88.

**Related Publications:** None available

**Topics:**

- Capecitabine
- HER-2 negative patients
- Metastatic breast cancer

**Keywords:** Capecitabine, HER-2 negative patients, metastatic breast cancer

**Title:** GeparQuattro: A randomized Phase III study exploring the efficacy of capecitabine given concomitantly or in sequence to EC – Doc with or without trastuzumab as neoadjuvant treatment of primary breast cancer.  
**A joint study of GBG and AGO / GBG 40.**

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**Summary:** *Design:*

Prospective, randomized, Phase III trial

*Study Population:*

Locally advanced (T4 a–d, N0-3, M0) or operable (T  $\geq$  2 cm, N0-2, M0) primary breast cancer. Tumour lesion in the breast with a palpable size of  $\geq$ 2 cm or a sonographically size of  $\geq$ 1 cm in maximum diameter. Stage of disease in which also adjuvant chemotherapy would be considered.

*Primary Objectives:*

- To compare the pCR rates of NACT with *versus* without capecitabine and 8 *versus* 12 cycles in patients with primary breast cancer.
- To compare the pCR rates in patients with HER-2/neu positive tumours receiving trastuzumab simultaneously to NACT to patients with HER-2/neu negative tumours receiving NACT only.

*Treatment:*

All patients will receive 4 cycles of EC:

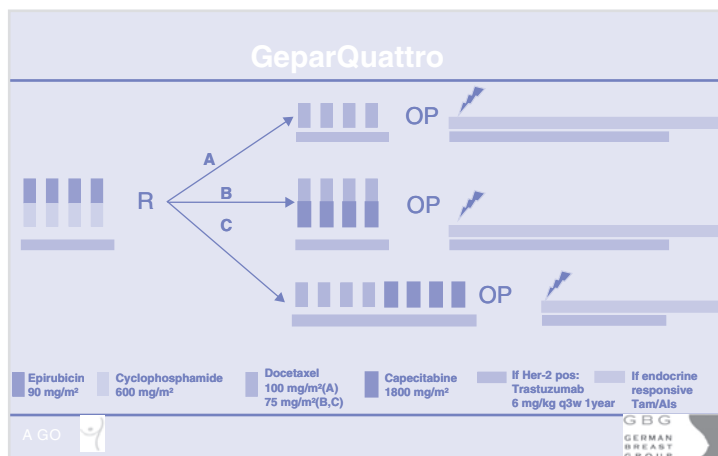
- Epirubicin 90 mg/m<sup>2</sup> given *simultaneously with*
- Cyclophosphamide 600 mg/m<sup>2</sup>, all day 1 q day 21

Thereafter they will be randomized to:

- *Arm A:* Docetaxel 100 mg/m<sup>2</sup> day 1 q day 21 for 4 cycles (EC-Doc)
- *Arm B:* Docetaxel 75 mg/m<sup>2</sup> day 1 q day 21 for 4 cycles *concomitantly given with* Capecitabine 1800 mg/m<sup>2</sup> day 1–14 q day 21 for 4 cycles (EC-DocX)
- *Arm C:* Docetaxel 75 mg/m<sup>2</sup> day 1 q day 21 for 4 cycles *followed by* Capecitabine 1800 mg/m<sup>2</sup> day 1–14 q day 21 for 4 cycles (EC-Doc-X)

Patients with HER-2/neu positive tumors will receive trastuzumab 6 mg/kg intravenous (i.v.) every 3 weeks concomitantly to cytotoxic treatment, starting with a loading dose of 8 mg/kg i.v. on day 1 of the first EC-cycle. A total number of 8 (in the EC-Doc and EC-DocX arm) or 12 (in the EC-Doc-X arm) infusions will be given preoperatively.



**Scheme:****Update:**

- Enrolment period: August 2005–December 2006.
- Planned patients: 1042.
- Enrolled patients: 510.

**Related Publications:**

None available

**Topics:**

- Anthracyclines
- Capecitabine
- Locally advanced breast cancer
- Taxanes
- Trastuzumab

**Keywords:**

Preoperative chemotherapy

**Title:** Randomized study comparing 6 × FEC with 3 × FEC followed by 3 × docetaxel in high-risk node-negative patients with operable breast cancer: comparison of efficacy and evaluation of clinico-pathological and biochemical markers as risk selection criteria.  
A joint study of AGO, the EORTC Receptor and Biomarker Group, and the GBG (GBG 42 / NNBC 3-Europe).

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**Summary:** *Design:*

The trial is a prospective, randomized multi-centre open-label Phase III trial that is designed to detect a difference in efficacy between two chemotherapy regimens (FE100C\*6 *versus* FE100C\*3 followed by Docetaxel100\*3) in high-risk node-negative breast cancer patients. Additionally, risk assessment by the traditional clinico-pathological factors and by tumour-biological factors (aPA/PAI-1) will be compared.

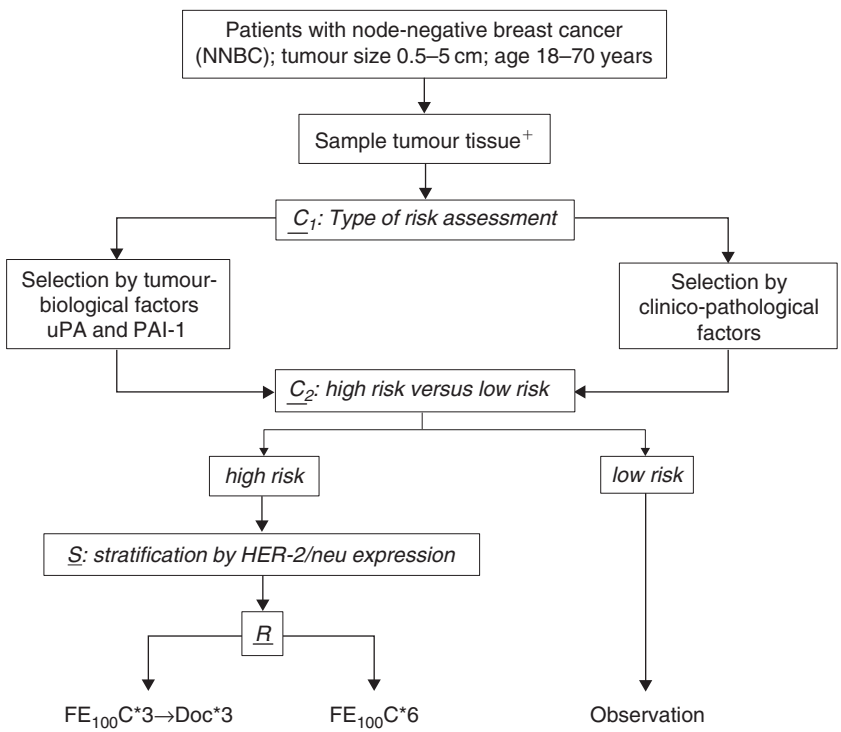
**Study Population:**

Primary breast cancer, tumour size  $\geq 0.5$  cm and  $\leq 5$  cm (pT1b-pT2, pN0, M0), age  $\geq 18$  years,  $\leq 70$  years

**Aims/Endpoints:**

- Amongst the chemotherapy treated population:
  - The primary endpoint of the study is Disease-Free Survival (DFS).
  - The secondary endpoints are: Overall Survival (OS) and safety.
- Amongst the entire population of registered patients:
  - DFS in each low-risk group (or of each patient group stratified for type of risk selection, respectively).
  - The proportion of node-negative breast cancer patients grouped into each low-risk group.
  - Side-effects of chemotherapy in each patient group.

**Scheme:**



Patients with hormone sensitive disease should receive endocrine treatment according to the latest recommendations of the AGO. \* = fresh tumour tissue in centres with biological risk assessment, additionally in all patients paraffin blocks for central review.

**Update:** ● 999 patients enrolled as of October 2006.

**Related Publications:**

Jänicke F, Prechtel A, Thomssen C, *et al.* for the German “Chemo-N0” Study Group. Randomized adjuvant chemotherapy trial in high-risk, lymph node-negative breast cancer patients identified by urokinase-type plasminogen activator and plasminogen activator inhibitor type 1. *J Natl Cancer Inst* 2001; 93(12): 913–920.

Look MP, van Putten WLJ, Duffy MJ, *et al.* Pooled analysis of prognostic impact of uPA and PAI-1 in 8377 breast cancer patients. *J Natl Cancer Inst* 2002; 94(2): 116–128.

Harbeck N, Kates RE, Gauger K, *et al.* Urokinase-type plasminogen activator (uPA) and its inhibitor PAI-1: novel tumor-derived factors with a high prognostic and predictive impact in breast cancer. *Thromb Haemost* 2004; 91(3): 450–456. Review.

Hayes DF. Prognostic and predictive factors revisited. *Breast* 2005; 14(6): 493–499. *Epub* 2005 Oct 18. Review.

**Topics:**

- Taxanes
- Prognostic factors
- Node-negative breast cancer
- UPA
- PAI-1

**Keywords:** Taxanes, prognostic factors, node-negative breast cancer, uPA, PAI-1