

FAST FACTS

AFT – 25 COMPARISON OF OPERATIVE TO MONITORING AND ENDOCRINE THERAPY (COMET) TRIAL FOR LOW RISK DCIS: A PHASE III PROSPECTIVE RANDOMIZED TRIAL

PATIENT SELECTION/POPULATION

Inclusion Criteria

1. New diagnosis of DCIS without invasive cancer; date of diagnosis defined as the date of the first pathology report that diagnosed the patient with DCIS
2. Unilateral, bilateral, unifocal, or multifocal DCIS
3. ADH/borderline DCIS
4. A patient who has had a lumpectomy with positive margins as part of their treatment for a current DCIS diagnosis is eligible (post-excision mammogram required at enrollment to establish a new baseline)
5. No previous history of breast cancer (DCIS or invasive cancer) in either breast prior to current DCIS diagnosis
6. 40 years of age or older at time of DCIS diagnosis
7. ECOG performance status 0 or 1
8. No contraindication for surgery
9. Baseline imaging:
 - unilateral DCIS: contralateral normal mammogram \leq 6 months of registration and ipsilateral breast imaging \leq 120 days of registration (must include ipsilateral mammogram; can also include US or breast MRI)
 - bilateral DCIS: bilateral breast imaging \leq 120 days of registration (must include bilateral mammogram; can also include ultrasound or breast MRI)
10. Pathologic criteria:
 - All grade I DCIS (irrespective of necrosis/comedonecrosis)
 - All grade II DCIS (irrespective of necrosis/comedonecrosis)
 - Absence of invasion or microinvasion
 - Diagnosis confirmed on core needle, vacuum-assisted or surgery \leq 120 days of registration
 - ER(+) and/or PR(+) by IHC (\geq 10% staining or Allred score \geq 4)
 - HER2 0, 1+, or 2+ by IHC if HER2 testing is performed
11. Histology slides reviewed and agreement between **two** clinical pathologists (not required to be at same institution) that pathology fulfils COMET eligibility criteria. In cases of disagreement between the two pathology reviews about whether or not a case fulfils the eligibility criteria, a third pathology review will be required.
12. At least two sites of biopsy for those cases where mammographic extent of calcifications exceeds 4 cm, with second biopsy benign or both sites fulfilling pathology eligibility criteria
13. Amenable to follow up examinations
14. Ability to read, understand and evaluate study materials
15. Reads and speaks Spanish or English

Exclusion Criteria

1. All grade III DCIS
2. Male DCIS
3. Concurrent diagnosis of invasive or microinvasive breast cancer in either breast
4. Documented mass on examination or imaging at site of DCIS **prior to** biopsy yielding diagnosis of DCIS, with exception of fibroadenoma at a distinct/separate site from site of DCIS (or diagnosis of mass as a cyst). In cases of uncertainty about whether the mass was present on physical examination prior to biopsy, the following criteria should be applied: if mammogram noting abnormal findings is diagnostic MMG = symptomatic/if mammogram noting abnormal findings is screening MMG = asymptomatic
5. Bloody nipple discharge (ipsilateral breast)
6. Mammographic finding of BIRADS 4 or greater within 6 months of registration at site other than that of known DCIS, without pathologic assessment

7. Use of investigational cancer agents within 6 weeks prior to diagnosis
8. Any serious and/or unstable pre-existing medical, psychiatric, or other existing condition that would prevent compliance with the trial or consent process
9. Pregnancy. If a woman has been confirmed as pregnant she will not be eligible to take part in the trial. If she suspects there is a chance that she may be pregnant, a pregnancy test should be undertaken; however, a pregnancy test for all women of child-bearing potential is not mandatory
10. Documented history of prior tamoxifen, aromatase inhibitor, or raloxifene in last 6 months

Stratification Factors

- age at diagnosis: <55, 55-65, >65
- maximum diameter of microcalcifications: <2cms, 2-5cms, >5cms
- DCIS nuclear grade: I or II

Schedule of Eligibility Screening – Refer to Table 2

- History/physical
- Weight, height
- Pulse, Blood Pressure
- Breast imaging
- Bilateral MMG
- PRO surveys

Study Flow Diagram

