

## **Listing of PORTEC-4a Protocol Amendments (version 2.6, 3 August 2021)**

### **Listing of changes in version 2.6:**

#### **1. Administrative changes in the protocol**

- Addition of two more participating German sites and 1 Swiss site with their PIs and pathologist. An update on study coordinators, participating groups, centers, local investigators and trial coordinators (page 2, pages 29-30).
- Removal of one German site (page 2, pages 29)
- Exclusion criteria: Addition of the word “expected’ to the exclusion criterium Interval between the operation and start of radiotherapy exceeding 8 weeks (page 6; and section 5.2 page 13)
- Specification that after September 2021 no new centers will be activated, since the trial is nearing completion of accrual (page 6).
- Specification of the total number of patients who will be recruited, which has been increased to 550 evaluable patients, including 50 patients of the pilot phase (page 6; section 4, page 13, and section 12, page 24). Furthermore it is described which patients of the previous PORTEC-4 trial will be included into the PORTEC-4a trial (section 12, page 24).
- The total study duration has been further specified to 7 year follow up after the inclusion of the last patient (page 6).
- Specification of the timeframe in which the results of the molecular profile for patients in the experimental arm should be completed and send to the trial office in accordance with current timelines (page 19).
- Addition of the possibility to perform virtual on-site monitoring because of the COVID pandemic, and specification that as of 2021 international sites with high inclusion rates are being virtually monitored (page 23)

#### **2. Changes in ABR form:**

- Section C: updated information in this section:
  - C4: Specification of the total number of patients: a total of 550 eligible and evaluable patients, including 22 patients from the previous PORTEC-4 trial
  - C11: section C11a: 450, section C11b: 100, section C11c: 550
- Section D: updated information in this section:
  - D2: Total number of patients 550.
  - D5a/b: Specification of the interval between the operation and start of radiotherapy. Expected interval between the operation and start of radiotherapy exceeding 8 weeks and ‘Een te verwachten interval tussen de operatie en start van brachytherapie > 8 wk’.
- Section K: updated information in this section:
  - Onderzoeksopszet/Study design: The number of patients to be recruited in the PORTEC-4a trial will be 550, of whom 22 patients from the previous PORTEC-4 trial