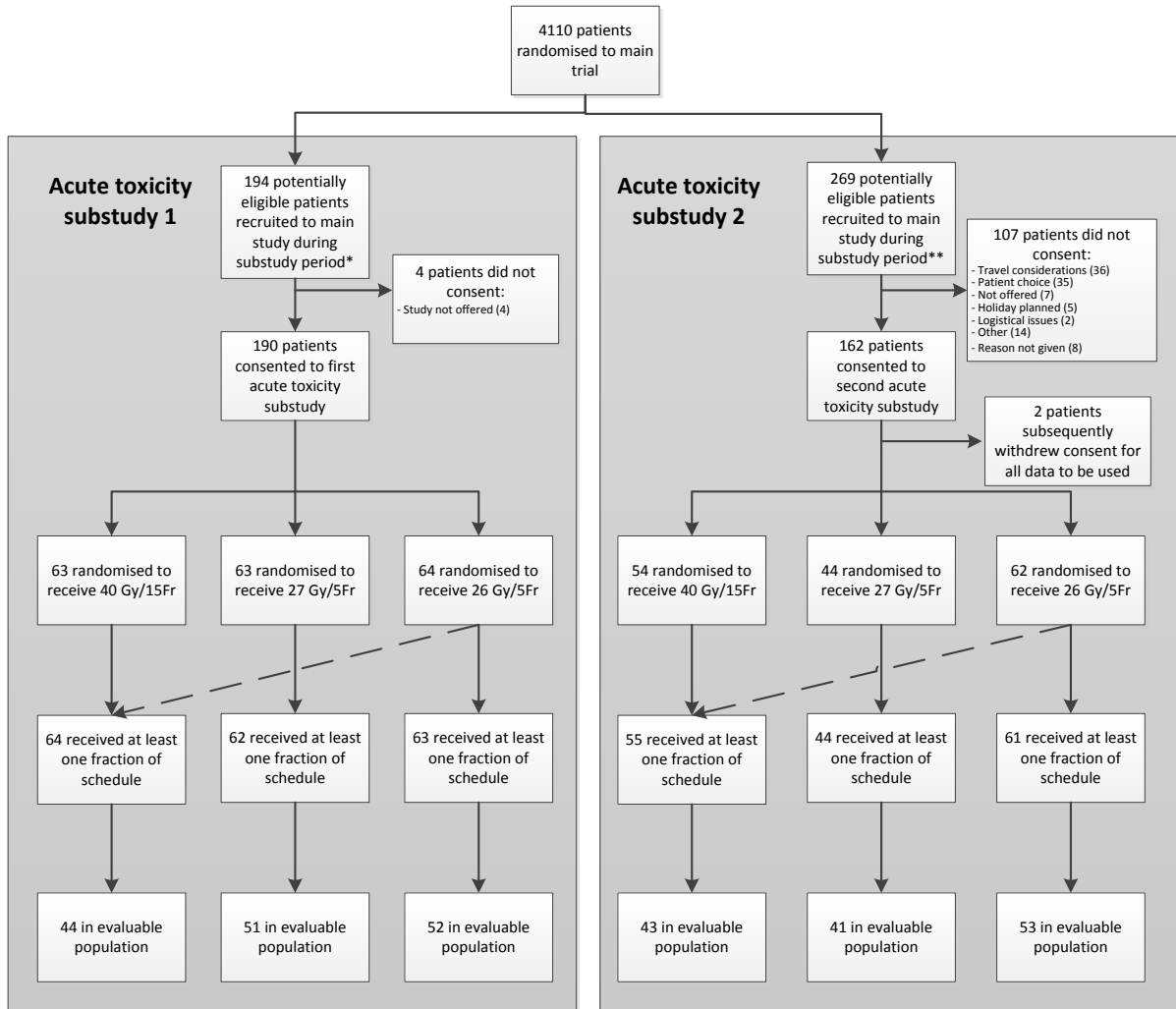


SUPPLEMENTARY FILES

Appendix 1: Flow diagram of participation in the first and second acute toxicity substudies



Appendix 2: Baseline characteristics of patients in first and second acute toxicity substudies

| Baseline characteristics | Acute toxicity substudy 1 | | | Acute toxicity substudy 2 | | |
|---------------------------|---------------------------|---------------|---------------|---------------------------|---------------|---------------|
| | 40 Gy/15F | 27 Gy/5F | 26 Gy/5F | 40Gy/15fr | 27Gy/5fr | 26Gy/5fr |
| | N=63 N (%) | N=63 N (%) | N=64 N (%) | N=54 N (%) | N=44 N (%) | N=62 N (%) |
| Age | | | | | | |
| Median (IQR) | 63 (57 - 69) | 65 (54 - 68) | 63 (55 - 68) | 60 (54-66) | 61 (56-70) | 61 (54-64) |
| Range | 46-87 | 41-82 | 44-86 | 47-81 | 49-84 | 42-78 |
| Age category | | | | | | |
| <50 | 5 (8) | 11 (17) | 10 (16) | 3 (5) | 2 (5) | 6 (10) |
| ≥50 | 58 (92) | 52 (83) | 54 (84) | 52 (95) | 42 (95) | 57 (90) |
| Tumour grade | | | | | | |
| 1 | 15 (24) | 16 (25) | 17 (27) | 14 (25) | 10 (23) | 16 (25) |
| 2 | 28 (44) | 31 (49) | 33 (52) | 27 (49) | 23 (52) | 34 (54) |
| 3 | 20 (32) | 16 (25) | 14 (22) | 14 (25) | 11 (25) | 13 (21) |
| Risk group | | | | | | |
| Low (≥50 & grade 1 or 2) | 40 (63) | 39 (62) | 43 (67) | 38 (69) | 31 (70) | 45 (71) |
| High (<50 and/or grade 3) | 23 (37) | 24 (38) | 21 (33) | 17 (31) | 13 (30) | 18 (29) |
| Boost planned | | | | | | |
| No | 54 (86) | 52 (83) | 59 (92) | 55 (100) | 44 (100) | 63 (100) |
| Yes | 9 (14) | 11 (17) | 5 (8) | 0 | 0 | 0 |

Appendix 3: Modified RTOG and CTCAEv4.03 definitions used in first and second acute skin toxicity sub-studies

Acute toxicity substudy 1

| *Modified RTOG | Description |
|-----------------------|--|
| Grade 1 | Faint/dull erythema |
| Grade 2 | Tender/bright erythema +/- dry desquamation |
| Grade 3 | Patchy moist desquamation, moderate oedema |
| Grade 4 | Confluent moist desquamation, pitting oedema |

Acute toxicity substudy 2

| CTCAEv4.03 | Description |
|-------------------|--|
| Grade 1 | Faint erythema or dry desquamation |
| Grade 2 | Moderate to brisk erythema; patchy moist desquamation, mostly confined to skin fold; moderate oedema |
| Grade 3 | Moist desquamation in areas other than skin folds and creases; bleeding induced by minor trauma or abrasion |
| Grade 4 | Life threatening consequences; skin necrosis or ulceration of full thickness dermis; spontaneous bleeding from involved site; skin graft indicated |

*The inclusion of oedema in the modified RTOG system proved unhelpful in this particular context, since the primary concern was to score moist desquamation; hence the second substudy was undertaken using CTCAE criteria

Appendix 5: Missed assessment form



Missed Assessment Form

| | | | | | | | | | |
|--------------------|----------------------|----------------------|----------------------|----------------------|----------------------|----------------------|----------------------|----------------------|----------------------|
| RT Centre | <input type="text"/> | Trial number | FF | <input type="text"/> | <input type="text"/> | <input type="text"/> | <input type="text"/> | <input type="text"/> | <input type="text"/> |
| Referring Hospital | <input type="text"/> | | | | | | | | |
| Patient's Initials | <input type="text"/> | <input type="text"/> | <input type="text"/> | Date of Birth | <input type="text"/> | <input type="text"/> | <input type="text"/> | <input type="text"/> | <input type="text"/> |
| | | | | | Day | Month | Year | | |

TO BE COMPLETED IF A PATIENT DOES NOT ATTEND FOR A TOXICITY ASSESSMENT

Was the patient called if they failed to turn up for a toxicity assessment? Yes* No**

** If yes, please complete the rest of this form*

*** If no, please specify reason*

Date of telephone assessment
Day Month Year

The patient should be asked the following questions:

Please state reason for not attending for an assessment

Has there been any change in skin colour since last week?

Yes, less pink/red Yes, more pink/red No, same colour

Is there been any evidence of skin weeping or oozing?

Yes* No

** If yes, please specify the site*

Skin creases/folds Other

| | | | | | | | | | |
|------------|----------------------|---------------------|----------------------|----------------------|----------------------|----------------------|----------------------|----------------------|----------------------|
| Signature | <input type="text"/> | Date form completed | <input type="text"/> | <input type="text"/> | <input type="text"/> | <input type="text"/> | <input type="text"/> | <input type="text"/> | <input type="text"/> |
| Print Name | <input type="text"/> | | | | | | | | |
| | | | | | Day | Month | Year | | |

| | | |
|--------------------------------|----------|----------|
| Received | Entered | Checked |
| TRAQS <input type="checkbox"/> | | |
| Initials | Initials | Initials |