For **Group-Specific StudyHandbook Amendment** # 1 to: GOG-0274/RTOG1174

NCI Protocol #: GOG-0274/RTOG1174
NCI Version Date: March 21, 2013

NOTE: Amendment #1 was submitted to CTEP including Updates which were previously broadcast to the Groups on December 10, 2012. (***RTOG Broadcast December 20, 2012).*** Changes that were made as updates are **noted** in the Summary of Change memo with ***

All changes include effective dates **(05/13/2013)**. **(*RTOG Broadcast May 14, 2013*)**

#	Section	Change
1.	TP	Version date and Version 2 have been added to footer. Version 2 has been updated in the header throughout the handbook. ***
2.	TOC	Has been updated and hyperlinked per NCI mandate.
3.	1.0	Titles have been corrected for study personnel. *** CTC has been defined as Clinical Trials Centre. ***
4.	1.2	"…and completion of the InForm training exercise," was added. "Inform" was replaced with "live OUTBACK" ***
5.	2.2	"The screening logs are located in the Investigator Site File. Completed screening logs are to be sent to the GCIG coordinating office on a regular basis." has been deleted. ***
6.	3.0	Grammar has been corrected and "NOTE: If at any time you are experiencing difficulty accessing the OUTBACK database please email the study mailbox at outback@ ctc.usyd.edu.au " has been added to this section. ***
7.	3.1	The screen shots for the InForm System have been updated Section "GOG Registration" has been added. "**NOTE"** for RTOG non US members has been added. ***
8.	4.1	In Example 1, "28 fractions" has been corrected to "25". ***
9.	4.1	In Example 2, "28 fractions" has been corrected to "25". ***
10.	4.1	In Example 3, "28 fractions" has been corrected to "25". ***
11.	4.1	In Example 4, "28 fractions" has been corrected to "25". ***
12.	5.2.6	Section number has been corrected. 3) doses will be based on subjects weight "at the time of start of adjuvant chemotherapy" replaced "baseline" ***
13	6.0	Section has been completely updated with current TR language and instructions.
14	6.11	Section numbering has been updated

#	Section	Change
15.	7.0	Grammar has been corrected. And "NOTE:If at any time you are experiencing difficulty accessing the OUTBACK database please email the study mailbox at outback@ ctc.usyd.edu.au" has been added to this section***
16.	7.1	"The NHMRC CTC will send two types of correspondence to GCIG coordinating centres for distribution to local sites regarding trial data: patient query reports and overdue eCRF/queries reports." Has been replaced with: "The NHMRC CTC will raise queries as necessary after reviewing eCRFs, and send out overdue form reports to sites on a regular basis." Overdue "sCRF queries" is now Overdue "Form" reports. Specific instructions have been updated. ***
17.	7.1	"eCRF/queries" is now "form".
18.	7.2	Specific completion guidelines have been edited.
19.	7.3	Instructions for completing QOL forms have been updated for GOG and RTOG institutions. ***
20.	8.0	SAE reporting to ANZGOG has been updated throughout to conform with InForm.
21.	9.0	"and these will be communicated to GCIG coordinating centres for distribution to local sites." Has been deleted. ***
22.	9.1	"RT and QA" have been added. Paragraph explaining QA process has been replaced with updated information. **Brachytherapy** paragraph regarding Data submission has been deleted. ***

For **Group-Specific Informed Consent** Amendment # 1 to: GOG-0274/RTOG1174

NCI Protocol #: GOG-0274/RTOG1174
NCI Version Date: March 21, 2013

#	Section	Change
1.	Title Page	NCI Version date has been updated.
2.	Study Chart	In the table for Arm A and Arm B "**What you do**" on Days 1, 8, 15, 22 and 29 "60" minutes has been corrected to "30",and "90" to "60". Day 34 "with" has been corrected to "will". For Arm B-Cycles 2-4 with Carboplatin and Paclitaxel. Days 1: infusion time has been inserted. Column has been added to state "4 weeks after additional 12 weeks of chemo is complete" Physical exam that may include pelvic and routine blood tests have been added.
3.	ICD	In the section **General Information About the Collection and Use of Specimens for Research,** "Some of your" has been added to the first sentence. -"(if available from a previous surgery or biopsy) and blood" has been replaced with, "specimens" in the last paragraph of the section.
4.	ICD	Throughout the section **Specific Information for this Research Study,** the specimen requirements have been updated for clarity.
5.	ICD	Throughout the section **What Will Happen To Your Specimens If You Agree,** specimen information has been updated for clarity.

#	Section	Change
6.	ICD	In the Section **Specific Information for Future Research**, “Reports of this research done in your specimens will not be given to you or your doctor, or be put in your health record.” has been deleted. The penultimate paragraph has been deleted

Summary of changes for OUTBACK protocol amendment v4.0 dated 2 November 2012

NB. Page numbers refer to the track changes version of protocol 4.0

1. Page 6: The synopsis has been updated
2. Page 13: The randomisation process has been clarified
3. Page 15: The section about patient withdrawal has been amended to explain about collection of follow-up data if allowed by the patient
4. Pages 16-22: The chemotherapy section has been updated to clarify dosing requirements and dose modifications
5. Page 26: The section about brachytherapy has been amended with regard to the use of interstitial brachytherapy if the primary tumour does not respond to EBRT
6. Page 29-30: Radiation quality control requirements have been clarified
7. Page 33: Information has been added about the use of a standardised checklist to record reasons for non-completion of quality of life questionnaires
8. Pages 34-36: The section about SAEs has been updated to reflect the current template of the NHMRC Clinical Trials Centre
9. Pages 39-41: The schedule of assessments has been amended to clarify assessment requirements
10. Pages 51-65: The PICs have been updated to clarify that the study drug paclitaxel is only provided to Australian sites, and that the amount of blood taken for the optional translational research is 35ml
11. Pages 66-75: The Quality of Life questionnaires have been compiled into a Quality of Life booklet, but the questionnaires have not been changed
12. Minor inconsistencies in formatting and spelling have been corrected throughout the protocol

SUMMARY OF CHANGES
Update: December 10, 2012
(Broadcast: December 18, 2012)

RTOG 1174 (ANZGOG 0902/GOG-0274), "A Phase III Trial of Adjuvant Chemotherapy as Primary Treatment for Locally Advanced Cervical Cancer Compared to Chemoradiation Alone: The OUTBACK Trial"

Study Chair: William Small, MD; 312-926-2520;WSmall@nmff.org

The US Group-Specific Study Handbook has been updated (vs. December 10, 2012) as follows:

Title Page: The update date has been added; version 2 has been added to header.

Section 1.0: Titles have been corrected for study personnel; CTC has been defined as Clinical Trials Centre.

Section 1.2: "…and completion of the InForm training exercise," was added. "Inform" was replaced with "live OUTBACK"

Section 2.2: "The screening logs are located in the Investigator Site File. Completed screening logs are to be sent to the GCIG coordinating office on a regular basis." was deleted.

Section 3.0: Grammar has been corrected and "NOTE: If at any time you are experiencing difficulty accessing the OUTBACK database please email the study mailbox at outback@ctc.usyd.edu.au." was added to the last paragraph

Section 3.1: Section "GOG Registration" has been added. "NOTE" for RTOG non US members has been added.

Section 4.0: Under "Total dose reporting" **"**28 fractions" has been corrected to "25" in Examples 1, 2, 3 and 4.

Section 5.2.6.4: doses will be based on subject's weight "at the time of start of adjuvant chemotherapy" replaced "baseline" in the 3rd bullet under "Dosing of Carboplatin"

Section 6.2 Section Footnotes 4 and 5 have been replaced

Section 6.13: **"**RTOG" has been added to the section head. "Note: the FFPE block or unstained slides must correspond with the H&E slide." has been added to the second bullet. Fifth bullet "(ST)" has been added after Specimen Transmittal. "Section 6.13.7" has been added for reference to shipping instructions.

Section 6.13.3: Incorrect symbol has been corrected to symbol for degree. "…and time stored" has been changed to "specific"

Section 6.13.4: the table, the address for specimen submission, and bullets with instructions have been updated for clarity

Section 6.13.5: "shaking" has been changed to "rattling" through-out the section and the bullets have been revised for clarity

Section 6.13.6: First bullet has been updated with instructions for handling Frozen Specimens; the FAX number has been removed

Section 6.13.7: Labeling instructions for Plasma have been updated; the 4th bullet under "Process" for plasma has been updated; Under "Process" for whole blood: duplicate word has been deleted in the 1st bullet;.the 2nd bullet was updated;. Symbols have been corrected in the 3rd and 4th bullet

Section 7.0: Grammar has been corrected and "NOTE: If at any time you are experiencing difficulty accessing the OUTBACK database please email the study mailbox at outback@ctc.usyd.edu.au." was added

Section 7.1: Under "Data Requests"**,** the NHMRC CTC will send two types of correspondence to GCIG coordinating centres for distribution to local sites regarding trial data: patient query reports and overdue eCRF/queries reports" has been replaced with "The NHMRC CTC will raise queries as necessary after reviewing eCRFs, and send out overdue form reports to sites on a regular basis."; Overdue "sCRF queries" is now Overdue "Form" reports; Specific instructions have been updated.

Section 7.3: Instructions for completing QOL forms have been updated for GOG and RTOG institutions; Instructions have been inserted for time when patient does not complete QOL questionnaire

Section 8.0: Outback e-mail address and instructions have been added

Section 9.0: The sentence**, "**and these will be communicated to GCIG coordinating centres for distribution to local sites" has been deleted

Section 9.1: RT and QA" have been added to the 1st paragraph; paragraph explaining QA process has been replaced with updated information; brachytherapy paragraph regarding data submission has been deleted.

SUMMARY OF CHANGES
Administrative Update: April 26, 2012
(Broadcast: April 26, 2012)

Group-Specific Study Handbook and Informed Consent for:

RTOG 1174 (ANZGOG 0902/GOG-0274), "A Phase III Trial of Adjuvant Chemotherapy as Primary Treatment for Locally Advanced Cervical Cancer Compared to Chemoradiation Alone: The OUTBACK Trial"

Study Chair: William Small, MD; 312-926-2520;WSmall@nmff.org

The Handbook has been updated as follows:

Section 6.13.1: RTOG has been added to the title of section.

Section 6.13.2: RTOG has been added to the title of section**.**

Section 6.13.3: "samples" has been deleted and replaced with "and Plasma aliquots".

Section 6.13.4: The mailing and courier addresses of the RTOG Biospecimen Resource have been updated for frozen and non-frozen specimens.

Section 6.13.5: "and" has been added in the third paragraph and "or Plugs" has been deleted

Section 6.13.6: "an FFPE Specimen Plug" has been deleted in the final paragraph and has been replaced with, "Frozen Biospecimen Collection"

The Group Specific Informed Consent has been updated as follows:

"Specific Information for Future Research" ,"*Making Your Choices About Future Research*"**:** RTOG has been added to the third and fifth questions for clarity