

North Central Cancer Treatment Group

N0321: Phase I/II Study of PS-341 in Combination with Paclitaxel, Carboplatin, and Concurrent Thoracic Radiation Therapy for Non-small Cell Lung Cancer (NSCLC)

Addendum 13 – April 16, 2010

Summary

- An updated Comprehensive Adverse Event and Potential Risk listing (CAEPR) for PS-341 (version 2.2, dated March 25, 2010) has been received from the National Cancer Institute (NCI). Therefore, the Drug Information section and consent forms have been updated accordingly.
- Administrative/editorial changes.

Replacement pages are included. Please incorporate into the protocol and keep this addendum with your protocol.

Title Page Updated to reflect Addendum 13 and revised NCI version date.

Protocol Resource Page

Page 2: **Rachael M. Meyers** replaces ~~Sarah Hanson~~ as the NCCTG *Research Base* Quality Assurance Specialist.

Kristine M. Hacker, R.N. replaces ~~Dianne M. Herrera, R.N.~~ as the NCCTG *Research Base* Nurse.

Section 15.0 Drug Information

Pages 39-42: Due to the receipt of the updated CAEPR for PS-341 from NCI, Section 15.16 has been replaced in its entirety.

Appendices IA-IB Consent Forms

Pages 5-6: Due to the receipt of the updated CAEPR for PS-341 from NCI, the risks sections of the consent forms have been updated as follows:

Likely related side effects of PS-341 (events occurring greater than 20% of the time):

- Fatigue **or tiredness** (*expanded wording*)
- Loss of appetite
- Nausea (feeling sick to your stomach)
- Vomiting (throwing up)
- Swelling and fluid buildup in the arms or legs
- Diarrhea (loose stools)
- Constipation (difficulty passing stool)
- Fever
- Decreased number of blood cells that help to clot the blood
- ~~Nerve damage causing numbness, tingling, burning (peripheral neuropathy)~~ **Inflammation (swelling and redness) or degeneration of the peripheral nerves (those nerves outside of brain and spinal cord) causing numbness, tingling, burning** (*reworded*)

- ~~Decrease in a red blood cell protein that carries oxygen in the body~~ **Lack of enough red blood cells (anemia)** (*reworded*)
- Infection
- **Weakness or paralysis (loss of muscle function) caused by damage to peripheral nerves (those nerves outside of brain and spinal cord)** (*moved from Less Likely and reworded*)

Less likely related side effects of PS-341 (events occurring greater than 3% but less than 20% of the time):

- Chills, shivering
- Heartburn
- ~~Abnormally slow bowel contraction~~ **Partial or complete blockage of the small and/or large bowel. Ileus is a functional rather than actual blockage of the bowel** (*reworded*)
- Irritation or sores in the lining of the throat
- Fever with dangerously low white blood cell count
- Double or blurred vision
- Back pain
- Excess fluid that accumulates in the pleural cavity, the fluid-filled space that surrounds the lungs
- ~~Weakness or paralysis caused by damage to nerves~~ (*moved to Likely and reworded*)
- Infection which occurs due to a decreased number of a type of white blood cell (**lymphocyte**) (*expanded wording*)
- Dehydration (loss of body fluids)
- Nerve damage causing numbness, tingling, burning (sensory neuropathy)
- Dizziness (**or sensation of lightheadedness, unsteadiness, giddiness, spinning or rocking**) (*expanded wording*)
- Anxiety, **feelings of dread or danger** (*expanded wording*)
- Cough
- Shortness of breath
- ~~Rash/flaking or shedding of outer layer of skin~~ **Skin rash with the presence of macules (flat discolored area) and papules (raised bump)** (*reworded*)
- Low blood pressure
- Pneumonia
- Difficulty sleeping or falling asleep
- Decreased total number of white blood cells (**lymphocyte**) (*expanded wording*)
- Decreased number of a type of white blood cell (**neutrophil/granulocyte**) (*expanded wording*)
- **Decrease in the total number of white blood cells (leukocytes)** (*newly added*)
- Bleeding of the digestive tract
- Nosebleed
- Blood infection
- Muscle weakness of the whole body
- Fainting
- Belly pain
- Bone pain
- Leg **and/or arm** pain (*expanded wording*)
- **Headache or Hhead pain** (*expanded wording*)

- Joint pain
- Muscle pain
- Nerve pain

Rare but serious side effects of PS-341 (events occurring less than 3% of the time):

- A hole in the digestive tract
- Syndrome associated with high blood pressure characterized by headache, confusion, seizures, and vision loss associated with imaging findings
- ~~Kidney failure~~ **Sudden or traumatic injury to the kidney** (*reworded*)
- **Progressive necrosis (tissue death) of a part (the white matter) of the brain without inflammation (swelling and redness)** (*newly added*)

Appendix IB **Consent Form**

Page 2: In the study table for Day 23, the second bullet has had a typographical error corrected as follows:

Paclitaxel given into the vein over 3 ~~y~~h hours followed by Carboplatin given over 30 minutes