

## North Central Cancer Treatment Group

N0821: A Phase II First-Line Study of a Combination of Pemetrexed, Carboplatin and Bevacizumab in Advanced Nonsquamous NSCLC Evaluating Efficacy and Tolerability in Elderly Patients (Age  $\geq 70$  yrs) with Good Performance Status (PS  $< 2$ )

Addendum 8 – October 21, 2011

### Summary

- An updated Investigator Brochure (IB) for Pemetrexed (LY31514) dated June 24, 2011 has been received. Therefore, The Drug Information section and the consent form have been updated accordingly.
- Administrative/editorial changes.

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

**Title page** Updated to reflect Addendum 8 and current NCI version date.

### **Protocol Resources**

Page 2: The NCCTG Research Base Nurse's name has changed from Kristine M. ~~Hacker~~, R.N. to Kristine M. **Sorgatz**, R.N.

### **Section 15.0** **Drug Information**

Page 41: Due to the receipt of the updated IB for Pemetrexed, Section 15.13 has been updated as follows:

- Likely ( $>20\%$ ): Neutropenia, anemia, nausea, vomiting, diarrhea, fatigue, anorexia, mild hair loss, abdominal pain, and itchy skin rash, dyspnea, weakness, ~~constipation~~, fever, dehydration, stomatitis/**pharyngitis**, leukopenia, less energy, cough, headache
- Less Likely ( $\leq 20\%$ ): Thrombocytopenia, short-lived increases in hepatic function tests, elevated creatinine, cellulitis, and urinary tract infection, mucositis, conjunctivitis, injection site reactions, thromboembolism, edema, infection in the blood with fever, decreased kidney function, inflammation of the skin, blood clots, burning, itchy red sore eyes, dyspepsia, dysgeusia, angina, heart attack, increased heart rate, peripheral neuropathy, pneumonitis/pneumonia, insomnia, gastrointestinal bleeding, bowel obstruction, nephropathy/acute renal failure, arrhythmia, **fever, constipation**
- Rare but serious ( $<2-3\%$ ): Allergic reaction, colitis

### **Appendix I** **Consent Form**

Pages 6-7: Due to the receipt of the updated IB for Pemetrexed, the risks sections of the consent form have been updated as follows:

#### **Likely (events that occur greater than 20% of time)**

- Low white blood cells which may lead to infection (Leukopenia)
- Low red blood cells which may lead to fatigue and/or less energy (Anemia)
- Feeling sick to the stomach (nausea)
- Throwing up (vomiting)
- Loss of body fluids (dehydration)

- Loose stools (diarrhea)
- Mouth **and throat sores** ~~ulcers (similar to canker sores)~~ (Stomatitis/**Pharyngitis**) (*expanded and reworded*)
- Loss of appetite (Anorexia)
- Mild hair loss (alopecia)
- Abdominal (stomach) pain
- Itchy skin rash
- Shortness of breath (Dyspnea)
- ~~Fever (Pyrexia)~~ (*moved to Less Likely*)
- Weakness (Asthenia)
- ~~Irregular or difficulty in passing stool (constipation)~~ (*moved to Less Likely*)
- Less energy
- Cough
- Headache
- Feeling tired (Fatigue)

**Less Likely (events that occur less than or equal to 20% of time)**

- Fall in the platelet count that may lead to the risk of bruising and bleeding after an injury (Thrombocytopenia)
- Change in liver function tests (tests that show how the liver is working)
- Decrease in kidney function
- Burning, itchy, red, sore eyes with lots of watering
- Infection in the blood with fever (caused by a fall in the number of white blood cells)
- Inflammation of the skin (swelling, redness, pain)
- Infection such as urinary tract infection
- Blood clots
- Increase in body fluid (edema, swelling)
- Sores on the skin or local redness, pain, and/or swelling at the site of the injection
- Infection of the skin and surrounding tissue (Cellulitis)
- Indigestion or heartburn (Dyspepsia)
- Change in taste sensation (Dysguesia)
- Chest pain that occurs when your heart doesn't get enough oxygen. It can be a warning sign of a heart attack (Angina – unstable)
- Lack of oxygen to the heart muscle which can cause damage to the heart (Heart attack)
- Increased heart rate
- Numbness, tingling, or inflammation of the nerves (usually in the hands and feet) which may be painful (Peripheral neuropathy)
- Inflammation/infection of the lungs (Pneumonitis/pneumonia)
- Difficulty falling or staying asleep (Insomnia)
- Bleeding from the stomach, large intestine and/or small intestine (Gastrointestinal bleeding)
- Blockage in the intestines (Bowel obstruction)
- Infection around the eye, also called pink eye (Conjunctivitis)
- Damage to the kidneys, which may be temporary or permanent (Nephropathy/acute renal failure)
- Abnormal heart beat (Arrhythmia)
- **Fever (Pyrexia)** (*moved from Likely*)
- **Irregular or difficulty in passing stool (constipation)** (*moved from Likely*)

**Rare but serious**

- **Fever, chills, swelling of body, shortness of breath** (Allergic reaction) (*expanded wording*)
- Inflammation in the colon (Colitis) has been reported in patients taking Pemetrexed