

North Central Cancer Treatment Group

N0775: A Randomized Phase II Trial of Temozolomide (TMZ) and Avastin® or ABI-007/Carboplatin (CBDCA) and Avastin® in Patients with Unresectable Stage IV Malignant Melanoma

Addendum 9 – May 21, 2010

Summary

ABI-007 risks in the consent form have been updated due to the release of an updated Investigator Brochure (IB), Version 12 dated February 5, 2010.

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

Title Page Reflects the addition of Addendum 9 and a new NCI version date.

Appendix I **Consent Form**

Pages 7-8: Due to the receipt of the updated IB for ABI-007 the risks section has been updated as follows:

Likely risks of ABI-007 (events occurring greater than 20% of the time)

- Fall in the number of white blood cells leading to a greater chance of infection (Leukopenia)
- Fall in the number of red blood cells leading to anemia (feeling tired and run down) (Anemia)
- Nausea (feeling sick to your stomach)
- Vomiting (throwing up)
- Diarrhea (loose stools)
- Constipation or difficult bowel movements
- Loss of appetite (Anorexia)
- Feeling of tiredness (Fatigue)
- Hair loss (Alopecia)
- Muscle pain (Myalgia)
- Joint pain (Arthralgia)
- Nerve damage with numbness and tingling, usually in the hands and feet (Peripheral neuropathy). This effect may go away when the drug is stopped but it may not go away for some patients.
- Changes in taste sensation (Dysgeusia)

Less likely risks of ABI-007 (events occurring less than or equal to 20% of the time)

- Fever (which may indicate an infection) (Pyrexia)
- Fall in the number of platelets in your blood leading to a greater chance of bleeding (Thrombocytopenia)
- ~~Imbalance of salts or chemicals in the blood leading to kidney damage~~ (moved to Rare but serious)
- Sores in the mouth and/or throat leading to painful swallowing
- Difficult or painful breathing caused by narrowing of the airways (Dyspnea)

- ~~Changes in your liver as seen on a blood test, which may lead to skin yellowing, confusion and swelling and soreness in your belly~~ **Abnormal liver function tests which may be a sign that your liver is not working properly** (*reworded*)
- ~~Low blood pressure~~ (*deleted*)
- Abnormalities in heart rhythm (abnormal electrocardiogram) (Arrhythmia)
- Infection
- Decreased level of phosphorus in the blood (Hypophosphatemia)
- Decreased Albumin in the blood (Hypoalbuminemia)
- Abnormally low calcium in the blood stream, that can result in muscle cramps, abdominal cramps, spasms (Hypocalcemia)
- High blood sugar (Hyperglycemia)
- Low blood sugar (Hypoglycemia)
- Decreased sodium levels in the blood (Hyponatremia)
- Elevated levels of potassium in the blood (Hyperkalemia)
- Increased magnesium in the blood (Hypermagnesemia)
- Nose bleed (Epistaxis)
- Dry skin (Xerosis)
- Rash
- An infection where the nail and skin meet at the side or the base of a finger or toenail, which can be painful (Paronychia)
- Mood changes
- Eye problems, blurred vision, watery eyes, dry eyes

Rare but serious risks of ABI-007 (*events occurring less than 2-3% of the time*)

- ~~Skin changes at the site of injection~~ (*deleted*)
- ~~Slowing of the heart rate (pulse)~~ (*deleted*)
- Lack of oxygen to the heart muscle which can cause damage to the heart (Heart attack)
- Liver failure
- ~~Death~~ (*deleted*)
- **Imbalance of salts or chemicals in the blood leading to kidney damage** (*moved from Less likely*)
- **Rare reports of congestive heart failure and left ventricular dysfunction may have been observed among patients receiving Abraxane. Most of the patients were previously exposed to cardiotoxic drugs, such as anthracyclines, or had underlying cardiac history.** (*newly added*)