REDOXS© Bulletin Serious Adverse Events

The Issue: All **UNEXPECTED** and **SERIOUS** Adverse Events must be reported to CERU with 48 hours of becoming aware of the event, regardless of the relationship of the study supplements to the event.

UNEXPECTED	SERIOUS
An Unexpected adverse event is any medical occurrence that is **NOT** expected due to the progression of the underlying disease or co-morbid illnesses	A Serious Adverse Event (experience) or reaction is any untoward medical occurrence that at any dose [(1)]: • Results in death • Is life threatening • Requires or prolongs in-patient hospitalization • Results in persistent or significant disability/incapacity • May require medical or surgical intervention to prevent one of the other outcomes to defining serious.

(1) Guidance for Industry, Clinical Safety Data Management Definitions and Standards for Expedited Reporting ICH Topic E2A Health Canada 1995.

Solution To be sure that we are compliant with these guidelines, two reports must be completed:

SAE INITIAL REPORT	SAE FINAL REPORT
✓ Must be faxed to CERU **within 48 hrs** of becoming aware of the event. ✓ Must be completed by the Site Investigator or the Study Coordinator in consultation with the Site Investigator; requires the signature of the Site Investigator. ✓ Must include SAEs that occur during the study period, i.e. from the time of randomization to the end of the study period (actual ICU discharge, death or Study Day 30). ✓ Certain aspects of the form may change (for example, the resolution date may not be known at the time of reporting) and this should be made clear in Final report.	✓ Must be faxed to CERU whichever comes first: • end of study period =day 30 (from admission to ICU) • time of ICU d/c • time of ICU death • **within 15 days from becoming aware of the event *if the event is fatal or life-threatening*** ✓ Must be completed by the Site Investigator by reviewing the Initial Report and the medical chart. ✓ Must include patients admitting diagnosis, co-morbidities, a chronological complete narration of the events leading to the SAE, the nature of the SAE, action taken with the study supplements, the outcome and the relationship to the study supplements.

Concerns: All SAEs must have an assessment concerning the relationship to investigational product. The relationship of the events to the products is based on the judgment of the Site Investigator.

What to do: The relationship to study supplements must be categorized as either *not related, unlikely related, possibly related or probably related.* **In the absence of a relationship to investigational product determination, or if there is any doubt concerning the relationship assessment, the relationship should be considered "possibly related" to the investigational product.** Please provide adequate description of the issues or the site investigator's rationale for the determination.