Appendix C Safety Monitoring Plan

Adverse Event Review and Monitoring

All clinically significant adverse events, whether or not unexpected, and whether or not considered to be associated with the use of the treatment, will be documented and discussed with the ENT specialist.

All clinically significant adverse events documented will be reviewed by the ENT specialist.

Increase in Adverse Event Rates

If an increase in the frequency of Grade 3 or 4 adverse events (based on Performance Status Criteria) is noted in the study, a report should be prepared by the investigator.

If at any time the Investigator stops enrollment or stops the study due to safety issues the ENT specialist must be notified immediately via phone call.

Performance Status Criteria

|  |  |  |  |
| --- | --- | --- | --- |
| **ECOG Performance Status Scale** | | **Karnofsky Performance Scale** | |
| Grade | Descriptions | Percent | Description |
| 0 | Normal activity  Fully active, able to carry on all pre-disease performance without restriction | 100 | Normal, no complaints, no evidence of disease |
| 90 | Able to carry on normal activity; minor signs or symptoms of disease |
| 1 | Symptoms, but ambulatory  Restricted in physically strenuous activity, but ambulatory and able to carry out work of a light or sedentary nature (*e.g.*, light housework, office work) | 80 | Normal activity with effort; some signs or symptoms of disease |
| 70 | Cares for self, unable to carry on normal activity or to do active work |
| 2 | In bed < 50% of the time  Ambulatory and capable of all self-care, but unable to carry out any work activities  Up and about more than 50% of waking hours | 60 | Requires occasional assistance, but is able to care for most of his/her needs |
| 50 | Requires considerable assistance and frequent medical care |
| 3 | In bed > 50% of the time  Capable of only limited self-care, confined to bed or chair more than 50% of waking hours | 40 | Disabled, requires special care and assistance |
| 30 | Severely disabled, hospitalization indicated  Death not imminent |
| 4 | 100% bedridden  Completely disabled  Cannot carry on any self-care  Totally confined to bed or chair | 20 | Very sick, hospitalization indicated  Death not imminent |
| 10 | Moribund, fatal processes progressing rapidly |
| 5 | Dead | 0 | Dead |