## Appendix 1: Full Inclusion/Exclusion criteria for PISCES III

## **Inclusion**

- 1. Written informed consent or witnessed informed consent in the event that the subject is unable to sign informed consent.
- Ischemic stroke that includes the supratentorial region occurring within 6 to 12 months of the time that surgical intervention will be performed (Qualifying Stroke Event)
- 3. Aged between 35 and 75 (inclusive)
- 4. Qualifying Stroke Event must be confirmed by CT or MRI
- Must have current Moderate or Moderately Severe disability as measured by modified Rankin Score = 3 or 4 due to the Qualifying Stroke Event
- 6. Must have some residual upper limb movement as defined by the GRASP manual e.g. ability to actively shoulder shrug against gravity and wrist extension (palpable by the investigator; or visible lift of the fingers with hand on table)
- 7. Must have sufficient cognitive and language abilities to comprehend verbal commands for study assessments
- 8. No medical conditions that would preclude neurosurgery with appropriate preparation and management.
- 9. Sufficient putamen, globus pallidus or caudate nucleus volume to enable delivery of the CTX0E03 DP
- 10. Able to attend study related Visits and complete any diary, telephone or questionnaire assessments
- 11. Females of childbearing potential (FOCBP), (or within 2 years of last menstrual cycle) must have a confirmed negative pregnancy test at time of treatment and agree to use two r reliable methods of contraception (e.g. oral contraceptive and condom, intra-uterine device and condom, diaphragm with spermicide and condom) for six months following surgery
- 12. Sexually active males with partners who are FOCBP must be willing to use a reliable method of contraception (e.g. barrier and spermicide or as described above) for six months following surgery.

## **Exclusion**

- 1. Permanent disability corresponding to a Modified Rankin Score of >1 prior to the Qualifying Stroke Event.
- 2.Stroke due to hemorrhage or related-to connective tissue disorder, congenital disorder of the cerebral vessels or a disorder of thrombosis Subjects with atrial fibrillation as a suspected cause of stroke are NOT excluded.
- 3. Neurosurgical pathway obstructed by vascular malformation or cavity
- 4. History of neurological or other disease resulting in significant functional impairment
- 5. Any contraindications to either CT scan or MRI
- Inability to stop or transition off valproic acid or other demethylating agents or HDAC inhibitors for 1 week before and 4 weeks after treatment with CTX0E03 DP
- 7. Use of selective serotonin reuptake inhibitors (SSRI), unless the subject is on a stable dose that has been started at least 2 months before screening
- 8. Use of antispasticity medications (excluding oral antispasticity medications if they have been taken regularly for at least four months prior to treatment with CTX0E03)
- 9. Inability to discontinue anticoagulation therapy
- 10. Severe comorbid disorder that has reasonable likelihood of limiting survival to less than 24 months.
- 11. History of malignant disease within the last 5 years, (excluding benign tumors such as non-melanoma skin cancer, cervical carcinoma in situ, superficial bladder cancer)
- 12. Any history of primary or secondary brain malignant disease.
- 13. Previous participation in a cell-based therapy study at any time or in any other study involving an investigational product or rehabilitation study within the last 30 days
- 14. Clinically significant laboratory values, including positive Class I HLA antibodies specific for CTX0E03, during screening
- 15. Inability to adhere to the study post-surgery upper limb standard PT regimen e.g. excessive spasticity or pain
- 16. Planned initiation of any other new physical therapy regimen within 6-months post-treatment
- 17. Any other conditions that, in the opinion of the investigators, would preclude safe or effective participation

CT (computed tomography); CTX0E03 (investigational product); FOCBP (female of child bearing potential); GRASP (graded repetitive arm supplementary program); HDAC (histone deacetylase inhibitor); HLA (human leukocyte antigen); MRI (magnetic resonance imaging); PT (Physical Therapy)