

Reminders for Key Screening and Day 1 Procedures

Screening Period

Brain MRI

References

- Protocol 228HV101
- *MRI Acquisition Procedures Manual, Version 1*

Preparation

- Ensure imaging site has been trained and calibrated equipment according to *MRI Acquisition Procedures Manual, Version 1*.

Timing

- Complete anytime between the screening visit and Day -1 BEFORE the lumbar puncture takes place on Day -1 to Day -2
- ~45 minutes – 1 hour duration

Transfer

- Send images to NeuroRx within 24 hours

Tip: Sequences

Brain MRI will include, at a minimum, these sequences:

- T1
- T2-FLAIR
- DWI
- Gradient echo
- DTI
- T1-weighted "neuromelanin"

SPECT with DATSCAN™

References

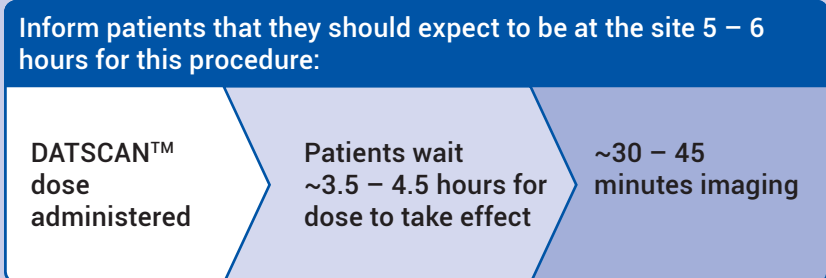
- Protocol 228HV101
- *Technical Operations Manual (TOM), Version 1.0, 09-August-2016*

Preparation

- Since DATSCAN™ doses are commercially available, allot time for imaging center to order and receive doses

Timing

Inform patients that they should expect to be at the site 5 – 6 hours for this procedure:



Transfer

- Send images to MNI within 24 hours

24 – 48 hours before IV infusion

MDS-UPDRS

Reference

- Protocol 228HV101

Rater

- Must be trained within 1 year prior to site initiation

Timing

- Anywhere from 25 – 40 minutes duration

Tip: Scheduling LP

Time of day of all LP CSF procedures throughout the study should match (± 1 hour). Choose time of day of this first LP CSF procedure with this in mind and discuss with subject prior to first LP.

Lumbar puncture (LP)

References

- Protocol 228HV101
- *CSF Sample Collection, Processing, and Shipping Instructions, Version 5.0, 03-August-2016*

Preparation

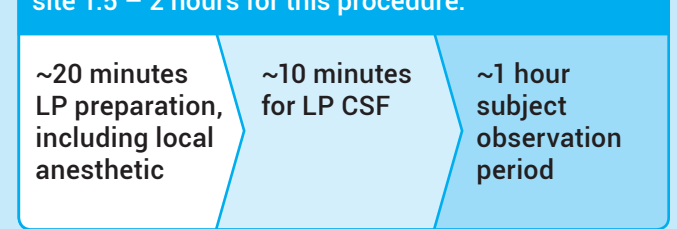
- Allot 20 minutes to prepare the patients for LB CSF
- Prepare materials, including logs and aliquot cryovials, according to *CSF Sample Collection, Processing, and Shipping Instructions*

Timing

Tip: Transportation

Due to the potential side effects after an LP CSF, such as headache, recommend to subjects that they may wish to be picked up at the site by a friend or family member.

On Day -1 to Day -2 AFTER baseline MRI Inform patients that they should expect to be at the site 1.5 – 2 hours for this procedure:



Day 1: IV infusion

References

- Protocol 228HV101

Continuous Monitoring

- Cardiac
- Adverse reactions

Timing

Assessments completed throughout Day 1 (Dosing Day) after dosing at 0 minutes:

- ≤ 10 minutes**

Assessments at this check-in point:

 - 12-lead ECG
 - Serum for PK sampling
 - Vital signs
- 30 minutes ± 5 minutes**

Assessments at this check-in point:

 - Plasma for biomarkers
 - Serum for PK sampling
 - Vital signs
- 1 hour ± 15 minutes**

Assessment at this check-in point:

 - Serum for PK sampling
- 2 hours ± 15 minutes**

Assessments at this check-in point:

 - Plasma for biomarkers
 - Serum for PK sampling
- 4 hours ± 15 minutes**

Assessments at this check-in point:

 - Serum for PK sampling
 - 12-lead ECG
 - Vital signs
- 8 hours ± 15 minutes**

Assessments at this check-in point:

 - Plasma and urine for biomarkers
 - Hematology and clinical chemistry
 - Urinalysis
 - Serum for PK sampling
- 12 hours ± 30 minutes**

Assessments at this check-in point:

 - Serum for PK sampling
 - 12-lead ECG
 - Vital signs