Myalgic Encephalomyelitis/Chronic Fatigue Syndrome (ME/CFS)
CDE Revision History Document

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January 2020 Revisions

- General Core CRF updated to replace 'Gender' question with ‘Sex assigned at birth' and 'Gender identity’.
- Two new CDEs added to CRF: C58676 (Sex assigned at birth) and C58677 (Gender identity). C00035 has been removed
- Start-Up document updated to reflect these changes.

August 2018

- Baseline/Covariate Information Subgroup
    - Updates were made to the DePaul Symptom Questionnaire (DSQ) and DePaul Pediatric Health Screening Questionnaire Informational Documents.
        - DSQ – update to the availability information and edits to the short description and scoring sections.
        - DePaul Pediatric Health Screening Questionnaire – changed name of instrument to DSQ-Pediatric Screening Questionnaire (DSQ-PSQ), updates to the availability section, edits to the short description and references and addition of special instruction, scoring and rationale/justification sections.
    - The Highlight Summary and Baseline/Covariate Information Subgroup Summary were updated to reflect the name change for the DSQ-PSQ.

July 2018

- Post-Exertional Malaise Subgroup
    - Text in the Guidance for PEM-focused Studies document was revised to remove a reference to the 2003 Canadian Consensus Criteria and add further clarification regarding the definition of PEM.
- Pain Subgroup
    - Subgroup summary was updated to include that the Fibromyalgia questions on the Pain Assessment CRF are Supplemental-Highly Recommended.

June 2018 Revisions

- Sleep Subgroup
    - Corrected Population error for all CDEs on the Sleep Questions for All Studies (F2706) and Sleep Focused Study Questionnaire (F2707). Population was changed from Adult to Adult;Pediatric.
    - Corrected Classification error for CDEs associated with the Sleep Focused Study Questionnaire CRF (F2707) as shown in the table below. The CRF, CDE Details, Sleep

Subgroup Summary and ME/CFS Start-up Resource Listing were updated to reflect this revision.

CDE ID	CDE Name	Current Classification	Corrected Classification
C17402	Site name	Supplemental	Supplemental–Highly Recommended
C19247	Subject ID		
C57101	MECFS prior history sleep problem indicator		
C57102	Sleep problem diagnosis type		
C57103	Issue present no diagnosis type		
C57104	Acclimation period before study indicator		
C57105	Acclimation process description text		
C57106	Medication help sleep problem indicator		
C57107	Medication taken neurologic psychiatric affect sleep indicator		
C57108	Medication neurologic psychiatric type text		
C57109	Medication affect sleep stop indicator		
C58439	Medication stopped other instructions given specify text		
C58438	Prior to study medication stopped duration		
C57110	Sleep wake time correspond subject sleep wake time indicator		
C57111	Objective measure heart rate variability indicator	Supplemental–Highly Recommended	Supplemental
C58440	Objective measure taken what measure text		
C58441	Objective measure method of measurement text		
C58442	Objective measure when performed text		
C58443	Objective measure describe equipment used text		

- Neurologic/Cognitive/CNS Imaging Subgroup
    - PET CRF: Updates were made to the PET CRF to reflect the current state of the science in this imaging modality and the use of PET imaging in ME/CFS. Questions regarding ligands were replaced with updated ones regarding radiopharmaceuticals. Questions related to uptake, distribution, and lateralization were removed. Questions related to hardware, participant groups, and acquisition were added. The question regarding study conclusion as it relates to detecting the presence or absence of something such as a tumor was replaced with a study conclusion question related to the detection of difference/abnormalities.

CDE ID	CDE Name	Added/Removed/Kept
C02494	Imaging study date and time	Kept (Question text changed to match CRF)
C08258	Imaging radioligand type	Removed

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CDE ID	CDE Name	Added/Removed/Kept
C02494	Imaging study date and time	Kept (Question text changed to match CRF)
C18711	Imaging radioligand other text	Removed
C08261	Imaging radioligand dose value	Removed
C13996	PET localization uptake type	Removed
C13997	PET localization distribution pattern type	Removed
C02411	Laterality type	Removed
C13998	PET localization cortical anatomic site	Removed
C18529	Imaging positron emission tomography ictal cortical anatomic site	Removed
C13999	PET localization study type	Removed
C02494	Imaging study date and time	Kept (Question text changed to match CRF)
C58527	Imaging radiopharmaceutical name	Added
C58528	Imaging radiopharmaceutical manufacturer text	Added
C58529	Imaging radiopharmaceutical synthesis process text	Added
C58530	Imaging radiopharmaceutical synthesis process source text	Added
C58531	Imaging radiopharmaceutical mechanism of action text	Added
C58532	Imaging radiopharmaceutical synthesis administration between average duration	Added
C58533	Imaging radiopharmaceutical dose	Added
C58534	Imaging radiopharmaceutical molar activity rate	Added
C58535	Imaging radiopharmaceutical mass inject measurement	Added
C58536	Imaging radiochemical minimum yield value	Added
C58537	Imaging radiochemical minimum purity percentage value	Added
C58538	Imaging radiopharmaceutical administration type	Added
C58539	Imaging radiopharmaceutical infusion duration	Added
C02496	Imaging scanner manufacturer name	Added
C17939	Scanner manufacturer other name	Added
C20247	Imaging scanner model name text	Added
C58540	Imaging participant group type	Added
C58541	Imaging patient participant group diagnosis text	Added
C58542	Imaging experiment condition task indicator	Added
C58543	Imaging experiment condition task text	Added
C58544	Imaging scan location type	Added
C08242	Imaging acquisition duration	Added
C58545	Imaging matrix size axis 1 measurement	Added
C58546	Imaging matrix size axis 2 measurement	Added
C58547	Imaging matrix size axis 3 measurement	Added
C10838	Imaging voxel size axis 1 measurement	Added

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CDE ID	CDE Name	Added/Removed/Kept
C02494	Imaging study date and time	Kept (Question text changed to match CRF)
C17858	Imaging voxel size axis 2 measurement	Added
C17859	Imaging voxel size axis 3 measurement	Added
C58548	Imaging Gaussian filter type	Added
C08244	Imaging slice thickness value	Added
C58549	Imaging acquisition plane type	Added
C58550	Imaging acquisition plane other text	Added
C58551	Imaging acquisition correct apply indicator	Added
C58552	Imaging acquisition correct apply text	Added
C58553	Imaging patient prepare text	Added
C58554	Imaging dynamic acquisition indicator	Added
C17931	Imaging magnetic resonance imaging performed indicator	Added
C58555	Imaging activity arterial blood measure indicator	Added
C58556	Imaging activity peripheral blood measure indicator	Added
C58557	Imaging metabolite measure indicator	Added
C58558	Imaging pharmacologic effect observe indicator	Added
C58559	Imaging adverse event radiopharmaceutical relate indicator	Added
C08236	Imaging scanner software name	Added
C02498	Imaging scanner software version number	Added
C13994	Imaging PET localization processing type	Kept (Removed SPM and added Other, specify to PVs)
C56595	Imaging PET localization processing other text	Added
C13995	PET localization rating type	Kept (Question text changed to match CRF)
C58560	Imaging analysis reference region use indicator	Added
C08270	Imaging reference region text	Added
C58561	Imaging analysis software package name	Added
C10843	Imaging analysis software version number	Added
C58562	Imaging study conclusion type	Added

May 2018 Revisions

- ME/CFS Data Standards webpage
  - Revised the Prospective Assessments for Suicidal Ideation and Behavior section to "Investigators should review the FDA's guidance regarding suicidal ideation and behavior."
- Start-up Resource Listing and Highlight Summary Documents
  - Removed FDA guidance regarding suicidal ideation and behavior in clinical trials language.

- Overall Working Group Summary
  - Updated language to describe disease prevalence, variation in case definitions, stratification of participants, and the need for validating how well measures reflect patients' experience of ME/CFS.
- Baseline/Covariate Information Subgroup
  - Added Modifiable Activity Questionnaire (MAQ) as Supplemental under the Outcomes and Endpoints Domain/ Activities of Daily Living/ Performance Sub-Domain. The Highlight Summary and Subgroup Summary were updated to include this new recommendation.
- Pain Subgroup
  - Revised CDEs to match CRF

CDE ID	CDE Name	Added, Removed or Modified	Modification
C22622	Wong-Baker FACES - Pain rating scale	Removed	
C22804	Pain present eleven-point intensity scale	Added	
C55573	Pain intensity visual analogue scale value	Added	
C58517	Tender point pair cervical vertebrae type	Added	
C58518	Tender point pair trapezius type	Added	
C58519	Tender point pair second rib type	Added	
C58520	Tender point pair lateral epicondyle type	Added	
C58521	Tender point pair knee type	Added	
C58522	Tender point pair occiput type	Added	
C58523	Tender point pair supraspinatus type	Added	
C58524	Tender point pair gluteal muscle type	Added	
C58525	Tender point pair trochanter major type	Added	
C58460	Fibromyalgia diagnosis criterion type	Modified	1. Definition is a duplicate of the Fibromyalgia diagnosis indicator CDE definition Previously: The indicator related to the diagnosis for fibromyalgia. Currently: The type of criterion used to diagnose fibromyalgia. 2. Added Other, specify to the PVs as listed on the CRF.
C58526	Fibromyalgia diagnosis other text	Added	

  - Updates to the Pain Assessment CRF – corrected grammar, added 2016 criterion as a PV to the Fibromyalgia diagnosis criterion type question to match the corresponding CDE, and added asterisk to Supplemental-Highly Recommended fibromyalgia questions.
- Neurologic/Cognitive/CNS Imaging Subgroup

- Updated Beck Anxiety Inventory NOC to include information about the overestimation of anxiety severity in patients with Postural Tachycardia Syndrome (POTS) and the corresponding reference.
- Updated language in subgroup summary from first person to third person.

April 2018 Revisions

- Updates to the Start-up Resource Listing and Highlight Summary Documents
  - Revised Fatigue Severity Scale of Sleep Disorders to Fatigue Severity Scale as the copyrighted name of the instrument is: Fatigue Severity Scale.
  - Included note that CDEs on Imaging CRFs are all Supplemental – Highly Recommended and are not listed individually.
  - Deleted NHANES CDEs as they are now classified as Exploratory.
  - Added Core and S-Highly Recommended CDEs that were missing from the list:

CDEID	CDE Name	Domain; Sub-Domain; Module
C00015	Education year count	Participant Characteristics; Demographics; General Core
C01521	Heart rate	Assessments and Examinations; Autonomic; Passive Standing Test Protocol
C01539	Temperature measurement	Assessments and Examinations; Physical/Neurological Examination; Physical Exam - Immune Module
C02014	Medication prior or concomitant name	Treatment/Intervention Data; Drugs; Medication and Other Treatments
C17402	Site name	Outcomes and End Points; Sleep; Sleep Questions For All Studies
C19247	Subject ID	Outcomes and End Points; Sleep; Sleep Questions For All Studies
C18027	Comment text	Assessments and Examinations; Autonomic; Passive Standing Test Protocol
C19500	Assessment performed date	Treatment/Intervention Data; Drugs; Medication and Other Treatments
C57889	Cerebellar gait normal gait assessment status	Assessments and Examinations; Physical/Neurological Examination; Physical Examination
C57891	Cerebellar tandem Romberg assessment status	Assessments and Examinations; Physical/Neurological Examination; Physical Examination
C58440	Objective measure taken what measure text	Outcomes and End Points; Sleep; Sleep Focused Study Questionnaire
C58441	Objective measure method of measurement text	Outcomes and End Points; Sleep; Sleep Focused Study Questionnaire
C58442	Objective measure when performed text	Outcomes and End Points; Sleep; Sleep Focused Study Questionnaire
C58443	Objective measure describe equipment used text	Outcomes and End Points; Sleep; Sleep Focused Study Questionnaire

- Fatigue Subgroup

- Fatigue Severity Scale - Corrected name of Fatigue Severity Scale by removing "of Sleep Disorders" in the NOC, and Subgroup Summary Document and updated the availability link.
- Checklist for Individual Strength-Fatigue (CIS) - revised NOC to remove cut-offs per subgroup in response to internal review comments. Deleted:

Section	Text
Scoring	"(e.g., subjective fatigue cut-off= 35 or 40 for CFS)"
Rationale/Justification	"therefore higher score of 40 suggested for CFS for subjective fatigue."
Rationale/Justification	"The 35 points cut-off score for severe fatigue is appropriate, but, given the 17% false positive rate, should be adjusted to 40 for research in CFS. (Worm-Smeitink et al., 2017)."

- Sleep Subgroup
  - Updated subgroup summary: Changes include rewording, and adding purpose for instruments.
  - Updated Sleep Questions for All Studies CRF: Reorganization of questions and changes to classifications and instructions. The DSQ and treatment questions are Core and the NHANES questions are Exploratory. The instructions were updated to match.
  - Updated Sleep Questions for All Studies CDEs to match changes on CRF: The DSQ and treatment questions remained Core and the NHANES questions are now Exploratory. Question order was updated to match reorganization of CRF.
- Pain Subgroup
  - Updated subgroup summary to include information regarding the assessment of fibromyalgia, an explanation of how pain subgroup recommendations fit with other subgroup recommendations, and noted the need for a method for quantitatively assessing pain.
- Neuroendocrine Subgroup
  - Updated population for the following CDEs from Adult to Adult; Pediatric on the Neuroendocrine Labs CRF:
    - C10896 – Pregnancy test performed indicator
    - C01714 – Pregnancy test not applicable reason
    - C01702 – Pregnancy test date and time
    - C01704 – Pregnancy test specimen type
    - C01710 – Pregnancy test qualitative result value
- Biomarkers Subgroup
  - Corrected page numbering footers.

March 2018 Revisions

- ME/CFS Overall Working Group Summary added to the website.
- Hyperlink to CDE definition was added to the Data Standards Page.
- Revisions to the Roster on the History and Acknowledgments Page:
  - Corrected Dr. Younger's degree to PhD.
  - Removed duplicate listing for Dr. Sullivan under the Neurologic/Cognitive/CNS Imaging Subgroup.
  - Corrected Dr. Biagionni's degree to MD.

- Baseline/Covariate Information Subgroup
  - Update name of the Pediatric ME/CFS Screening Questionnaire to the DePaul Pediatric Health Screening Questionnaire. This change was made in the NOC as well as on the Subgroup Summary and Highlight Document.
  - QC of the following NOCs:
    - DePaul Symptom Questionnaire (DSQ)
    - ME/CFS Symptom Checklist
    - DePaul Pediatric Health Screening Questionnaire
- Fatigue Subgroup
  - PROMIS Fatigue Short Form NOC – spelled out computerized adaptive testing (CAT) the first time it is used.
  - Modified Fatigue Impact Scale (MFIS) NOC - Replaced subscript references with parenthetical and deleted duplicate references
- Sleep Subgroup
  - QC of the following NOCs:
    - Stanford Sleepiness Scale
    - Sleep Disorders Screening Checklist
    - Holland Sleep Disorders Questionnaire (HSDQ)
    - Epworth Sleepiness Scale - Children's Version
    - Nonrestorative Sleep Scale (NRSS)
    - Global Sleep Assessment Questionnaire (GSAQ)
    - Restorative Sleep Questionnaire (RSQ)
    - Adolescent Sleep Hygiene Scale (ASHS)
- Pain Subgroup
  - 2016 Fibromyalgia criterion added as a PV and references updated for C58460 Fibromyalgia diagnosis criterion type
- Biomarkers Subgroup
  - The link, http://www.equator-network.org/, was revised in the F2795 Biomarkers Guidelines and F2796 Biomarkers Reference Table.
  - Update the following documents on the ME/CFS page under the Laboratory Tests and Biospecimens/Biomarkers subdomain:
    - F2743 Biomarker-Related Sample and Collection Questions: Add questions on type of vial used for specimen collection.
    - F2795 Biomarkers Guidelines: Edits to the Gene expression/transcriptome section to include details on quantitative polymerase chain reaction (qRT-PCR) and Next Generation sequencing (NGS).
    - CDEs for F2743 also revised:
      - C57377: PV added for dose decreased
      - CDEs added:
        - C58461
        - C57375
        - C58463
        - C58464
        - C58466
        - C58467
        - C58468

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