SUPPLEMENTARY MATERIAL

Reconstruction of Quantitative Susceptibility Mapping and Susceptibility Map-Weighted Imaging

The k-space data of the multicho data image combination images were transferred to a personal computer. For quantitative susceptibility mapping (QSM) reconstruction, the STI Suite software package^{1,2} was used with error tolerances of 0.01 and a threshold for a binary mask of the ill-conditioned k-space region of 0.1. The mask for the STI suite was automatically determined using an active contour algorithm.³ Susceptibility map-weighted imaging (SMWI),⁴ which is a recently proposed susceptibility contrast enhancing method that multiplies a QSM-generated mask with a magnitude (or root sum of squares of multi-echo magnitude) image, was reconstructed using the QSM and magnitude images. This approach has demonstrated to provide an improved contrast to noise ratio of the nigrosome-1. More detailed data processing and reconstruction parameters can be found elsewhere.⁴ SMWI was used for visual assessment of the substantia nigra (SN).

The reconstructed QSM images were transferred to a multimodality solution (Syngo.via version VA30, Siemens Healthcare, Forchheim, Germany) for reslicing of the images at an increment of 0.5 mm. As not all of the QSM images were symmetric due to the participant's head shape and/or position, they were repositioned to be as symmetric as possible at the level of the lower border of the red nucleus. This procedure was done to help investigators draw the regions of interest (ROIs) by comparing each side, particularly in healthy subjects.

For normalization of the QSM susceptibility values, an oval ROI greater than 10 mm² was placed in the region of the decussation of the superior cerebellar peduncle on each resliced QSM images. This region is presumed to be located in the center of the midbrain at the level of inferior tip of the SN, which was determined by comparison with sagittal magnetization-prepared rapid gradient-echo imaging (Supplementary Fig. 2 in the online-only Data Supplement). We chose this region because it was close to the SN and is less affected by magnetic susceptibility anisotropy due to its horizontal direction.⁵ Normalization was achieved by subtracting the mean susceptibility values measured in the decussation of the superior cerebellar peduncle.

REFERENCES

- 1. Li W, Wang N, Yu F, Han H, Cao W, Romero R, et al. A method for estimating and removing streaking artifacts in quantitative susceptibility mapping. *Neuroimage* 2015;108:111-122.
- Li W, Avram AV, Wu B, Xiao X, Liu C. Integrated Laplacian-based phase unwrapping and background phase removal for quantitative susceptibility mapping. NMR Biomed 2014;27:219-227.
- 3. Chan TF, Vese LA. Active contours without edges. IEEE Trans Image Process 2001;10:266-277.
- 4. Nam Y, Gho SM, Kim DH, Kim EY, Lee J. Imaging of nigrosome 1 in substantia nigra at 3T using multiecho susceptibility map-weighted imaging (SMWI). J Magn Reson Imaging 2017;46:528-536.
- 5. Lee J, Shmueli K, Fukunaga M, van Gelderen P, Merkle H, Silva AC, et al. Sensitivity of MRI resonance frequency to the orientation of brain tissue microstructure. *Proc Natl Acad Sci U S A* 2010;107:5130-5135.