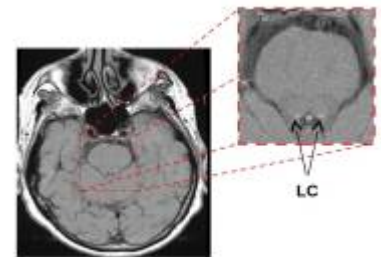


# Neuromelanin magnetic resonance imaging

Neuromelanin-sensitive MRI (NM-MRI) purports to detect the content of **neuromelanin** (NM), a product of **dopamine** metabolism that accumulates with age in dopamine neurons of the **substantia nigra** (SN). Interindividual variability in dopamine function may result in varying levels of NM accumulation in the SN; however, the ability of NM-MRI to measure dopamine function in nonneurodegenerative conditions has not been established.



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Neuromelanin sensitive MRI may be the method of choice for the follow-up of **meningeal melanocytoma** <sup>1)</sup>.

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Cassidy et al. validated that NM-MRI **signal intensity** in postmortem **midbrain** specimens correlated with regional NM concentration even in the absence of **neurodegeneration**, a prerequisite for its use as a proxy for dopamine function. They then validated a **voxelwise** NM-MRI approach with sufficient anatomical sensitivity to resolve SN subregions. Using this approach and a multimodal dataset of molecular **PET** and **fMRI** data, they further showed the NM-MRI signal was related to both **dopamine** release in the dorsal striatum and resting blood flow within the SN. These results suggest that NM-MRI signal in the SN is a proxy for function of dopamine neurons in the nigrostriatal pathway. As a proof of concept for its clinical utility, we show that the NM-MRI signal correlated to severity of psychosis in schizophrenia and individuals at risk for schizophrenia, consistent with the well-established dysfunction of the nigrostriatal pathway in psychosis. The results indicate that noninvasive NM-MRI is a promising tool that could have diverse research and clinical applications to investigate in vivo the role of dopamine in neuropsychiatric illness <sup>2)</sup>.

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A study aimed to evaluate the accuracy and diagnostic test performance of the U-net-based segmentation method in neuromelanin magnetic resonance imaging (NM-MRI) compared to the established manual segmentation method for **Parkinson's disease diagnosis**.

NM-MRI datasets from two different 3T-scanners were used: a “principal dataset” with 122 participants and an “external validation dataset” with 24 participants, including 62 and 12 PD patients, respectively. Two radiologists performed SNpc manual segmentation. Inter-reader precision was determined using Dice coefficients. The U-net was trained with manual segmentation as ground truth and Dice coefficients used to measure accuracy. Training and validation steps were performed on the principal dataset using a 4-fold cross-validation method. We tested the U-net on the external validation dataset. SNpc hyperintense areas were estimated from U-net and manual segmentation masks, replicating a previously validated thresholding method, and their diagnostic test performances for PD determined.

For SNpc segmentation, U-net accuracy was comparable to inter-reader precision in the principal dataset (Dice coefficient: U-net,  $0.83 \pm 0.04$ ; inter-reader,  $0.83 \pm 0.04$ ), but lower in external validation dataset (Dice coefficient: U-net,  $0.79 \pm 0.04$ ; inter-reader,  $0.85 \pm 0.03$ ). Diagnostic test

performances for PD were comparable between U-net and manual segmentation methods in both principal (area under the receiver operating characteristic curve: U-net, 0.950; manual, 0.948) and external (U-net, 0.944; manual, 0.931) datasets.

U-net segmentation provided relatively high accuracy in the evaluation of the SNpc in NM-MRI and yielded diagnostic performance comparable to that of the established manual method<sup>3)</sup>.

## References

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