

Table S1 – ROI and whole-brain cross-sectional studies*

Author	Year	Participants			ROI analysis				Whole-brain analysis				Threshold	
		Controls	PreHD	HD	FA	MD	AD	RD	FA	MD	AD	RD		
Mascalchi	2004	21	2	19	N/A	HDGC: ↑ in BG, deep WM and whole WM	N/A	N/A	N/A	N/A	N/A	N/A	N/A	p<0.05
Reading	2005	7	7	Not included	N/A	N/A	N/A	N/A	PreHD: ↓ in the superior frontal, post-central and pre-central WM	N/A	N/A	N/A	N/A	p<0.005 uncorrected, k>50 voxels
Rosas	2006	29	15	17	PreHD and HD: ↓ in CC and ↑ IC and BG	N/A	N/A	N/A	PreHD and HD: ↑ in BG ↓ in CC, EC, cerebral peduncles, brainstem, thalamus, subcortical WM	PreHD and HD: ↑ in BG	N/A	N/A	N/A	p<0.05
Kloppel	2008	20	25	Not included	N/A	PreHD: ↓ in BG	N/A	N/A	N/A	N/A	N/A	N/A	N/A	p<0.001
Douaud	2009	10	Not included	14	HD: ↑ in BG	HD: ↑ in BG	N/A	N/A	HD: ↑ in BG	HD: ↑ in BG, CR, subcortical WM	N/A	N/A	N/A	<u>WHOLE-BRAIN</u> p<0.01 FDR corrected for the MD maps p<0.05 FDR corrected for FA maps <u>ROI</u> p<0.05
Mandelli	2009	25	15	9	N/A	HD: ↑ in BG PreHD: ↓ in BG	N/A	N/A	N/A	N/A	N/A	N/A	N/A	p<0.05

Della Nave	2010	15	1	14	N/A	N/A	N/A	N/A	HDGC: ↓ in CC, fornix, EC, IFOF, ILF and ↑ in supralenticular frontal WM	HDGC: ↑ areas with ↓ FA plus AF and CP	HDGC: ↑ in areas with ↓ FA	HDGC: Similar to MD	p<0.05 TFCE corrected
Sritharan	2010	17	Not included	18	N/A	HD: ↑ in BG, thalamus and CC	N/A	N/A	N/A	N/A	N/A	N/A	p<0.005 Bonferroni corrected
Stoffers	2010	25	39	Not included	N/A	N/A	N/A	N/A	PreHD: ↓ widespread	PreHD: ↑ widespread	PreHD: ↑ thalamic radiations, IC, ED	PreHD: ↑ widespread	p<0.05
Rosas	2010	40	19	21	PreHD and HD: ↓ in CC	N/A	PreHD and HD: ↑ in CC	PreHD and HD: ↑ in CC	N/A	N/A	N/A	N/A	p<0.05 Bonferroni-corrected
Bohanna	2011	17	Not included	10	HD: ↓ CC, fornix, EC, CR and SS	HD: ↑ same as ↓ FA plus IC and cerebral peduncles	HD: ↑ CR, IC, EC, fornix	HD: ↑ CC, fornix, EC, IC, CR, SS, cingulum, SLF	N/A	N/A	N/A	N/A	p<0.05 corrected for multiple comparisons cluster forming threshold of t>3.
Di Paola	2012	34	17	17	N/A	N/A	N/A	N/A	PreHD: ↓ in CC HD: ↓ in CC	N/A	PreHD: No differences HD: ↑ in CC	PreHD: ↑ in CC HD: ↑ in CC	p<0.05 TFCE corrected
Hobbs	2013	40	Not included	61	HD: No differences	HD: ↑ in BG	HD: ↑ in BG	HD: ↑ in BG	HD: ↑ in BG in HD ↓ in WM	HD: ↑ widespread	HD: ↑ widespread	HD: ↑ widespread	<u>WHOLE-BRAIN ROI</u> p<0.001 p<0.05
Delmaire	2013	24	Not included	27	N/A	N/A	N/A	N/A	HD: ↑ in BG and ↓ in frontal lobe, CR	HD: ↑ widespread	N/A	N/A	p<0.05 FWE corrected

Georgiou-Karistianis	2013	36	35	36	PreHD and HD: ↑ in BG	PreHD and HD: ↑ in BG	N/A	N/A	N/A	N/A	N/A	N/A	p<0.05
Sanchez-Castaneda	2013	30	17	12	PreHD: ↑ in BG HD: ↑ in BG and ↓ in thalamus and hippocampus	PreHD and HD: ↑ MD in BG, hippocampus	N/A	N/A	N/A	N/A	N/A	N/A	p<0.05 Bonferroni corrected
Matsui	2014	34	53	Not included	PreHD: No differences	PreHD: No differences	PreHD: No differences	PreHD: ↑ orbitofrontal and a inferior frontal lobe region	N/A	N/A	N/A	N/A	p<0.05 FDR corrected
Novak	2014	21	17	19	N/A	N/A	N/A	N/A	HDGC: ↓ CC, SLF, ILF, IC, EC, cingulum	HDGC: ↑ CC, ILF, SLF, IC, EC, cingulum, AF and CP	N/A	N/A	p<0.05 FWE corrected
Phillips	2014	50	25	25	PreHD: No differences HD: ↓ in deep WM	N/A	PreHD: ↑ in deep WM HD: ↑↑ in deep WM	PreHD: ↑ in deep WM HD: ↑↑ in deep WM	N/A	N/A	N/A	N/A	p<0.05 FDR corrected
Odish	2014	24	22	10	N/A	N/A	N/A	N/A	PreHD: No differences total WM HD: No differences in total WM	PreHD: No differences in total GM or total WM HD: ↑ in total GM and total WM	PreHD: ↑ in total WM, no differences in total GM HD: ↑ in total GM and total WM	PreHD: No differences in total GM or total WM HD: ↑ in total GM and total WM	p≤0.001 Bonferroni corrected
Rees	2014	12	Not included	22	HD: ↓ in cerebellar GM	HD: ↑ in cerebellar GM and WM	HD: ↑ in cerebellar GM and WM	HD: ↑ in cerebellar GM and WM	N/A	N/A	N/A	N/A	p<0.05

Gregory	2015	36	Not included	48	HD: ↓ in CC and brainstem	N/A	HD: ↑ in CC, CR, EC, IC, PTR, SLF, SS	HD: ↑ in CC, CR, EC, IC, SLF, brainstem	N/A	N/A	N/A	N/A	p<0.05 FDR corrected
Syka	2015	14	Not included	14	HD: No differences	HD: No differences	N/A	N/A	N/A	N/A	N/A	N/A	p<0.05 FDR corrected
Faria	2016	79	194	Not included	PreHD: No differences	PreHD: ↑ in deep WM, corticostriatal tract	PreHD: ↑ in deep WM	PreHD: ↑ in deep WM and corticostriatal tract	PreHD: No differences	PreHD: ↑ in PTR, CC and occipital WM	Same as MD	Same as MD	p<0.05 FWE corrected
Harrington	2016	67	37	Not included	PreHD: ↓ in all tracts	PreHD: ↑ in all tracts	PreHD: ↑ in all tracts	PreHD: ↑ in all tracts	N/A	N/A	N/A	N/A	p<0.05 FDR corrected
Muller	2016	32	Not included	34	N/A	N/A	N/A	N/A	HD: ↑ in BG and ↓ in CC, EC, IC and thalamus	N/A	HD: ↑ in BG and dorsal WM	HD: ↑ in BG and dorsal WM	p<0.05 FDR corrected
Gorges	2017	13	12	Not included	N/A	N/A	N/A	N/A	PreHD: No differences	N/A	PreHD: No differences	PreHD: No differences	p<0.05 FDR corrected
Saba	2017	11	12	11	N/A	N/A	N/A	N/A	PreHD: No differences HD: ↓ in CC, EC, IC, ATR, PTR, SLF, ILF, IFOF	N/A	N/A	N/A	p<0.05 FWE corrected
Gregory	2020	20	20	40	N/A	N/A	N/A	N/A	HDGC: ↓ widespread	HDGC: ↑ in CC	HDGC: ↑ in subcortical WM	HDGC: ↑ in CC	p<0.05 TFCE corrected
Sweidan	2020	11	Not included	13	N/A	N/A	N/A	N/A	HD: ↓ widespread	HD: ↓ widespread	N/A	N/A	p<0.05 TFCE corrected

*Only bilateral differences noted. HDGC used when manifest HD and preHD were grouped together in a single group

AD: Axial diffusivity, AF: Arciform Fibers, ATR: Anterior Thalamic Radiations, CC: Corpus Callosum, CP: Cerebral Peduncles, CR: Corona Radiata, CST: Corticospinal Tract, DTI: Diffusion Tensor Imaging, FA: Fractional Anisotropy, IC: Internal Capsule, EC: External Capsule, HD: Huntington's Disease, HDGC: Huntington's disease gene-carriers, IFOF: Inferior Fronto Occipital Fasciculus, ILF: Inferior Longitudinal Fasciculus, MD: Mean Diffusivity, preHD: Premanifest Huntington's Disease, PTR: Posterior Thalamic Radiations, RD: Radial Diffusivity, SS: Sagittal Stratum, SLF: Superior Longitudinal Fasciculus, SFOF: Superior Fronto Occipital Fasciculus, UNC: uncinata fasciculi.