

Plasma phospho-tau181 in familial Alzheimer's disease

EYO (years)	P-tau181 (p-value)	NfL (p-value)
-20	0.27	0.18
-19	0.20	0.12
-18	0.13	0.065
-17	0.085	0.033
-16	0.050	0.015
-15	0.027	0.0055
-14	0.014	0.0017
-13	0.0061	0.00046
-12	0.0025	0.00010
-11	0.00096	<0.0001
-10	0.00033	<0.0001
-9	0.00011	<0.0001

Supplementary Table 1S: Evidence for divergence between mutation carriers and non-carriers

in plasma biomarkers at sequential EYO timepoints

P-values from tests of whether the estimated difference between carriers and non-carriers for geometric mean i) plasma p-tau181 and ii) plasma NfL is statistically significantly different from zero, after adjusting for age and gender, for integer values of EYO between –20 to –9 years. P-values expressed to two significant figures unless $p < 0.0001$.

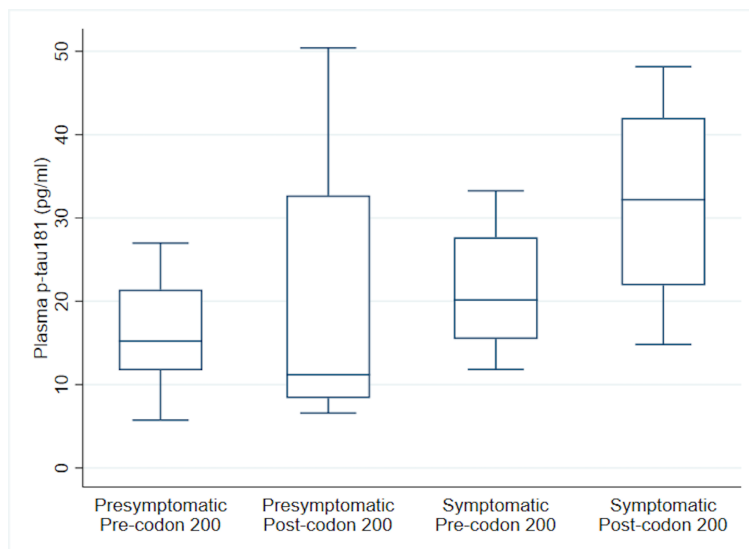
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Median/IQR of person-level p-tau181 medians		
	Presymptomatic	Symptomatic
<i>APP</i>	14.5 (8.4, 17.6) N=6	17.0 (16.4, 19.0) N=5
<i>PSEN1</i>	14.2 (9.9, 19.9) N=18	25.1 (19.1, 39.8) N=14
<i>PSEN1</i> Pre-codon 200	15.3 (13.5, 18.6) N=10	23.5 (18.3, 29.5) N=8
<i>PSEN1</i> Post-codon 200	11.2 (9.7, 22.8) N=8	32.2 (21.9, 42.0) N=6

Supplementary Table 2S: Plasma p-tau181 levels according to genetic subgroup

Median and IQRs of observed plasma p-tau181 values grouped by genetic mutation and mutation position. All data were used and medians of person-level medians calculated to allow for repeated measures at the individual level.

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Supplementary Figure 15: Observed baseline p-tau181 values for *PSEN1* individuals by disease status and by mutation position pre- vs. post-codon 200. A box shows the median, and 25th and 75th percentiles; the whiskers extend to the largest value less than or equal to the upper quartile + 1.5 times the interquartile range, and the smallest value greater than or equal to the lower quartile - 1.5 times the interquartile range; values outside this range are shown individually.