

# Risk Polymorphisms of *FNDC5*, *BDNF*, and *NTRK2* and Poor Education Interact and Aggravate Age-Related Cognitive Decline

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**Supplementary Table S1.** Significant CANTAB results in three-way ANOVA. Values are shown as mean Z score  $\pm$  SD.

Tests	Education	Age	SNP NTRK2 (n)	Mean $\pm$ SD	CI 95%	d
PAL total errors adjusted (score)	Lower	Young	AA+AG (25)	-0.73 $\pm$ 0.40	-0.23, 0.44	0.32
			GG (31)	-0.83 $\pm$ 0.22		
		Older	AA+AG (16)	1.01 $\pm$ 1.14	-0.64, 0.18	0.23
			GG (21)	1.24 $\pm$ 0.92		
	Higher	Young	AA+AG (20)	-0.86 $\pm$ 0.27	-0.34, 0.37	0.05
			GG (31)	-0.87 $\pm$ 0.18		
		Older	AA+AG (28)	<b>0.86 <math>\pm</math> 0.97</b>	<b>0.59, 1.27***</b>	<b>1.19</b>
			GG (27)	<b>-0.07 <math>\pm</math> 0.52</b>		
PAL mean trials to success (score)	Lower	Young	AA+AG (25)	-0.83 $\pm$ 0.32	-0.26, 0.35	0.16
			GG (31)	-0.88 $\pm$ 0.29		
		Older	AA+AG (16)	<b>0.87 <math>\pm</math> 0.94</b>	<b>-0.88, -0.12**</b>	<b>0.58</b>
			GG (21)	<b>1.38 <math>\pm</math> 0.83</b>		
	Higher	Young	AA+AG (20)	-0.94 $\pm$ 0.40	-0.32, 0.33	0.03
			GG (31)	-0.95 $\pm$ 0.22		
		Older	AA+AG (28)	<b>0.83 <math>\pm</math> 0.77</b>	<b>0.45, 1.07***</b>	<b>1.07</b>
			GG (27)	<b>0.06 <math>\pm</math> 0.67</b>		
PAL first trial memory score (score)	Lower	Young	AA+AG (25)	0.74 $\pm$ 0.72	-0.46, 0.28	0.12
			GG (31)	0.83 $\pm$ 0.77		
		Older	AA+AG (16)	-1.02 $\pm$ 0.80	-0.57, 0.34	0.16
			GG (21)	-0.90 $\pm$ 0.72		
	Higher	Young	AA+AG (20)	0.99 $\pm$ 0.63	-0.32, 0.46	0.10
			GG (31)	0.92 $\pm$ 0.79		
		Older	AA+AG (28)	<b>-0.82 <math>\pm</math> 0.57</b>	<b>-1.23, -0.48***</b>	<b>1.54</b>
			GG (27)	<b>0.03 <math>\pm</math> 0.53</b>		

Significant results in bold. \*:  $p \leq 0.05$ ; \*\*:  $p \leq 0.01$ ; \*\*\*:  $p \leq 0.001$ . CI: confidence interval; d: effect size - Cohen's d. SNP: single nucleotide polymorphism.

**Supplementary Table S2.** Significant CANTAB results in three-way ANOVA with Bootstrapping. Values are shown as mean Z score  $\pm$  SD.

Tests	Education	Age	SNP NTRK2 (rs2289656)	Mean $\pm$ SD	CI 95%	d
<b>With Bootstrapping 5000 samples</b>						
<b>PAL total errors adjusted (score)</b>	<b>Lower</b>	Young	AA+AG	-0.73 $\pm$ 0.40	-0.49, 0.29	0.32
			GG	-0.83 $\pm$ 0.22		
		Older	AA+AG	1.01 $\pm$ 1.14	-0.90, 0.44	0.23
			GG	1.24 $\pm$ 0.92		
	<b>Higher</b>	Young	AA+AG	-0.86 $\pm$ 0.27	-0.09, 0.14	0.05
			GG	-0.87 $\pm$ 0.18		
		<b>Older</b>	<b>AA+AG</b>	<b>0.86 <math>\pm</math> 0.97</b>	<b>0.53, 1.37***</b>	<b>1.19</b>
			<b>GG</b>	<b>-0.07 <math>\pm</math> 0.52</b>		
<b>PAL mean trials to success (score)</b>	<b>Lower</b>	Young	AA+AG	-0.83 $\pm$ 0.32	-0.11, 0.21	0.16
			GG	-0.88 $\pm$ 0.29		
		Older	AA+AG	0.87 $\pm$ 0.94	-1.11, 0.07	0.58
			GG	1.38 $\pm$ 0.83		
	<b>Higher</b>	Young	AA+AG	-0.94 $\pm$ 0.40	-0.18, 0.20	0.03
			GG	-0.95 $\pm$ 0.22		
		<b>Older</b>	<b>AA+AG</b>	<b>0.83 <math>\pm</math> 0.77</b>	<b>0.38, 1.15***</b>	<b>1.07</b>
			<b>GG</b>	<b>0.06 <math>\pm</math> 0.67</b>		
<b>PAL first trial memory score (score)</b>	<b>Lower</b>	Young	AA+AG	0.74 $\pm$ 0.72	-0.47, 0.30	0.12
			GG	0.83 $\pm$ 0.77		
		Older	AA+AG	-1.02 $\pm$ 0.80	-0.60, 0.39	0.16
			GG	-0.90 $\pm$ 0.72		
	<b>Higher</b>	Young	AA+AG	0.99 $\pm$ 0.63	-0.31, 0.45	0.10
			GG	0.92 $\pm$ 0.79		
		<b>Older</b>	<b>AA+AG</b>	<b>-0.82 <math>\pm</math> 0.57</b>	<b>-1.13, -0.59***</b>	<b>1.54</b>
			<b>GG</b>	<b>0.03 <math>\pm</math> 0.53</b>		

Significant results in bold. \*:  $p \leq 0.05$ ; \*\*:  $p \leq 0.01$ ; \*\*\*:  $p \leq 0.001$ . CI: confidence interval; d: effect size - Cohen's d. SNP: single nucleotide polymorphism.