

SUPPLEMENTARY TABLES

Supplementary Table 1. The demographic characteristics and MMSE scores of study subjects.

Characteristic	MMSE score (Mean ± SD)	P
Gender		
Male	27.5±2.3	
Female	27.3±2.6	< 0.001
Education level ¹		
"1" for no formal education,	1: 27.1±3.1	
"2" for homeschooling,	2: 27.4±2.4	
"3" for elementary school,	3: 25.2±3.2	
"4" for middle school,	4: 26.7±2.5	< 0.001
"5" for high school,	5: 27.7±1.9	
"6" for college, and	6: 28.2±1.7	
"7" for graduate school	7: 28.5±1.6	
Physical activity		
Yes	27.5±2.4	
No	27.2±2.6	< 0.001
Smoking		
Yes	27.2±2.6	
No	27.4±2.5	0.004
Alcohol drinking		
Yes	27.4±2.3	
No	27.4±2.5	0.279
Chronic conditions		
Yes	27.4±2.4	
No	27.0±2.8	< 0.001

MMSE, Mini-Mental State Examination; SD, standard deviation.

Data are presented as mean ± standard deviation.

¹Education level is defined as the following seven levels: no formal education, homeschooling, elementary school, middle school, high school, college, and graduate school.

Supplementary Table 2. Linear regression models of associations between the MMSE scores and 46 SNPs in the NCAM1-TTC12-ANKK1-DRD2 locus, which have an evidence of association (P < 0.05).

CHR	Gene	SNP	A1	A2	Region	MAF	Dominant model			Recessive model		
							BETA	SE	P	BETA	SE	P
11	<i>NCAM1-LOC105369498</i>	rs143926817	G	A	intergenic	0.016	-0.27	0.09	1.94E-03	-0.27	1.08	0.806
11	<i>NCAM1</i>	rs10789931	T	C	intron	0.468	-0.06	0.03	0.079	-0.14	0.04	1.62E-04
11	<i>NCAM1</i>	rs11214442	G	A	intron	0.304	0.13	0.03	2.06E-05	0.14	0.05	0.011
11	<i>NCAM1</i>	rs61905366	G	A	intron	0.312	0.12	0.03	1.20E-04	0.14	0.05	6.61E-03
11	<i>NCAM1</i>	rs10891485	G	A	intron	0.303	0.13	0.03	2.54E-05	0.13	0.05	0.015
11	<i>NCAM1</i>	rs1940725	T	C	intron	0.182	0.12	0.03	2.45E-04	0.04	0.09	0.627
11	<i>NCAM1</i>	rs7128707	T	A	intron	0.443	0.11	0.03	5.81E-04	0.09	0.04	0.020
11	<i>NCAM1</i>	rs1940717	A	G	intron	0.121	0.07	0.04	0.070	0.26	0.13	0.045
11	<i>NCAM1</i>	rs4424705	T	A	intron	0.443	0.12	0.03	4.74E-04	0.09	0.04	0.020
11	<i>NCAM1</i>	rs12363408	A	C	intron	0.444	0.12	0.03	3.68E-04	0.09	0.04	0.021
11	<i>NCAM1</i>	rs1884	G	C	intron	0.443	0.12	0.03	4.55E-04	0.09	0.04	0.019
11	<i>NCAM1</i>	rs61902793	G	A	intron	0.444	0.12	0.03	3.92E-04	0.09	0.04	0.018
11	<i>NCAM1</i>	rs11214468	C	T	intron	0.444	0.12	0.03	3.12E-04	0.09	0.04	0.015
11	<i>NCAM1</i>	rs12365502	C	A	intron	0.444	0.12	0.03	3.76E-04	0.09	0.04	0.015
11	<i>NCAM1</i>	rs1940735	T	G	intron	0.440	0.12	0.03	2.00E-04	0.09	0.04	0.024
11	<i>NCAM1</i>	rs10750023	A	G	intron	0.080	0.11	0.04	9.25E-03	-0.04	0.20	0.858
11	<i>NCAM1</i>	rs1836799	C	T	intron	0.374	0.07	0.03	0.023	0.00	0.04	0.911
11	<i>NCAM1</i>	rs1245089	C	G	intron	0.081	0.11	0.04	8.62E-03	-0.05	0.19	0.790
11	<i>NCAM1</i>	rs1816537	C	A	intron	0.091	0.10	0.04	0.015	0.25	0.18	0.169
11	<i>NCAM1</i>	rs12788208	A	G	intron	0.091	0.09	0.04	0.021	0.25	0.18	0.155

11	<i>NCAMI</i>	rs56268408	T	C	intron	0.099	0.09	0.04	0.024	-0.13	0.16	0.415
11	<i>NCAMI</i>	rs189464800	G	A	intron	0.023	-0.15	0.07	0.042	0.12	0.73	0.868
11	<i>NCAMI</i>	rs10891508	A	G	intron	0.376	-0.06	0.03	0.064	-0.13	0.04	4.10E-03
11	<i>NCAMI</i>	rs72995965	T	A	intron	0.100	0.09	0.04	0.022	-0.11	0.16	0.513
11	<i>NCAMI</i>	rs873803	A	G	intron	0.139	-0.08	0.04	0.031	-0.26	0.11	0.023
11	<i>NCAMI</i>	rs10891509	G	T	intron	0.376	-0.06	0.03	0.063	-0.13	0.04	4.15E-03
11	<i>NCAMI</i>	rs595231	C	T	intron	0.391	-0.08	0.03	0.016	-0.05	0.04	0.279
11	<i>NCAMI</i>	rs144186126	T	G	intron	0.022	-0.07	0.08	0.333	-1.70	0.81	0.036
11	<i>NCAMI</i>	rs146103467	T	C	intron	0.011	-0.08	0.10	0.446	-4.76	1.71	5.38E-03
11	<i>NCAMI</i>	rs146627958	G	A	intron	0.022	-0.19	0.08	0.013	1.60	0.73	0.029
11	<i>NCAMI-TTC12</i>	rs185398311	T	C	intergenic	0.012	-0.21	0.10	0.045	-1.83	2.43	0.450
11	<i>NCAMI-TTC12</i>	rs56159811	T	C	intergenic	0.056	-0.05	0.05	0.314	-0.56	0.27	0.034
11	<i>TTC12</i>	rs186496476	A	G	2KB upstream	0.016	-0.18	0.09	0.043	1.90	1.40	0.173
11	<i>TTC12</i>	rs138333675	A	G	missense	0.029	-0.01	0.07	0.879	-2.54	0.51	5.39E-07
11	<i>TTC12</i>	rs184198959	T	C	missense	0.013	-0.21	0.10	0.034	-1.83	2.42	0.450
11	<i>DRD2</i>	rs4936270	T	C	intron	0.137	0.00	0.04	0.894	-0.32	0.11	4.78E-03
11	<i>DRD2</i>	rs4274224	G	A	intron	0.167	-0.01	0.03	0.750	-0.18	0.09	0.049
11	<i>DRD2</i>	rs4245148	T	C	intron	0.137	0.00	0.04	0.993	-0.31	0.11	5.66E-03
11	<i>DRD2</i>	rs4460839	C	T	intron	0.137	0.00	0.04	0.966	-0.29	0.11	9.32E-03
11	<i>DRD2</i>	rs4581480	C	T	intron	0.137	0.00	0.04	0.992	-0.29	0.11	9.08E-03
11	<i>DRD2</i>	rs7122454	C	G	intron	0.393	0.08	0.03	9.57E-03	0.03	0.04	0.553
11	<i>DRD2</i>	rs10891550	C	G	intron	0.393	0.08	0.03	9.11E-03	0.03	0.04	0.476
11	<i>DRD2</i>	rs11214611	G	A	intron	0.393	0.08	0.03	9.95E-03	0.03	0.04	0.468
11	<i>DRD2</i>	rs4350392	A	C	intron	0.377	0.07	0.03	0.030	0.02	0.04	0.639
11	<i>DRD2</i>	rs4245149	A	G	intron	0.377	0.07	0.03	0.032	0.02	0.04	0.646
11	<i>DRD2</i>	rs10789944	A	C	intron	0.377	0.07	0.03	0.036	0.02	0.04	0.668

A1, minor allele; A2, major allele; CHR, chromosome; BETA, Beta coefficients; MAF, minor allele frequency; MMSE, Mini-Mental State Examination; SE, standard error.

Analysis was obtained after adjustment for covariates including age, gender, education, physical activity, smoking, alcohol drinking, and chronic conditions.

P values of < 0.05 are shown in bold.

Supplementary Table 3. Linear regression models of associations between the MMSE scores and 44 SNPs in the DRD3-LOC107986115-ZNF80-TIGIT-MIR568-ZBTB20 locus, which have an evidence of association (P < 0.05).

CHR	Gene	SNP	A1	A2	Region	MAF	Dominant model			Recessive model		
							BETA	SE	P	BETA	SE	P
3	<i>DRD3</i>	rs144005981	A	G	intron	0.016	0.02	0.09	0.782	-2.74	1.21	0.024
3	<i>DRD3</i>	rs2971569	G	T	intron	0.386	0.08	0.03	0.017	0.05	0.04	0.262
3	<i>DRD3</i>	rs7638876	C	T	intron	0.341	0.07	0.03	0.035	0.07	0.05	0.161
3	<i>DRD3</i>	rs62267154	C	G	intron	0.252	0.08	0.03	0.011	0.07	0.06	0.238
3	<i>DRD3</i>	rs36212178	G	A	intron	0.238	0.08	0.03	8.61E-03	0.08	0.07	0.218
3	<i>DRD3</i>	rs1587756	C	T	intron	0.252	0.08	0.03	0.011	0.07	0.06	0.238
3	<i>DRD3</i>	rs1587757	G	A	intron	0.238	0.08	0.03	8.46E-03	0.08	0.07	0.210
3	<i>DRD3</i>	rs1587758	G	A	intron	0.252	0.08	0.03	0.010	0.07	0.06	0.230
3	<i>DRD3</i>	rs36212515	C	T	intron	0.251	0.08	0.03	8.44E-03	0.08	0.06	0.199
3	<i>DRD3</i>	rs12633630	C	A	intron	0.261	0.08	0.03	8.65E-03	0.06	0.06	0.304
3	<i>DRD3</i>	rs56362817	A	C	intron	0.252	0.08	0.03	0.010	0.08	0.06	0.228
3	<i>DRD3</i>	rs55953427	A	G	intron	0.252	0.08	0.03	0.011	0.08	0.06	0.223
3	<i>DRD3</i>	rs6804925	T	A	intron	0.252	0.08	0.03	0.011	0.08	0.06	0.223
3	<i>DRD3</i>	rs1394016	A	G	intron	0.356	0.06	0.03	0.043	0.08	0.05	0.066
3	<i>DRD3</i>	rs17605608	A	G	intron	0.261	0.08	0.03	9.83E-03	0.06	0.06	0.299
3	<i>DRD3</i>	rs57504078	G	A	intron	0.238	0.08	0.03	7.79E-03	0.08	0.07	0.209
3	<i>DRD3</i>	rs62268962	T	A	intron	0.262	0.08	0.03	9.38E-03	0.05	0.06	0.391
3	<i>DRD3-LOC107986115</i>	rs7649438	G	C	intergenic	0.262	0.08	0.03	0.011	0.06	0.06	0.343
3	<i>ZNF80-TIGIT</i>	rs78899496	A	G	intergenic	0.026	0.16	0.07	0.023	-0.26	0.52	0.609
3	<i>ZNF80-TIGIT</i>	rs965970177	T	G	intergenic	0.010	-0.09	0.11	0.431	-7.93	2.42	1.07E-03
3	<i>ZBTB20</i>	rs75995250	A	G	intron	0.086	0.09	0.04	0.042	0.40	0.18	0.022
3	<i>ZBTB20</i>	rs1219190601	T	G	intron	0.012	-0.38	0.10	2.07E-04	2.05	1.71	0.232
3	<i>ZBTB20</i>	rs76546326	C	T	intron	0.013	0.07	0.10	0.447	2.86	1.40	0.040

3	ZBTB20	rs150020524	A	G	intron	0.024	-0.01	0.07	0.851	-2.13	0.70	2.37E-03
3	ZBTB20	rs77867745	A	G	intron	0.038	-0.04	0.06	0.536	0.94	0.47	0.043
3	ZBTB20	rs116180437	T	A	intron	0.048	-0.05	0.05	0.345	-0.99	0.34	3.29E-03
3	ZBTB20	rs145272406	C	T	intron	0.015	-0.12	0.09	0.177	-4.27	0.99	1.59E-05
3	ZBTB20	rs114295131	A	C	intron	0.015	-0.12	0.09	0.174	-4.27	0.99	1.59E-05
3	ZBTB20	rs77949732	T	A	intron	0.015	-0.11	0.09	0.222	-4.27	0.99	1.59E-05
3	ZBTB20	rs146425021	A	G	intron	0.029	-0.04	0.07	0.542	-1.33	0.65	0.039
3	ZBTB20	rs201716537	C	T	intron	0.010	0.08	0.11	0.474	-3.76	1.40	7.28E-03
3	ZBTB20	rs145607299	T	C	intron	0.011	0.07	0.10	0.497	-3.51	1.21	3.82E-03
3	ZBTB20	rs931723	C	G	intron	0.204	0.08	0.03	0.012	-0.06	0.08	0.428
3	ZBTB20	rs4146802	A	C	intron	0.022	0.02	0.07	0.797	-1.57	0.63	0.012
3	ZBTB20	rs138639994	T	G	intron	0.016	-0.10	0.09	0.261	-1.46	0.73	0.047
3	ZBTB20	rs9827199	T	C	intron	0.359	0.05	0.03	0.107	0.11	0.05	0.018
3	ZBTB20	rs143323356	A	G	intron	0.046	-0.11	0.05	0.034	0.15	0.36	0.681
3	ZBTB20	rs10222496	A	C	intron	0.404	0.02	0.03	0.525	0.11	0.04	7.86E-03
3	ZBTB20	rs73228389	G	C	intron	0.408	0.02	0.03	0.516	0.12	0.04	4.92E-03
3	ZBTB20	rs144357867	G	A	intron	0.016	-0.10	0.09	0.242	-2.65	1.08	0.014
3	ZBTB20	rs7643617	A	C	intron	0.406	0.03	0.03	0.430	0.11	0.04	9.38E-03
3	ZBTB20	rs1473580	C	G	intron	0.413	0.03	0.03	0.406	0.09	0.04	0.034
3	ZBTB20	rs9851136	A	C	intron	0.418	0.02	0.03	0.483	0.09	0.04	0.021
3	ZBTB20	rs73230612	C	T	intron	0.413	0.02	0.03	0.446	0.09	0.04	0.023

A1, minor allele; A2, major allele; CHR, chromosome; BETA, Beta coefficients; MAF, minor allele frequency; MMSE, Mini-Mental State Examination; SE, standard error.

Analysis was obtained after adjustment for covariates including age, gender, education, physical activity, smoking, alcohol drinking, and chronic conditions.

P values of < 0.05 are shown in bold.

Supplementary Table 4. Linear regression models of associations between the MMSE scores and 11 SNPs in the DRD1 locus, which have an evidence of association (P < 0.05).

CHR	Gene	SNP	A1	A2	Region	MAF	Dominant model			Recessive model		
							BETA	SE	P	BETA	SE	P
5	ARL2BPP6-DRD1	rs76373994	G	A	Intergenic	0.030	-0.16	0.07	0.017	-0.66	0.45	0.143
5	ARL2BPP6-DRD1	rs2253357	C	T	Intergenic	0.133	-0.07	0.04	0.041	-0.03	0.12	0.820
5	ARL2BPP6-DRD1	rs188263122	C	T	Intergenic	0.016	0.06	0.09	0.503	-2.84	0.92	1.92E-03
5	DRD1	rs686	G	A	3' UTR	0.152	-0.07	0.03	0.040	-0.07	0.10	0.522
5	DRD1	rs4532	C	T	3' UTR	0.152	-0.07	0.03	0.039	-0.07	0.10	0.474
5	DRD1	rs265981	A	G	3' UTR	0.145	-0.08	0.03	0.020	-0.08	0.11	0.459
5	DRD1	rs267410	A	C	2KB upstream	0.478	0.03	0.03	0.372	0.09	0.04	0.018
5	DRD1-SFXN1	rs183458028	A	C	Intergenic	0.016	0.09	0.09	0.295	-1.62	0.70	0.020
5	DRD1-SFXN1	rs267416	C	T	Intergenic	0.447	0.03	0.03	0.437	0.09	0.04	0.023
5	DRD1-SFXN1	rs2644644	G	C	Intergenic	0.451	0.01	0.03	0.855	0.08	0.04	0.042
5	DRD1-SFXN1	rs147030807	G	A	Intergenic	0.011	-0.27	0.11	9.65E-03	-0.88	1.71	0.609

A1, minor allele; A2, major allele; CHR, chromosome; BETA, Beta coefficients; MAF, minor allele frequency; MMSE, Mini-Mental State Examination; SE, standard error.

Analysis was obtained after adjustment for covariates including age, gender, education, physical activity, smoking, alcohol drinking, and chronic conditions.

P values of < 0.05 are shown in bold.

Supplementary Table 5. Linear regression models of associations between the MMSE scores and 3 SNPs in the *DRD4* locus, which have an evidence of association ($P < 0.05$).

CHR	Gene	SNP	A1	A2	Region	MAF	Dominant model			Recessive model		
							BETA	SE	P	BETA	SE	P
11	<i>DRD4-DEAF1</i>	rs183494193	T	A	intergenic	0.012	-0.17	0.10	0.088	-2.36	1.09	0.030
11	<i>DRD4-DEAF1</i>	rs111419860	C	T	intergenic	0.394	-0.02	0.03	0.512	0.09	0.04	0.028
11	<i>DRD4-DEAF1</i>	rs7109899	G	A	intergenic	0.391	-0.02	0.03	0.612	0.12	0.04	6.68E-03

A1, minor allele; A2, major allele; CHR, chromosome; BETA, Beta coefficients; MAF, minor allele frequency; MMSE, Mini-Mental State Examination; SE, standard error.

Analysis was obtained after adjustment for covariates including age, gender, education, physical activity, smoking, alcohol drinking, and chronic conditions.

P values of < 0.05 are shown in bold.

Supplementary Table 6. Linear regression models of associations between the MMSE scores and 12 SNPs in the *DRD5-SLC2A9* locus, which have an evidence of association ($P < 0.05$).

CHR	Gene	SNP	A1	A2	Region	MAF	Dominant model			Recessive model		
							BETA	SE	P	BETA	SE	P
4	<i>SLC2A9</i>	rs28523967	G	A	intron	0.151	0.07	0.03	0.053	0.26	0.10	0.011
4	<i>SLC2A9</i>	rs2030287	T	G	intron	0.151	0.07	0.03	0.055	0.26	0.10	0.012
4	<i>SLC2A9</i>	rs16888725	C	T	non coding transcript	0.150	0.07	0.03	0.046	0.29	0.10	4.84E-03
4	<i>SLC2A9</i>	rs118072948	A	G	intron	0.025	-0.14	0.07	0.047	0.46	0.54	0.395
4	<i>SLC2A9</i>	rs36075927	A	C	intron	0.148	0.08	0.03	0.028	0.32	0.11	3.08E-03
4	<i>SLC2A9</i>	rs11946054	C	T	intron	0.083	0.05	0.04	0.193	0.39	0.19	0.046
4	<i>SLC2A9</i>	rs10004947	C	T	intron	0.083	0.05	0.04	0.189	0.39	0.19	0.046
4	<i>SLC2A9</i>	rs145231664	T	C	intron	0.020	0.16	0.08	0.047	0.89	0.73	0.225
4	<i>SLC2A9</i>	rs56014085	A	G	intron	0.083	0.06	0.04	0.186	0.39	0.19	0.046
4	<i>SLC2A9</i>	rs56038393	A	G	intron	0.065	0.00	0.05	0.937	-0.49	0.22	0.028
4	<i>SLC2A9</i>	rs142956580	T	C	intron	0.015	-0.19	0.09	0.034	1.89	1.21	0.121
4	<i>SLC2A9</i>	rs7681699	A	G	intron	0.012	-0.29	0.10	3.75E-03	-3.25	2.42	0.179

A1, minor allele; A2, major allele; CHR, chromosome; BETA, Beta coefficients; MAF, minor allele frequency; MMSE, Mini-Mental State Examination; SE.

Analysis was obtained after adjustment for covariates including age, gender, education, physical activity, smoking, alcohol drinking, and chronic conditions.

P values of < 0.05 are shown in bold.

Supplementary Table 7. Linkage disequilibrium (LD) among 3 SNPs including rs11214442 in *NCAM1*, rs10891485 in *NCAM1*, and rs138333675 in *TTC12*.

Chr	Gene 1	SNP 1	Gene 2	SNP 2	r2
12	<i>NCAM1</i>	rs11214442	<i>NCAM1</i>	rs10891485	0.997
12	<i>NCAM1</i>	rs11214442	<i>TTC12</i>	rs138333675	0.003
12	<i>NCAM1</i>	rs10891485	<i>TTC12</i>	rs138333675	0.003

The measure of LD is based on the squared allelic correlation r^2 .

Supplementary Table 8. Linkage disequilibrium (LD) among 3 SNPs including rs145272406, rs114295131, and rs77949732 in *ZBTB20*.

Chr	Gene 1	SNP 1	Gene 2	SNP 2	r2
12	<i>ZBTB20</i>	rs145272406	<i>ZBTB20</i>	rs114295131	1.000
12	<i>ZBTB20</i>	rs145272406	<i>ZBTB20</i>	rs77949732	1.000
12	<i>ZBTB20</i>	rs114295131	<i>ZBTB20</i>	rs77949732	1.000

The measure of LD is based on the squared allelic correlation r^2 .