

**SUPPLEMENTAL E-TABLES**

Supplemental: e-Table 1: Secondary Neurocognitive Outcomes Sub Scores

Secondary Neurocognitive Outcomes	Group Difference at Age 6				Group Difference at Age 8				Group Difference at Age 10				Group Difference at Age 12			
	Est.	95% CI	ES*	p-val	Est.	95% CI	ES*	p-val	Est.	95% CI	ES*	p-val	Est.	95% CI	ES*	p-val
<b>Vocabulary</b>	-0.66	(-1.34,0.01)	-0.24	0.053	-0.71	(-1.26,-0.16)	-0.31	0.011	-0.76	(-1.27,-0.25)	-0.36	0.003	-0.81	(-1.37,-0.24)	-0.34	0.006
<b>Matrix Reasoning</b>	-0.84	(-1.59,-0.09)	-0.27	0.029	-0.65	(-1.24,-0.06)	-0.27	0.030	-0.47	(-0.94,0.01)	-0.24	0.054	-0.28	(-0.74,0.17)	-0.15	0.220
<b>Similarity</b>	-0.63	(-1.24,-0.02)	-0.25	0.043	-0.60	(-1.12,-0.08)	-0.28	0.023	-0.57	(-1.06,-0.07)	-0.28	0.024	-0.54	(-1.08,0.01)	-0.24	0.056
<b>Block Design</b>	-0.47	(-1.24,0.31)	-0.14	0.240	-0.34	(-1.04,0.35)	-0.12	0.332	-0.22	(-0.90,0.46)	-0.08	0.524	-0.10	(-0.83,0.64)	-0.03	0.794
<b>CPT2 Commissions</b>	1.85	(-0.62,4.32)	0.18	0.142	1.84	(-0.19,3.87)	0.22	0.075	1.84	(-0.28,3.95)	0.21	0.088	1.83	(-0.84,4.50)	0.17	0.179
<b>CPT2 Detectability</b>	1.10	(-1.37,3.56)	0.11	0.383	1.05	(-0.94,3.04)	0.13	0.301	1.00	(-1.00,3.00)	0.12	0.327	0.95	(-1.54,3.45)	0.09	0.455
<b>Concept Formation</b>	-3.88	(-7.19,-0.56)	-0.28	0.022	-3.45	(-6.26,-0.64)	-0.30	0.016	-3.03	(-5.61,-0.45)	-0.28	0.021	-2.60	(-5.30,0.10)	-0.23	0.059

\* Effect sizes (Cohen's d) were approximately calculated as 2 times t-value divided by square root of sample size minus one, where t- values were calculated as point estimates of group difference from mixed effects modeling divided by their robust maximum likelihood standard errors.

Supplemental e-Table 2: Results of LME conditional on replacement status

Primary Outcomes	Group Difference at Age 6				Group Difference at Age 8				Group Difference at Age 10				Group Difference at Age 12			
	Estimate	95% CI	ES*	p-val	Estimate	95% CI	ES*	p-val	Estimate	95% CI	ES*	p-val	Estimate	95% CI	ES*	p-val
<i>Neurocognitive</i>																
FSIQ	<b>-4.18</b>	<b>(-7.41,-0.95)</b>	<b>-0.31</b>	<b>0.011</b>	<b>-3.86</b>	<b>(-6.73,-0.99)</b>	<b>-0.32</b>	<b>0.008</b>	<b>-3.53</b>	<b>(-6.28,-0.79)</b>	<b>-0.31</b>	<b>0.012</b>	<b>-3.21</b>	<b>(-6.10,-0.32)</b>	<b>-0.27</b>	<b>0.030</b>
PIQ	-3.43	(-7.30, 0.45)	-0.21	0.083	-2.73	(-6.05, 0.59)	-0.20	0.107	-2.04	(-5.05, 0.97)	-0.16	0.184	-1.35	(-4.36, 1.67)	-0.11	0.383
VIQ	<b>-3.82</b>	<b>(-7.09,-0.54)</b>	<b>-0.28</b>	<b>0.020</b>	<b>-3.89</b>	<b>(-6.75,-1.03)</b>	<b>-0.33</b>	<b>0.008</b>	<b>-3.96</b>	<b>(-6.72,-1.20)</b>	<b>-0.35</b>	<b>0.005</b>	<b>-4.04</b>	<b>(-7.04,-1.03)</b>	<b>-0.32</b>	<b>0.009</b>
<i>Structural MRI</i>																
Total brain volume	-15524	(-35588, 4541)	-0.18	0.129	<b>-21322</b>	<b>(-41790,-854)</b>	<b>-0.25</b>	<b>0.041</b>	<b>-25727</b>	<b>(-46808,-4646)</b>	<b>-0.29</b>	<b>0.017</b>	<b>-28738</b>	<b>(-50224,-7252)</b>	<b>-0.32</b>	<b>0.009</b>
Gray matter	-8637	(-20596, 3323)	-0.17	0.157	<b>-11987</b>	<b>(-23905, -70)</b>	<b>-0.24</b>	<b>0.049</b>	<b>-14542</b>	<b>(-26798,-2286)</b>	<b>-0.28</b>	<b>0.020</b>	<b>-16300</b>	<b>(-28744,-3856)</b>	<b>-0.31</b>	<b>0.010</b>
White matter	-6263	(-15766, 3241)	-0.16	0.196	-8381	(-18183, 1421)	-0.20	0.094	<b>-10368</b>	<b>(-20609,-128)</b>	<b>-0.24</b>	<b>0.047</b>	<b>-12225</b>	<b>(-22786,-1664)</b>	<b>-0.28</b>	<b>0.023</b>

\* Effect sizes (Cohen's d) were approximately calculated as 2 times t-value divided by square root of sample size minus one, where t-values were calculated as point estimates of group difference from mixed effects modeling divided by their robust maximum likelihood standard errors.

Supplemental e-Table 3 Correlations between Brain and Neurocognitive Scores in T1D and Control Groups (longitudinal mixed effect modeling)				
Age	6 years	8 years	10 years	12 years
<b>Total Brain Volume</b>				
Diabetes				
FSIQ	0.245 (0.086) <b>P=0.004</b>	0.272 (0.074) <b>P=0.000</b>	0.283 (0.070) <b>P=0.000</b>	0.277 (0.075) <b>P=0.000</b>
PIQ	0.339 (0.085) <b>P=0.000</b>	0.347 (0.075) <b>P=0.000</b>	0.347 (0.072) <b>P=0.000</b>	0.336 (0.078) <b>P=0.000</b>
VIQ	0.043 (0.087) <b>P=0.004</b>	0.090 (0.078) P=0.246	0.133 (0.076) 0.081	0.166 (0.085) P=0.052
Control				
FSIQ	0.407 (0.125) <b>P=-0.001</b>	0.397 (0.117) <b>P=0.001</b>	0.374 (0.118) <b>P=0.002</b>	0.345 (0.123) <b>P=0.005</b>
PIQ	0.282 (0.148) P=0.057	0.311 (0.137) P=0.024	0.325 (0.128) <b>P=0.011</b>	0.323 (0.120) <b>P=0.007</b>
VIQ	0.418 (0.114) <b>P=0.000</b>	0.377 (0.104) <b>P=0.000</b>	0.332 (0.116) <b>P=0.004</b>	0.290 (0.136) <b>P=0.033</b>
<b>Gray Matter Volume</b>				
Diabetes				
FSIQ	0.225 (0.081) <b>P=0.006</b>	0.266 (0.070) <b>P=0.000</b>	0.286 (0.068) <b>P=0.000</b>	0.285 (0.076) <b>P=0.000</b>
PIQ	0.314 (0.086) <b>P=0.000</b>	0.336 (0.073) <b>P=0.000</b>	0.348 (0.071) <b>P=0.000</b>	0.347 (0.078) <b>P=0.000</b>
VIQ	0.043 (0.083) P=0.605	0.094 (0.074) P=0.205	0.137 (0.075) P=0.069	0.166 (0.088) P=0.061
Control				
FSIQ	0.367 (0.125) <b>P=0.003</b>	0.375 (0.112) <b>P=0.001</b>	0.365 (0.112) <b>P=0.001</b>	0.032 (0.12) <b>P=0.004</b>
PIQ	0.247 (0.144) P=0.086	0.294 (0.132) <b>P=0.026</b>	0.321 (0.126) <b>P=0.011</b>	0.325 (0.121) <b>P=0.007</b>
VIQ	0.402 (0.129) <b>P=0.002</b>	0.368 (0.111) <b>P=0.001</b>	0.329 (0.114) <b>P=0.004</b>	0.290 (0.130) <b>P=0.026</b>
<b>White Matter Volume</b>				

Diabetes				
FSIQ	0.249 (0.094) <b>P=0.008</b>	0.258 (0.083) <b>P=0.002</b>	0.257 (0.077) <b>P=0.001</b>	0.243 (0.078) P=-0.002
PIQ	0.342 (0.086) <b>P=0.008</b>	0.333 (0.079) <b>P=0.000</b>	0.319 (0.077) <b>P=0.000</b>	0.294 (0.082) <b>P=0.000</b>
VIQ	0.040 (0.096) P=0.667	0.078 (0.086) P=0.365	0.117 (0.082) P=0.154	0.149 (0.086) P=0.084
Control				
FSIQ	0.398 (0.136) <b>P=0.003</b>	0.372 (0.126) <b>P=0.003</b>	0.342 (0.124) <b>P=0.006</b>	0.309 (0.127) <b>P=0.015</b>
PIQ	0.282 (0.155) P=0.069	0.291 (0.140) <b>P=0.037</b>	0.293 (0.127) <b>P=0.021</b>	0.286 (0.118) <b>P=0.015</b>
VIQ	0.380 (0.114) <b>P=0.001</b>	0.338 (0.106) <b>P=0.001</b>	0.297 (0.122) <b>P=0.015</b>	0.256 (0.145) P=0.077