

Supplementary Table 2. Mean values for CBD, Δ^9 -THC eCBs and NAEs, per treatment group (pmol/mL), alongside individual and adjusted p-values representing changes in concentration (65 and 160min) from t=0

eCB/NAE	Serum levels of eCBs/NAEs (pmol/mL) ¹				2-way ANOVA (comparison to t=0)			
	Medication	Post drug administration			65-min		160-min	
		0min	65min	160min	p-value ²	p _{corr} ³	p-value ²	p _{corr} ³
AEA	Placebo	1.866 ± 0.200	1.559 ± 0.132	1.850 ± 0.154	0.0840	0.1765	0.9354	0.9822
	THC 10mg	2.554 ± 0.237	1.831 ± 0.130	2.182 ± 0.156	<u>0.0013</u>	<u>0.0014</u>	0.1670	0.0877
	THC 20mg	1.370 ± 0.098	1.031 ± 0.100	1.476 ± 0.113	0.0552	0.1160	0.5516	0.5791
	CBD 600mg	1.992 ± 0.198	1.703 ± 0.209	1.828 ± 0.175	0.1162	0.1534	0.1461	0.1534
	CBD 800mg	1.201 ± 0.149	1.530 ± 0.309	1.914 ± 0.302	0.0979	0.0514	<u>0.0029</u>	<u>0.0030</u>
	CBD 800mg + THC 20mg	1.169 ± 0.149	1.628 ± 0.268	2.480 ± 0.421	0.0624	<u>0.0328</u>	<u>0.0076</u>	<u>0.0080</u>
2-AG	Placebo	3.003 ± 0.461	3.046 ± 0.507	3.256 ± 0.832	0.8938	0.9385	0.6970	0.9385
	THC 10mg	3.448 ± 0.437	3.951 ± 0.758	4.746 ± 1.130	0.2396	0.2516	0.1154	0.2423
	THC 20mg	2.790 ± 0.326	2.919 ± 0.498	2.435 ± 0.366	0.8147	0.8554	0.4193	0.8554
	CBD 600mg	3.887 ± 0.683	3.305 ± 0.516	3.872 ± 0.824	<u>0.0331</u>	0.0694	0.9772	0.9999
	CBD 800mg	2.066 ± 0.338	2.717 ± 0.448	2.876 ± 0.664	0.1064	0.1773	0.1689	0.1773
	CBD 800mg + THC 20mg	2.633 ± 0.412	2.654 ± 0.401	3.020 ± 0.486	0.9421	0.9892	0.0483	0.1015
OEA	Placebo	9.322 ± 0.601	8.463 ± 0.669	10.088 ± 0.805	0.1496	0.2557	0.2435	0.2557
	THC 10mg	7.857 ± 1.002	6.296 ± 0.572	8.162 ± 0.773	0.2200	0.4621	0.6554	0.6882
	THC 20mg	7.857 ± 1.319	10.049 ± 0.783	9.758 ± 0.950	0.7295	0.9346	0.8901	0.9346
	CBD 600mg	6.894 ± 0.704	6.065 ± 0.567	6.714 ± 0.734	0.0995	0.2088	0.8536	0.8962
	CBD 800mg	6.692 ± 0.885	9.458 ± 1.480	9.577 ± 1.347	<u>0.0126</u>	<u>0.0132</u>	<u>0.0072</u>	<u>0.0132</u>
	CBD 800mg + THC 20mg	7.200 ± 0.870	12.225 ± 2.476	13.594 ± 2.549	0.0577	<u>0.0303</u>	<u>0.0223</u>	<u>0.0234</u>
PEA	Placebo	14.047 ± 1.703	15.586 ± 1.435	16.638 ± 1.871	0.2958	0.3106	0.1373	0.2883
	THC 10mg	20.320 ± 3.435	15.923 ± 1.892	21.190 ± 1.737	0.1152	0.2419	0.8200	0.8610
	THC 20mg	22.929 ± 2.065	23.294 ± 1.813	21.583 ± 2.013	0.8612	0.9042	0.5529	0.9042
	CBD 600mg	17.000 ± 1.713	19.622 ± 4.120	17.630 ± 1.740	0.5708	0.7796	0.7425	0.7796
	CBD 800mg	16.443 ± 2.063	22.907 ± 3.912	22.784 ± 3.456	<u>0.0455</u>	<u>0.0478</u>	<u>0.0193</u>	<u>0.0405</u>
	CBD 800mg + THC 20mg	16.294 ± 1.255	24.455 ± 3.912	28.775 ± 4.679	0.0991	0.0520	<u>0.0181</u>	<u>0.0190</u>

¹Concentrations were expressed as mean ± standard error (SEM); ²Unadjusted and ³adjusted p-values for multiple comparisons using the two-stage Benjamini, Krieger and Yekutieli FDR method (p_{corr}) are reported for changes in eCB and NAE expression (65 and 160min) from baseline (0min), with significant values highlighted (p < 0.05, underlined) and emphasised when adjusted for FDR (p_{corr} < 0.05, **bold italics underlined**). Abbreviations, eCB: endocannabinoid; NAE: N-acylethanolamine; HVs: healthy volunteers; AEA: anandamide; 2-AG: 2-arachidonoylglycerol; OEA: oleoylethanolamine; PEA: palmitoylethanolamine.