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| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
|  | Control | LPS-NaCl | | LPS-Etomidate | | | LPS-Ketamine | | | |
| Mean (SD) | Mean (SD) | p versus control | Mean (SD) | p vs control | p vs LPS-NaCl | Mean (SD) | p vs control | p vs LPS-NaCl | p vs LPS-Etomidate |
| CRH | 17.9 (5.8) | 31.7 (4.7) | 0.006 | 27.3 (5.9) | 0.03 | 0.437 | 29.1 (9.1) | 0.014 | 0.698 | 0.692 |
| POMC | 8.4 (0.9) | 20.5 (3.1) | <0.0001 | 23.7 (5) | <0.0001 | 0.089 | 19.4 (3.6) | <0.0001 | 0.74 | 0.047 |
| ACTH (pg/mL) | 9 (5.2) | 36 (15.2) | 0.118 | 89.9 (35.7) | <0.0001 | 0.001 | 2.5 (3.6) | 0.374 | 0.022 | <0.0001 |
| P450scc | 22.5 (9.5) | 32.3 (4.4) | 0.027 | 19 (6.7) | 0.399 | 0.005 | 18.8 (2.7) | 0.375 | 0.004 | 0.963 |
| 3β-HSD | 17.2 (2.3) | 15.1 (0.7) | 0.255 | 11.5 (4.7) | 0.005 | 0.054 | 14.2 (1.7) | 0.109 | 0.611 | 0.139 |
| 21-HS | 11.5 (2) | 16.8 (5.3) | 0.012 | 8.2 (0.8) | 0.056 | 0.0002 | 8.1 (1.2) | 0.034 | <0.0001 | 0.807 |
| 11β-HS | 5.7 (0.5) | 11 (4.1) | 0.0014 | 4.3 (0.5) | 0.317 | 0.0002 | 7.2 (1.1) | 0.31 | 0.012 | 0.054 |
| CORT-B (ng/mL) | 98.3 (62.6) | 642.4 (165) | <0.0001 | 412.7 (189) | 0.012 | 0.032 | 259.8 (161) | 0.1451 | 0.002 | 0.186 |

Multiple comparisons between groups 12 hours after LPS administration. Results are presented as mean ± SD and comparison was performed through a two-way ANOVA test with Bonferroni post-hoc analysis. n=5 per group. CRH: corticotropin releasing hormone, POMC: pro-opiomelanocortin, ACTH: adrenocorticotropin hormone, P450scc: P450 side chain cleavage, 3β-HSD : 3β hydroxysteroid deshydrogenase, 21-HS : 21 hydroxylase, 11β-HS : 11β hydroxylase, CORT-B: corticosterone.

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| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
|  | Control | LPS-NaCl | | LPS-Etomidate | | | LPS-Ketamine | | | |
|  | Mean (SD) | Mean (SD) | p versus control | Mean (SD) | p vs control | p vs LPS-NaCl | Mean (SD) | p vs control | p vs LPS-NaCl | p vs LPS-Etomidate |
| CRH | 1.6 (0.4) | 5.6 (2.6) | 0.057 | 5.7 (3.3) | 0.044 | 0.887 | 5 (4.3) | 0.087 | 0.824 | 0.717 |
| POMC | 6.1 (1.3) | 9.9 (3) | 0.013 | 10.9 (2.5) | 0.003 | 0.486 | 9.6 (1.3) | 0.021 | 0.819 | 0.359 |
| ACTH (pg/mL) | 5,8 (5.8) | 12,8 (4.9) | 0.257 | 10.6 (11.4) | 0.430 | 0.717 | 6.3 (3.2) | 0.891 | 0.208 | 0.358 |
| P450scc | 17.1 (6.4) | 4 (0.9) | 0.0002 | 6.7 (1.4) | 0.001 | 0.320 | 25.1 (5.2) | 0.008 | <0.0001 | <0.0001 |
| 3β-HSD | 31.0 (3.3) | 13.1 (3) | <0.0001 | 25.8 (2.8) | 0.014 | <0.0001 | 11.6 (3) | <0.0001 | 0.437 | <0.0001 |
| 21-HS | 15.3 (1.5) | 4.7 (1) | <0.0001 | 7.1 (0.9) | <0.0001 | 0.009 | 7.6 (1.6) | <0.0001 | 0.003 | 0.564 |
| 11β-HS | 9.7 (2.5) | 3.1 (0.6) | <0.0001 | 4.3 (0.8) | <0.0001 | 0.234 | 4.4 (1.2) | <0.0001 | 0.203 | 0.931 |
| CORT-B (ng/mL) | 72.9 (15) | 176.6 (76.8) | 0.327 | 218.1 (289) | 0 .176 | 0.691 | 159.8 (125) | 0.409 | 0.872 | 0.577 |

Multiple comparisons between groups 48 hours after LPS administration. Results are presented as mean +/- SD and comparison was performed through a two-way ANOVA test with Bonferroni post-hoc analysis. n=5 per group. CRH: corticotropin releasing hormone, POMC: pro-opiomelanocortin, ACTH: adrenocorticotropin hormone, P450scc: P450 side chain cleavage, 3β-HSD: 3β hydroxysteroid deshydrogenase, 21-HS: 21 hydroxylase, 11β-HS: 11β hydroxylase, CORT-B: corticosterone.

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| --- | --- | --- | --- | --- | --- | --- | --- | --- |
|  | CRH | POMC | ACTH | P450scc | 3β-HSD | 21-HS | 11β-HS | CORT-B |
| Interactions | 0.23 | 0.002 | <0.0001 | <0.0001 | <0.0001 | <0.0001 | <0.0001 | 0.03 |
| Groups | <0.0001 | <0.0001 | 0.0004 | <0.0001 | <0.0001 | 0.0008 | 0.0068 | 0.0006 |
| Time-points | 0.0056 | <0.0001 | <0.0001 | 0.0051 | <0.0001 | <0.0001 | 0.0013 | 0.0008 |

P values of the two-ANOVA test including the two time-points (12 and 48 hours) and the four groups (control, LPS-NaCl, LPS-etomidate, LPS-ketamine). The results for interaction between time-points and groups are presented as well as the comparisons between groups and between time-points.