Supplemental Digital Content Table 1. Level of Sedation, Presence of Nausea or Vomiting, Treatment of Nausea and Vomiting, and Presence of Itching, Hallucinations, and Dizziness

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| --- | --- | --- | --- | --- |
|  | Methadone Group | Methadone/Ketamine Group | Difference (99% CI) | P Value |
| Level of sedation |  |  |  |  |
|  At PACU admission | 2 (1 to 2)\* | 2 (1 to 2) | 0 (0 to 0) | 0.800 |
|  1 h after admission | 1 (1 to 2) | 1 (1 to 2) | 0 (0 to 0) | 0.804 |
|  Postoperative day 1 – AM | 0 (0 to 0) | 0 (0 to 0) | 0 (0 to 0) | 0.083 |
|  Postoperative day 1 - PM  | 0 (0 to 0) | 0 (0 to 0) | 0 (0 to 0) | 0.069 |
|  Postoperative day 2 – AM | 0 (0 to 0) | 0 (0 to 0)† | 0 (0 to 0) | 0.926 |
|  Postoperative day 2 – PM | 0 (0 to 0)\* | 0 (0 to 0)‡ | 0 (0 to 0) | 0.167 |
|  Postoperative day 3 – AM | 0 (0 to 0)§ | 0 (0 to 0)ǁ | 0 (0 to 0) | 0.788 |
|  Postoperative day 3 – PM | 0 (0 to 0)\*\* | 0 (0 to 0)†† | 0 (0 to 0) | > 0.999 |
| Nausea |  |  |  |  |
|  At PACU admission | 1/60 ( 2%) | 0/66 ( 0%) | 2% (- 8 to 13%) | 0.952‡‡ |
|  1 h after admission | 3/61 ( 5%) | 0/66 ( 0%) | 5% (- 5 to 18%) | 0.216‡‡ |
|  Postoperative day 1 – AM | 20/61 (33%) | 18/66 (27%) | 6% (-15 to 26%) | 0.628 |
|  Postoperative day 1 - PM  | 14/61 (23%) | 12/66 (18%) | 5% (-14 to 24%) | 0.656 |
|  Postoperative day 2 – AM | 12/61 (20%) | 13/64 (20%) | -1% (-19 to 18%) | > 0.999 |
|  Postoperative day 2 – PM | 10/60 (17%) | 7/61 (11%) | 5% (-12 to 23%) | 0.576 |
|  Postoperative day 3 – AM | 5/58 ( 9%) | 2/56 ( 4%) | 5% (- 9 to 20%) | 0.468‡‡ |
|  Postoperative day 3 – PM | 2/49 ( 4%) | 3/33 ( 9%) | -5% (-26 to 11%) | 0.635‡‡ |
| Vomiting |  |  |  |  |
|  At PACU admission | 0/60 ( 0%) | 0/66 ( 0%) | - | - |
|  1 h after admission | 1/61 ( 2%) | 0/66 ( 0%) | 2% (- 8 to 13%) | 0.961‡‡ |
|  Postoperative day 1 – AM | 7/61 (12%) | 4/66 ( 6%) | 5% (- 9 to 21%) | 0.442 |
|  Postoperative day 1 - PM  | 3/61 ( 5%) | 4/66 ( 6%) | -1% (-14 to 12%) | > 0.999‡‡ |
|  Postoperative day 2 – AM | 4/61 ( 7%) | 2/64 ( 3%) | 3% (- 9 to 17%) | 0.635‡‡ |
|  Postoperative day 2 – PM | 2/60 ( 3%) | 0/61 ( 0%) | 3% (- 7 to 15%) | 0.488‡‡ |
|  Postoperative day 3 – AM | 1/58 ( 2%) | 0/56 ( 0%) | 2% (- 9 to 13%) | > 0.999‡‡ |
|  Postoperative day 3 - PM | 0/49 ( 0%) | 0/33 ( 0%) | - | - |
| Medications to treat nausea and vomiting |  |  |  |
|  At PACU admission | 0/60 ( 0%) | 0/66 ( 0%) | - | - |
|  1 h after admission | 2/61 ( 3%) | 0/66 ( 0%) | 3% (- 6 to 15%) | 0.457‡‡ |
|  Postoperative day 1 - AM | 16/61 (26%) | 15/66 (23%) | 4% (-16 to 23%) | 0.801 |
|  Postoperative day 1 - PM  |  7/61 (12%) | 7/66 (11%) | 1% (-15 to 17%) | > 0.999 |
|  Postoperative day 2 - AM |  7/61 (12%) |  3/64 ( 5%) | 7% (- 7 to 22%) | 0.285 |
|  Postoperative day 2 - PM |  3/60 ( 5%) | 3/60 ( 5%) |  0% (-14 to 14%) | > 0.999‡‡ |
|  Postoperative day 3 - AM | 1/58 ( 2%) | 2/56 ( 4%) | -2% (-15 to 10%) | 0.974‡‡ |
|  Postoperative day 3 - PM | 1/49 ( 2%) | 2/33 ( 6%) | -4% (-24 to 10%) | 0.708‡‡ |
| Itching |  |  |  |  |
|  At PACU admission | 1/60 ( 2%) | 1/66 ( 2%) | 0% (-10 to 12%) | > 0.999‡‡ |
|  1 h after admission | 3/61 ( 5%) | 2/66 ( 3%) | 2% (-10 to 15%) | 0.925‡‡ |
|  Postoperative day 1 - AM | 14/61 (23%) | 15/66 (23%) | 0% (-19 to 20%) | > 0.999 |
|  Postoperative day 1 - PM  | 12/61 (20%) | 14/66 (21%) | - 2% (-20 to 18%) | > 0.999 |
|  Postoperative day 2 - AM | 12/61 (20%) | 21/64 (33%) | -13% (-33 to 7%) | 0.144 |
|  Postoperative day 2 - PM |  9/60 (15%) | 14/61 (23%) |  - 8% (-27 to 11%) | 0.377 |
|  Postoperative day 3 - AM | 6/58 (10%) | 11/56 (20%) | - 9% (-28 to 9%) | 0.258 |
|  Postoperative day 3 - PM | 4/49 ( 8%) |  4/33 (12%) | - 4% (-27 to 14%) | 0.818‡‡ |
| Hallucinations |  |  |  |  |
|  At PACU admission | 0/61 ( 0%) | 2/62 ( 3%) | - 3% (-15 to 7%) | 0.504‡‡ |
|  1 h after admission | 1/60 ( 2%) | 2/66 ( 3%) | - 1% (-13 to 10%) | > 0.999‡‡ |
|  Postoperative day 1 - AM | 3/61 ( 5%) | 4/66 ( 6%) | - 1% (-14 to 12%) | > 0.999‡‡ |
|  Postoperative day 1 - PM  | 2/61 ( 3%) | 3/66 ( 5%) | - 1% (-14 to 11%) | > 0.999‡‡ |
|  Postoperative day 2 - AM | 2/61 ( 3%) |  3/64 ( 5%) | - 1% (-14 to 11%) | > 0.999‡‡ |
|  Postoperative day 2 - PM |  0/60 ( 0%) |  1/63 ( 2%) |  - 2% (-12 to 9%) | > 0.999‡‡ |
|  Postoperative day 3 - AM | 0/58 ( 0%) | 1/56 ( 2%) | - 2% (-14 to 9%) | 0.983‡‡ |
|  Postoperative day 3 - PM | 0/49 ( 0%) |  1/33 ( 3%) | - 3% (-22 to 9%) | 0.805‡‡ |
| Dizziness |  |  |  |  |
|  At PACU admission | 1/60 ( 2%) | 0/62 ( 0%) | 2% (- 8 to 13%) | 0.984‡‡ |
|  1 h after admission | 0/60 ( 0%) | 1/66 ( 2%) | - 2% (-12 to 9%) | > 0.999‡‡ |
|  Postoperative day 1 - AM | 9/61 (15%) | 14/65 (22%) | - 7% (-25 to 12%) | 0.451 |
|  Postoperative day 1 - PM  | 13/61 (21%) | 24/66 (36%) | -15% (-35 to 6%) | 0.095 |
|  Postoperative day 2 - AM | 8/61 (13%) |  13/64 (20%) | - 7% (-25 to 11%) | 0.403 |
|  Postoperative day 2 - PM | 11/60 (18%) |  11/63 (17%) |  1% (-18 to 20%) | > 0.999 |
|  Postoperative day 3 - AM | 5/58 ( 9%) | 7/56 (13%) | - 4% (-21 to 12%) | 0.712 |
|  Postoperative day 3 - PM | 2/49 ( 4%) |  6/33 (18%) | -14% (-37 to 4%) | 0.086‡‡ |

Data are reported as median (interquartile range) or number of patients (%). Data reported as median (interquartile range) were compared between groups at the various times using the Mann-Whitney U test and data reported as number of patients (%) were compared between groups at the various times using the Pearson chi-square test or, when at least one of the cells of the contingency table had an expected n < 5, the Fisher exact probability test. No within-group (i.e., across time) comparisons have been made. Level of sedation on a 0 to 3 scale: 0 = fully awake to 3 = severely sedated. N = 61 in the methadone group and n = 66 in the methadone/ketamine group, except where indicated.

\*n = 60. †n = 64. ‡n = 63. §n = 58. ǁn = 56. \*\*n = 49. ††n = 33.

‡‡Friedman exact probability test

PACU = postanesthesia care unit