\*\*Table 1. Cohort summary\*\*

|  | **Main Summary** | | | | | **MEDPRN** | | |
| --- | --- | --- | --- | --- | --- | --- | --- | --- |
| **Variable** | | **N** | **0, N = 5,6301** | **1, N = 621** | **p-value2** | **0** | **1** | **(Missing)** |
| **Age** | | 5,692 | 55 (45, 64) | 56 (48, 67) | 0.105 |  |  |  |
| **Sex** | | 5,692 |  |  | 0.339 |  |  |  |
| Male | |  | 2,613 (46%) | 25 (40%) |  |  |  |  |
| Female | |  | 3,017 (54%) | 37 (60%) |  |  |  |  |
| **BMI** | | 5,685 | 27.3 (24.2, 31.0) | 30.5 (24.3, 35.4) | 0.004 |  |  |  |
| (Missing) | |  | 7 | 0 |  |  |  |  |
| **BMD** | | 4,685 | 0.97 (0.87, 1.07) | 0.93 (0.83, 1.03) | 0.103 |  |  |  |
| (Missing) | |  | 995 | 12 |  |  |  |  |
| **MEDPRN** | | 58 | 0 (NA%) | 37 (64%) | >0.999 |  |  |  |
| (Missing) | |  | 5,630 | 4 |  |  |  |  |
| **MEDPER** | | 56 |  |  | >0.999 |  |  |  |
| 1 | |  | 0 (NA%) | 42 (75%) |  | 21 | 17 | 4 |
| 2 | |  | 0 (NA%) | 7 (13%) |  | 0 | 7 | 0 |
| 3 | |  | 0 (NA%) | 7 (13%) |  | 0 | 7 | 0 |
| (Missing) | |  | 5,630 | 6 |  |  |  |  |
| (Missing) | |  |  |  |  | 0 | 6 | 5,630 |
| **CHEM\_GP1** | | 62 |  |  | >0.999 |  |  |  |
| Diphenylpropylamine derivatives | |  | 0 (NA%) | 11 (18%) |  | 2 | 8 | 1 |
| Natural opium alkaloids | |  | 0 (NA%) | 28 (45%) |  | 7 | 19 | 2 |
| Other opioids | |  | 0 (NA%) | 21 (34%) |  | 12 | 9 | 0 |
| Phenylpiperidine derivatives | |  | 0 (NA%) | 2 (3.2%) |  | 0 | 1 | 1 |
| (Missing) | |  | 5,630 | 0 |  |  |  |  |
| (Missing) | |  |  |  |  | 0 | 0 | 5,630 |
| **CHEM\_NM1** | | 62 |  |  | >0.999 |  |  |  |
| DEXTROPROPOXYPHENE | |  | 0 (NA%) | 11 (18%) |  | 2 | 8 | 1 |
| HYDROMORPHONE | |  | 0 (NA%) | 1 (1.6%) |  | 0 | 1 | 0 |
| MORPHINE | |  | 0 (NA%) | 3 (4.8%) |  | 1 | 1 | 1 |
| OXYCODONE | |  | 0 (NA%) | 24 (39%) |  | 6 | 17 | 1 |
| PETHIDINE | |  | 0 (NA%) | 2 (3.2%) |  | 0 | 1 | 1 |
| TRAMADOL | |  | 0 (NA%) | 21 (34%) |  | 12 | 9 | 0 |
| (Missing) | |  | 5,630 | 0 |  |  |  |  |
| (Missing) | |  |  |  |  | 0 | 0 | 5,630 |
| 1Median (IQR) or Frequency (%) | | | | | | | | |
| 2Wilcoxon rank sum test; Pearson's Chi-squared test; Fisher's exact test | | | | | | | | |