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| **Supplemental Digital Content 1:**  Number of subjects per ABP-700 dosing regimen and allocated premedication. Blue: ANVN-01, Green: ANVN-03, Maroon: ANVN-05, Orange: ANVN-02, Yellow: ANVN-04 | | | | | | |
|  | **No premed** | **Fentanyl** | **Remifentanil** | **Midazolam** | **Sufentanil** | **Fentanyl + Midzolam** |
| **Bolus** |  |  |  |  |  |  |
| 30 μg/kg | 5 |  |  |  |  |  |
| 100 μg/kg | 5 |  |  |  |  |  |
| 175 μg/kg | 5 |  |  |  |  |  |
| 250 μg/kg | 5 | 5 |  |  |  |  |
| 350 μg/kg | 5 | 5 |  |  |  |  |
| 500 μg/kg | 5 |  |  |  |  |  |
| 750 μg/kg | 5 |  |  |  |  |  |
| 1000 μg/kg | 5 |  |  |  |  |  |
| 250 μg/kg |  | 8 | 4 | 16 | 8 | 8 |
| 350 μg/kg |  |  |  | 16 |  | 8 |
| Slow Bolus- 175 μg/kg/min for 4 minutes |  | 16 |  |  |  |  |
| **7 min infusion** |  |  |  |  |  |  |
| 100 μg/kg/min | 7 | 8 | 8 | 8 |  |  |
| 120 μg/kg/min | 8 | 8 |  | 8 |  |  |
| 140 μg/kg/min | 8 | 8 | 8 | 16 |  |  |
| 160 μg/kg/min | 8 |  |  |  |  |  |
| 140 μg/kg/min for 2 minutes - 100 μg/kg/min for 2 minutes- 90 μg/kg/min for 3 minutes |  | 7 | 8 |  |  |  |
| **30 min infusion (μg/kg/min)** |  |  |  |  |  |  |
| 30 μg/kg/min | 5 |  |  |  |  |  |
| 40 μg/kg/min | 5 |  |  |  |  |  |
| 50 μg/kg/min | 5 | 5 |  |  |  |  |
| 60 μg/kg/min | 5 |  |  |  |  |  |
| 350 μg/kg bolus followed immediately by 50 μg/kg/min infusion |  | 5 |  |  |  |  |
| two 250 μg/kg bolus, separated by 2 min followed immediately by 50 μg/kg/min infusion |  | 5 |  |  |  |  |
| 100 μg/kg/min for 3 min, 75 μg/kg/min for 6 min, 58 μg/kg/min for 21 min | 4 | 4 |  |  |  |  |
| 25 μg/kg/min ABP-700for 10 minutes followed by 20 μg/kg/min for 20 minutes |  | 4 |  |  |  |  |
| 50 μg/kg/min ABP-700 for 5 minutes followed by 30 μg/kg/min for 25 minutes |  | 8 |  |  |  |  |
| 70 μg/kg/min ABP-700 for 3 minutes followed by 40 μg/kg/min for 27 minutes |  | 8 | 7 |  |  |  |
| 80 μg/kg/min ABP-700 for 3 minutes followed by 45 μg/kg/min for 27 minutes |  | 8 | 8 |  |  |  |
| 90 μg/kg/min ABP-700 for 3 minutes followed by 50 μg/kg/min for 27 minutes |  | 4 |  |  |  |  |
| **Titrate to effect: adjustment infusion rate if MOAA/S ≤ 4** |  |  |  |  |  |  |
| 80 μg/kg/min ABP-700 for 1-3 minutes followed by 45 μg/kg/min for 29-27 minutes |  |  | 1 |  |  |  |
| 80 μg/kg/min ABP-700 for 3-4 minutes followed by 50 μg/kg/min for 27-26 minutes |  |  | 2 |  |  |  |
| 80 μg/kg/min ABP-700 for 4-5 minutes followed by 60 μg/kg/min for 26-25 minutes |  |  | 5 |  |  |  |