Appendix 1

Hyperthermia therapy using Mitomycin C.

Bladder hyperthermia was achieved using the Combat BRS device (Combat Medical, UK) (**Supplemental Figure 1**). The device uses an aluminum heat exchanged and peristaltic pump to heat and recirculate chemotherapy solutions in the bladder. We used the disposable package which contained a 16 F 3-way catheter with a temperature monitoring thermistor in its tip. Mitomycin C (MMC) was reconstituted and diluted using sterile technique in a laminar flow hood and stored in a capped 60 mL syringe in a double bag Ziploc bag transport system. Each treatment used 40 mg of MMC in 50 ml of water for a continuous 60 minutes.

Institutions included in the study:

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| Hospital | Country | Number of Patients |
| Hospital Universitario de Canarias | Spain | 163 |
| Hospital Universitario 12 Octubre | Spain | 148 |
| St. Barbara Hospital | Germany | 98 |
| Hospital Universitario Infanta Sofía, | Spain | 92 |
| Netherlands Cancer Institute | Netherlands | 91 |
| Hospital Universitario de Torrejón | Spain | 84 |
| Antonio Cardarelli Hospital | Italy | 77 |
| San Luigi Hospital Orbassano | Italy | 57 |
| Hospital Universitario de Burgos | Spain | 56 |
| Queen Alexandra Hospital | United Kingdom | 56 |
| Hospital Monforte de Lemos | Spain | 44 |
| Hospital Universitario La Fe | Spain | 38 |
| Hospital General Rio Carrion | Spain | 13 |
| Fundació Puigvert, Autonomous University of Barcelona | Spain | 11 |