Supplementary Materials: Crotoxin Conjugated to SBA-15 Nanostructured Mesoporous Silica Induces Long-Last Analgesic Effect in the Neuropathic Pain Model in Mice

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Table S1. Evaluation of lethality induced by CTX and CTX:SBA-15**.** The survival time of each animal was followed for 48 hours after administration (i.p.) of CTX (600, 300, 150, 75, 37.5 and 18.75 μg/kg) or complex CTX:SBA-15 (424, 212, 106, 53 and 26.5 μg/kg).

Compound	LD_{50} (μg/kg)	95% Confidence Intervals (μg/kg)
CTX	106	63.2–177.9
CTX:SBA-15	143.2	70.4–291.2

CTX- Crotoxin; CTX:SBA-15- CTX absorbed to SBA-15.

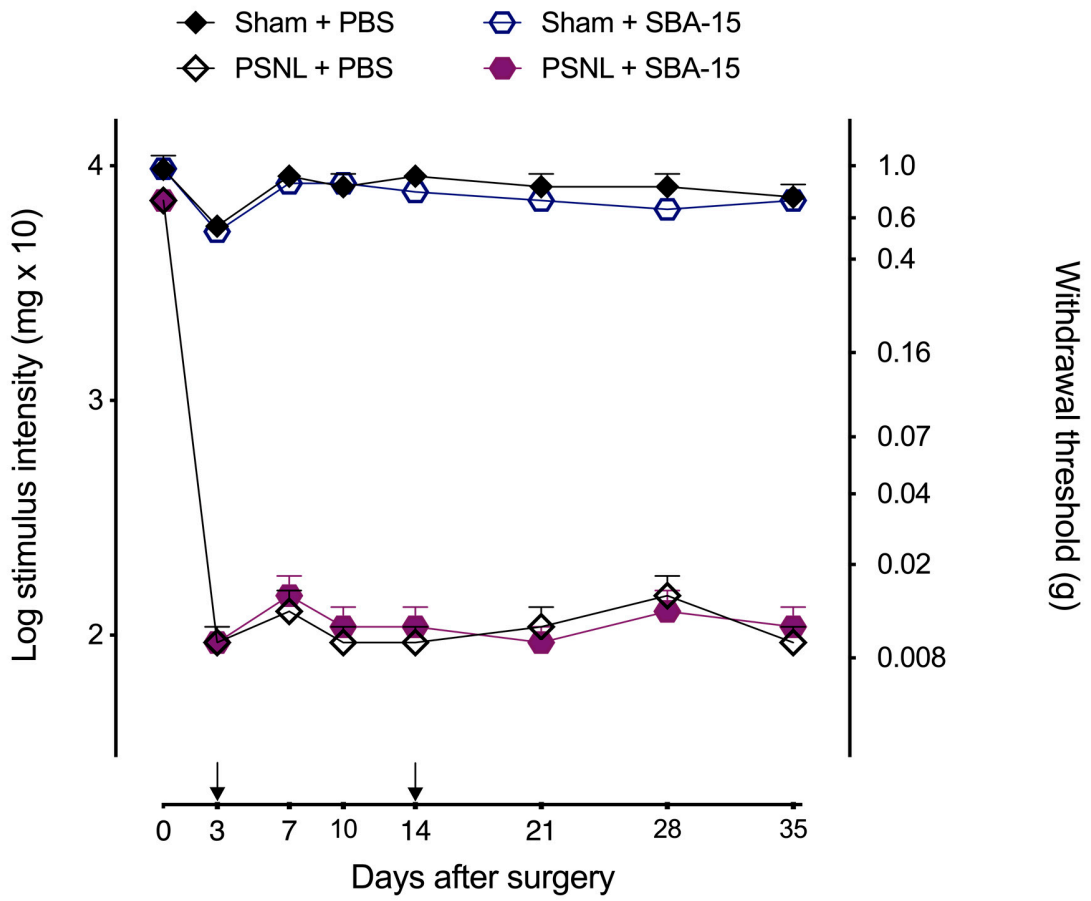

Figure S1. Effects of SBA-15 on mechanical hypernociception induced by PSNL. The pain threshold of the animals underwent surgery (sham and PSNL) was evaluated from day 0 (baseline) to 35th days after PSNL. Measurements were performed 1 hour after a single administration of SBA-15 (486 μg/kg, s.c.) on the 3rd

and 14th days after surgery. Arrows indicate the days of SBA-15 administration. The nociceptive threshold was assessed using the von Frey filaments. Results are expressed as mean (± SEM) n = 6.

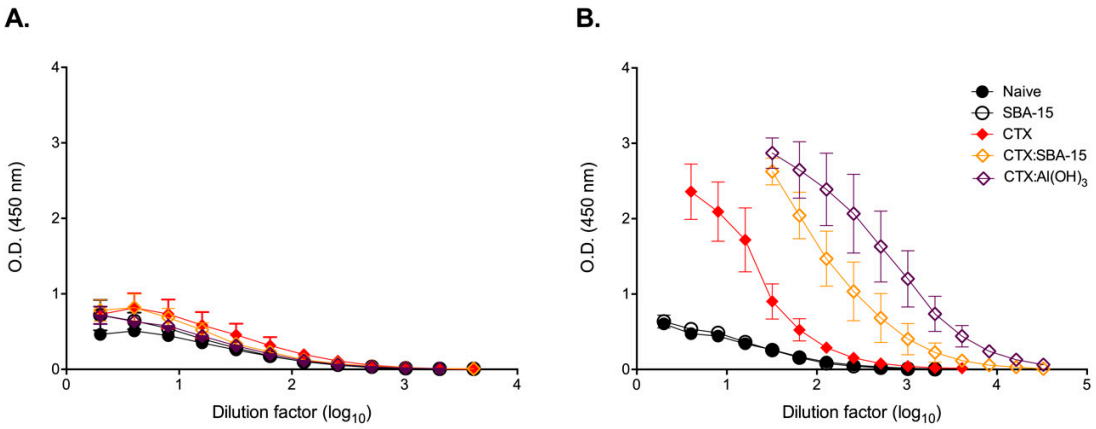

Figure S2. Titration of anti-CTX antibodies after administration of CTX and CTX:SBA-15 in HIII mice. Representative curves obtained after the first dose (A) and booster dose (B) of SBA-15 (486 μg/kg, s.c.), CTX (40 μg/kg, s.c.), CTX:SBA-15 (54 μg/kg, s.c.) or CTX:Al(OH)3 (54 μg/kg, s.c.). Results are expressed as the mean (± SEM) n = 4–5.