

**Supplementary Table 1.** Certainty of Evidence according to GRADE Approach

No. of studies	Certainty assessment							No. of patients		Effect size (95% CI)	Certainty
	Study design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other considerations	[intervention]	[comparison]			
Therapeutic impact of theophylline/aminophylline on PDPH severity											
6	Randomized trials	Serious*	Not serious <sup>†</sup>	Serious <sup>‡</sup>	Serious <sup>§</sup>	Very strong association	194	195	SMD: -1.67 (-2.29 to -0.85)	⊕⊕⊕⊕ Moderate	
Prophylactic impact of aminophylline on PDPH											
9	Randomized trials	Not serious	Not serious	Serious <sup>‡</sup>	Very serious <sup>  </sup>	None	153/539 (28.4%)	168/539 (31.2%)	RR: 0.74 (0.42 to 1.31)	⊕○○○ Very low	

GRADE: Grading of Recommendations Assessment, Development, and Evaluation, PDPH: post-dural puncture headache; CI: confidence interval, SMD: standardized mean difference, RR: relative risk. \*Serious risk of bias since 50% of the included studies showed a high risk of bias. Downgraded. <sup>†</sup>Serious inconsistency because of significant substantial between-study heterogeneity. However, I<sup>2</sup> was reduced in the subgroup analysis. Not downgraded. <sup>‡</sup>Serious indirectness since most of the studies were done in Asia. Downgraded. <sup>§</sup>Serious imprecision since the effect size was lower than the minimally important difference for clinical relevance (1.8–5.2), and the 95%CI was wide. Downgraded. <sup>||</sup>Very serious imprecision since the non-significant effect size and wide 95% CI. Downgraded.