

Table S2 Characteristics, quality and safety of clonidine studies.

Non randomised controlled trials				
Study	Overview	Safety	Efficacy	Quality
Ambrose 2000[51]	<p>Population: 30 ventilated children aged 10 years and under.</p> <p>Interventions: Group 1: 10 unparalysed ventilated patients were administered a clonidine infusion of 0.2 -1.0 mcg/kg/hr.</p> <p>Group 2: a further 10 unparalysed ventilated patients were administered a clonidine infusion of 0.2-2mcg/kg/hr.</p> <p>Group 3: 10 postoperative cardiac surgical patients were administered a fixed infusion rate of 1.0mcg/kg/hr.</p> <p>All patients were administered a background dose of midazolam</p>	<p>Cardiovascular/respiratory: There were no “adverse effects on cardiovascular performance” in the two groups of unparalysed ventilated patients (group 1 and 2). The postoperative cardiac surgical patients (Group 3) had no significant change over 6 hours in heart rate, blood pressure or derived cardiac index.</p> <p>“Bradycardia and hypotension were not recorded in any patient”</p> <p>Withdrawal: not assessed</p> <p>Neurological: not assessed</p>	<p>Group 1: Two patients failed to maintain adequate sedation at the maximum infusion of clonidine 1 mcg/kg/hr, but the other 8 patients were adequately sedated for the entire study period.</p> <p>Group 2: Adequate sedation was recorded in 602 of the 672 hourly assessments (89.5%) and there were no failures.</p> <p>Group 3: Not described</p>	<p>Ascertainment of AE data Cardiovascular: actively sought. Methods for haemodynamic assessment clearly described.</p> <p>Withdrawal: Not assessed Neurological: not assessed</p> <p>Reporting of AE data: Data presented numerically by intervention group. 2 patients were withdrawn from group 1 as there was a failure to maintain adequate sedation. Data up until withdrawal were included in the safety analysis for these patients.</p>
Case reports/case series				

Study	Case series/case report	Summary	Adverse effects
Lyons 1996[52]	Case report	11 year old boy who suffered second and third degree burns to 78% of his body was intubated and ventilated using synchronised intermittent mandatory ventilation. Clonidine was introduced intravenously for pain relief after a rise in morphine requirements at a rate of 7.5 to10mcg every 4 hours	When the rate of clonidine infusion was increased from 7.5 to10mcg every 4 hours haemodynamic parameters were not adversely affected.