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<td>Author(s)</td>
<td>Cheuk, KLD; Wong, VCN; Wraige, E; Baxter, P; Cole, A</td>
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Surgery for scoliosis in Duchenne muscular dystrophy (Review)

Cheuk DKL, Wong V, Wraige E, Baxter P, Cole A

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Surgery for scoliosis in Duchenne muscular dystrophy (Review)

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ABSTRACT

Background
Scoliosis in people with Duchenne muscular dystrophy is usually progressive and treated with surgery. However, it is unclear whether the existing evidence is sufficiently scientifically rigorous to support a recommendation for spinal surgery for most people with Duchenne muscular dystrophy and scoliosis. This is an updated review and an updated search was undertaken in which no new studies were found.

Objectives
To determine the effectiveness and safety of spinal surgery in people with Duchenne muscular dystrophy with scoliosis. We intended to test whether spinal surgery is effective in increasing survival, improving respiratory function, improving quality of life and overall functioning; and whether spinal surgery is associated with severe adverse effects.

Search methods
We searched the specialized registers of the Cochrane Neuromuscular Disease Group (31 July 2012), MEDLINE (January 1966 to July 2012), EMBASE (January 1947 to July 2012), CENTRAL (2012, Issue 7 in the Cochrane Library), CINAHL Plus (January 1937 to July 2012), Proquest Dissertation and Thesis Database (January 1980 to July 2012), and the National Institute of Health Clinical Trials Database (July 2012). No language restrictions were imposed.

Selection criteria
We planned to include controlled clinical trials using random or quasi-random allocation of treatment evaluating all forms of spinal surgery for scoliosis in people with Duchenne muscular dystrophy in the review. The control interventions would have been no treatment, non-operative treatment, or a different form of spinal surgery.

Data collection and analysis
Two authors independently examined the search results and evaluated the study characteristics against inclusion criteria to decide which ones would be included in the review.

Main results
On searching, 47 studies were relevant but none met the inclusion criteria for the review, because they were not clinical trials but prospective or retrospective reviews of case series.
Authors’ conclusions

Since there were no randomized controlled clinical trials available to evaluate the effectiveness of scoliosis surgery in people with Duchenne muscular dystrophy, no evidence-based recommendation can be made for clinical practice. People with scoliosis should be informed about the uncertainty of benefits and potential risks of surgery for scoliosis. Randomized controlled trials are needed to investigate the effectiveness of scoliosis surgery, in terms of quality of life, functional status, respiratory function and life expectancy.

PLAIN LANGUAGE SUMMARY

Scoliosis surgery for people with Duchenne muscular dystrophy

Scoliosis, curvature of the spine, is common in people with Duchenne muscular dystrophy. It is usually progressive and surgery is often performed aiming to halt its progression, improve cosmetic appearance, facilitate care, preserve upper limb and respiratory function, and hopefully increase life expectancy. However, there were no randomized controlled clinical trials available to evaluate the effectiveness of scoliosis surgery. Randomized controlled clinical trials are needed in this group of patients to evaluate the benefits and risks of different surgical treatments. This is an updated review and an updated search was undertaken in which no new studies were found.

BACKGROUND

Duchenne muscular dystrophy (DMD) is an inherited X-linked muscular dystrophy caused by mutations in the dystrophin gene. It is characterized by progressive dystrophic changes in skeletal and cardiac muscle. Progressive weakness in affected children results in loss of ambulation at a mean age of 9.5 years (Van Essen 1997). There is progressive cardiomyopathy and respiratory failure occurs secondary to respiratory muscle weakness. The mean survival in the absence of ventilatory support is 19.5 years (Van Essen 1997). In 90% death is the result of respiratory failure and in 10% the result of cardiac involvement. Currently there is no proven effective curative treatment for this debilitating disease. A systematic review has found that glucocorticoid therapy improves muscle strength and function in the short-term. However, adverse effects were common and long-term benefits are uncertain (Manzur 2008).

Spinal deformity, especially scoliosis, is progressive in the majority of people with DMD (Galasko 1995; Miller 1985). From the onset of spinal deformity, progression can be extremely rapid and impair unsupported sitting ability and further compromise the respiratory and cardiac function (Hsu 1983). Kurz observed a 4% decrease in vital capacity for every 10% progression of the spinal curve in people with DMD (Kurz 1983). Galasko found that on average, vital capacity decreases by 8% per year in patients with scoliosis secondary to DMD (Galasko 1992). Long-term corticosteroid treatment may slow the progress of scoliosis in people with DMD and may reduce the need for surgery (Dooley 2010), but adverse effects are frequent (Alman 2004). Non-operative treatment such as bracing might not prevent the progression of this kind of spinal deformity because of the progressive nature of the underlying muscle disease (Cambridge 1987; Colbert 1987). Therefore, non-operative treatment is usually considered only in exceptional cases when a person refuses surgery or when a person has a very advanced deformity with poor general health (Forst 1997; Heller 1997; McCarthy 1999).

Spinal fusion surgery with instrumentation remains the mainstay of treatment for people with DMD with scoliosis. Commonly used techniques are either based on sublaminar segmental wiring, such as Luque instrumentation, or the modern variants based on segmental pedicle screw and hook fixation such as Isola, TSRH or Universal Spine system. Two stainless steel or titanium rods are contoured to the desired spinal shape, and the spine reduced onto the rods, either with the sublaminar wires or segmental screws and hooks. Pelvic fixation is rarely required in DMD scoliosis and the Galveston technique of rod insertion into the ileum, or more modern screw fixation can be used in some circumstances. Postoperative bracing is not required with modern fixation techniques.

The potential advantages of surgery described in the literature include increased comfort and sitting tolerance (Bridwell 1999; Cambridge 1987; Marchesi 1997; Matsumura 1997; Miller 1991; Miller 1992; Rice 1998; Rideau 1984; Shapiro 1992), cosmetic improvement (Bellen 1993; Bridwell 1999), no need for orthopaedic braces (Bellen 1993; Colbert 1987; Miller 1985; Noble Jamieson 1986), easier nursing care by parents (Bellen 1993) and pain relief (Bellen 1993; Galasko 1977; Miller 1991). Nevertheless, the effects of spinal surgery on respiratory function...
and life expectancy are still controversial. Some studies reported that spinal fusion had no effects on the natural deterioration of respiratory function of people with DMD (Kinali 2006; Miller 1988; Miller 1992; Shapiro 1992), at short-term and five-year follow-up (Miller 1991). In contrast, several studies (Galasko 1992; Galasko 1995; Rideau 1984; Velasco 2007) reported stabilization of vital capacity in people surgically treated for two to eight years. Regarding life expectancy, Galasko observed a lower mortality in people surgically treated (Galasko 1992; Galasko 1995). However, other studies reported that spinal surgery did not improve life expectancy (Chataigner 1998; Gayet 1999; Kennedy 1995; Kinali 2006; Miller 1988). Adverse effects and complications during and after surgery are not uncommon, including ventilator-associated pneumonia (iatrogenic, in the post-operative period), wound dehiscence, surgical wound infection, haemorrhage, loosening of fixation, pseudarthrosis, deteriorated respiratory function and increased difficulty with hand to head motions.

A randomized trial has demonstrated that although tendon surgery in people with DMD may correct deformities, it might also result in more rapid deterioration of function in some patients and there were no beneficial effects on strength or function (Manzur 1992). With increasing use of non-invasive ventilation (NIV) in DMD patients with respiratory insufficiency which may prolong the life expectancy, it is unclear to what extent increased survival is related to NIV rather than to other interventions, including scoliosis surgery. It remains uncertain whether the existing evidence is sufficiently scientifically rigorous to recommend spinal surgery for most patients with DMD and scoliosis. In this systematic review, we evaluated the effectiveness of various forms of spinal surgery to prolong life expectancy, retard the natural deterioration of respiratory function, and improve quality of life in people with DMD. We wanted to evaluate whether the benefits outweigh the risks of surgery in general and determine which patient subgroups are most likely to benefit. The review has been updated, most recently in 2012.

**OBJECTIVES**

The objectives of this systematic review were to determine the effectiveness and safety of spinal surgery in people with DMD with scoliosis. We intended to test the following hypotheses:

1. Whether spinal surgery is effective in increasing survival;
2. Whether spinal surgery can improve respiratory function in the short-term and long-term;
3. Whether spinal surgery can improve quality of life and overall functioning;
4. Whether spinal surgery is associated with severe adverse effects.

**METHODS**

**Criteria for considering studies for this review**

**Types of studies**

We planned to include controlled clinical trials using random or quasi-random allocation of treatment in the review.

**Types of participants**

People with Duchenne muscular dystrophy (defined as progressive limb girdle weakness with at least one of: (1) dystrophic changes on muscle biopsy with reduced or absent dystrophin staining; (2) deletion, duplication or point mutation of dystrophin gene) and all degrees of scoliosis documented by appropriate x-rays would be included.

It was possible that this definition might have resulted in the inclusion of some individuals with an intermediate or severe Becker phenotype. However, the inclusion of only biopsy proven dystrophin negative cases could potentially result in the loss of some important data.

**Types of interventions**

We planned to include trials evaluating all forms of spinal surgery for scoliosis in the review. The control interventions were to be no treatment, non-operative treatment, or a different form of spinal surgery.

**Types of outcome measures**

**Primary outcomes**

1. Survival: to allow for studies using different follow-up periods, we planned to use hazard ratios from survival data regression analysis.

**Secondary outcomes**

1. Respiratory function, as measured by pulmonary function tests such as forced vital capacity (FVC); medium-term (3 to 12 months), and long-term (more than 12 months). The results from studies with differing lengths of follow-up were to be weighted appropriately to allow for this.
2. Medium and long-term disability as measured by validated scales such as the Barthel index or Functional Independent Measure.
3. Medium and long-term quality of life as measured by validated scales such as the 36-Item Short-Form Health Status Survey (SF-36).
4. Rate of progression of scoliosis, as measured by change of Cobb angle per year.
5. Frequency of severe adverse effects and complications, such as death related to surgery, deep surgical wound infection, wound dehiscence, loosening of fixation, pneumonia, pseudarthrosis, need for revision surgery.

Data extraction and management

We would have extracted the following data:

(1) Study methods
(a) Design (e.g. randomized or quasi-randomized).
(b) Randomization method (including list generation)
(c) Method of allocation concealment
(d) Blinding method
(e) Stratification factors

(2) Participants
(a) Inclusion/exclusion criteria
(b) Number (total/per group)
(c) Age distribution
(d) Severity of scoliosis
(e) Level of scoliosis
(f) Baseline respiratory function
(g) Associated morbidities, e.g. cardiomyopathy
(h) Previous treatments, including corticosteroids
(i) Pre-treatment quality of life and functional status, as measured by validated scales

(3) Intervention and control
(a) Type of spinal surgery
(b) Type of control
(d) Details of control treatment including duration of non-operative treatment
(e) Details of co-interventions

(4) Follow-up data
(a) Duration of follow-up
(b) Loss to follow-up

(5) Outcome data as described above

(6) Analysis data
(a) Methods of analysis (intention-to-treat/per-protocol analysis)
(b) Comparability of groups at baseline (yes/no)
(c) Statistical techniques
We planned that data would be entered into Review Manager (RevMan) by one review author and then checked by the second author.
Assessment of risk of bias in included studies
We planned to evaluate the validity of the trials by the following criteria:

(1) Selection bias
(a) Was allocation of participants to treatment and control groups randomized?
(b) Was allocation concealed?

(2) Performance bias
(a) Were participants in the comparison groups treated differently apart from the study treatments?
(b) Was there blinding of participants and personnel?

(3) Attrition bias
(a) Were there systematic differences between the comparison groups in the loss of participants from the study?
(b) Were analyses by intention-to-treat?

(4) Detection bias
(a) Were those assessing outcomes of the intervention blinded to the assigned intervention?

(5) Reporting bias
(a) Were there systematic differences between reported and unreported findings (incomplete outcome data)?

We planned to summarize the quality of a trial into one of the three categories:
A. Low risk of bias: all the validity criteria met.
B. Moderate risk of bias: one or more validity criteria partly met but none are not met.
C. High risk of bias: one or more criteria not met.

Measures of treatment effect
We planned to use risk ratio (RR) estimations with 95% confidence intervals (CI) for binary outcomes. We planned to use mean difference estimations with 95% CI for continuous outcomes. All analyses would include all participants in the treatment groups to which they were allocated.

Dealing with missing data
We planned to contact authors of included studies to supply missing data. We would have assessed missing data and drop-outs/attrition for each included study, and assess and discuss the extent to which the results and conclusions of the review could be altered by the missing data. If less than 70% of patients allocated to the treatments were not reported on at the end of the trial, for a particular outcome, we would not use those data as they would have been considered to be too prone to bias.

Assessment of heterogeneity
We planned to assess clinical heterogeneity by comparing the distribution of important participant factors between trials (age, respiratory function, severity and level of scoliosis, associated diseases), and trial factors (randomization concealment, blinding of outcome assessment, losses to follow-up, treatment type, co-interventions). We would assess statistical heterogeneity by examining I² (Higgins 2002), a quantity which describes approximately the proportion of variation in point estimates that is due to heterogeneity rather than sampling error. In addition, we would use a Chi² test for homogeneity to determine the strength of evidence that heterogeneity was genuine.

Assessment of reporting biases
We would have drawn funnel plots (estimated differences in treatment effects against their standard error) if sufficient studies were found. Asymmetry could be due to publication bias, but could also be due to a relationship between trial size and effect size. In the event that a relationship was found, we would examine clinical diversity of the studies (Egger 1997).

Data synthesis
Where the interventions were the same or similar enough, we planned to synthesize results in a meta-analysis if there was no important clinical heterogeneity. If no significant statistical heterogeneity was present, we planned to synthesize the data using a fixed-effect model. Otherwise we would use a random-effects model for the meta-analysis.

Adverse events
Since adverse events were rarely adequately dealt with by randomized studies alone because the numbers were small and follow-up too short, we planned to discuss adverse events taking into account the non-randomized literature.

Cost-benefit analyses
We planned to consider cost-effectiveness of interventions where relevant data were available.

Subgroup analysis and investigation of heterogeneity
If data permitted, we planned to conduct sub-group analyses for:
1. different age groups (younger than 12 years, 12 to 18 years, older than 18 years);
2. different degrees of pre-existing respiratory impairment (mild, severe);
3. different severity of scoliosis (moderate, severe);
4. previous corticosteroid treatments (yes, no).

Sensitivity analysis

We planned to undertake sensitivity analyses to assess the impact of study quality. These would have been undertaken including:
1. all studies;
2. only those with low risk of selection bias;
3. only those with low risk of performance bias;
4. only those with low risk of attrition bias;
5. only those with low risk of detection bias.

Sensitivity analysis would also be performed including and excluding subjects who might have Becker muscular dystrophy or an intermediate phenotype to see whether this would alter any of the results.

RESULTS

Description of studies

See: Characteristics of excluded studies.

In July 2012, a total of 80 studies were found on electronic search of the databases (Cochrane Neuromuscular Disease Group Registry: 2 studies, MEDLINE: 17 studies, EMBASE: 11 studies, CENTRAL: 1 study, CINAHL Plus: 13 studies, Proquest Dissertation and Thesis Database: 35 studies, and NIH Clinical Trials Database: 1 study). An additional 32 studies were identified on searching the reference lists of relevant studies. After duplicates were removed, a total of 105 studies were screened. Fifty-eight of these studies were excluded as they did not focus on Duchenne muscular dystrophy or scoliosis surgery, or were narrative reviews. We examined the remaining 47 studies in detail but none of these satisfied the inclusion criteria. All these studies were prospective or retrospective case series and were not clinical trials. Most of these reviews also did not have a control group for comparisons. Where a control group was included, the controls were people who refused surgery or were assigned a different treatment modality by the treating surgeons without randomization or quasi-randomization. We therefore excluded these studies from further analyses because of significant propensity for confounding and bias. The flow of studies is shown in Figure 1.
Figure 1. Study flow diagram.

80 records identified through database searching

32 additional records identified through other sources

105 records after duplicates removed

105 records screened

58 records excluded

47 full-text articles assessed for eligibility

47 full-text articles excluded, with reasons

0 study included in qualitative synthesis

0 study included in quantitative synthesis (meta-analysis)
**Risk of bias in included studies**

Not applicable.

**Effects of interventions**

No controlled trials met the inclusion criteria of the review for further analyses.

**DISCUSSION**

Despite a comprehensive search strategy used for this review, no randomized controlled trial (RCT) of surgery for scoliosis in people with Duchenne muscular dystrophy was identified. Instead we found many retrospective reviews or case series of patients with Duchenne muscular dystrophy and scoliosis treated with surgery. These studies showed varying results and had different conclusions. Although most agreed that surgery can improve patients’ quality of life and functional status in terms of sitting posture, upper limb function and ease of care, most failed to show a significant improvement in respiratory function or long-term survival, and short-term and long-term postoperative complications occurred not uncommonly.

However, a closer look at the relevant studies excluded might be helpful for guiding future clinical trials of scoliosis surgery for patients with DMD (Table 1). These 47 case series included 5 to 70 patients who had undergone scoliosis surgery. Nine of these studies also included a comparison group of 21 to 115 patients without surgery (Eagle 2007; Galasko 1992; Galasko 1995; Kennedy 1995; Kinali 2006; Miller 1988; Miller 1991; Miller 1992; Sakai 1977).

**Outcome measures and comparisons**

The studies had different objectives and focused on different outcomes. Most studies aimed to investigate whether spinal surgery improves the degree of scoliosis in the short-term (immediate post-operative period) and in the long-term (years later). Most studies used Cobb angle and degree of pelvic obliquity as outcome measures and described early and late complications of surgery. Some studies also reported duration of hospitalization (Harper 2004; Rideau 1984; Sengupta 2002; Sussman 1984), peri-operative mortality (Alman 1999; Bentley 2001; Brook 1996; Cambridge 1987; Cervellati 2004; Chataigner 1998; Dubousset 1983; Eagle 2007; Gaine 2004; Galasko 1992; Galasko 1995; Gayet 1999; Granata 1996; Hahn 2008; Harper 2004; Heller 2001; Hopf 1994; Kennedy 1995; LaPrade 1992; Marchesi 1997; Marsh 2003; Matsumura 1997; Modi 2009; Rideau 1984; Sakai 1977; Sengupta 2002; Shapiro 1992; Thacker 2002; Weimann 1983) and length of survival (Eagle 2007; Kinali 2006; Miller 1992) in people who had undergone scoliosis surgery. Many studies reported the change in respiratory function after operation (Brook 1996; Cervellati 2004; Chataigner 1998; Dubousset 1983; Eagle 2007; Galasko 1992; Galasko 1995; Gayet 1999; Granata 1996; Kennedy 1995; Kinali 2006; Matsumura 1997; Mehdian 1989; Miller 1988; Miller 1991; Miller 1992; Rideau 1984; Shapiro 1992; Thacker 2002; Velasco 2007). The parameters used included vital capacity, peak expiratory flow rate and forced vital capacity in one second. A few studies also reported patient oriented subjective outcomes such as quality of life, self-image, cosmetic appearance, pain and patient satisfaction (Bentley 2001; Bridwell 1999; Granata 1996; Matsumura 1997; Miller 1991; Miller 1992; Rideau 1984). While most studies evaluated the outcomes of spinal surgery in general, some studies tried to compare different surgical techniques, such as Luque instrumentation versus Isola pedicle screw (Gaine 2004), sublaminar wiring versus intraspinal segmental wiring (LaPrade 1992), Luque instrumentation versus distal instrumentation with Galveston construct and rigid cross-linking (Brook 1996), Harrington-Luque instrumentation versus modified Luque instrumentation (Bentley 2001), Harrington instrumentation versus Luque instrumentation versus segmental spinal instrumentation with fusion (Sussman 1984), sublaminar instrumentation versus pedicle screw versus a hybrid system (Arun 2010), or autogenous versus allogenous bone graft (Nakazawa 2010). Some studies also compared the outcomes of spinal fusion to different extents (Alman 1999; Bridwell 1999; Gaine 2004; Mubarak 1993; Sengupta 2002; Modi 2010), such as fusion to L5 versus fusion to sacrum. Some studies compared surgical outcomes in patients with different pre-operative respiratory function (Harper 2004; Marsh 2003; Matsumura 1997; Sussman 1984).

**Outcomes on survival**

Most studies did not demonstrate obvious benefits of scoliosis surgery in terms of prolonging survival (Brook 1996; Cervellati 2004; Chataigner 1998; Gayet 1999; Granata 1996; Hahn 2008; Kennedy 1995; Kinali 2006; Mehdian 1989; Miller 1988; Miller 1991; Miller 1992; Shapiro 1992; Thacker 2002). There was one study showing that when combined with nocturnal ventilation, patients after spinal surgery has longer median survival (30 years) compared with patients on nocturnal ventilation alone (22.2 years) (Eagle 2007). There was another study showing that survival rate was higher at five years after surgery (61%) compared to those who refused surgery (23%) (Galasko 1995). In general the age at death in patients with or without surgery was highly variable in the case series. Although most deaths could be attributed to respiratory infection, respiratory failure, progressive cardiomyopathy...
and sudden cardiac death, the cause of death could not be ascertained in many cases. However, the age and causes of death did not seem to differ between patients with or without surgery. Peri-operative mortality is generally uncommon. Most studies reported no peri-operative mortality (Alman 1999; Bellen 1993; Bentley 2001; Bridwell 1999; Brook 1996; Cambridge 1987; Chataigner 1998; Dubouset 1983; Eagle 2007; Galasko 1992; Galasko 1995; Gayet 1999; Hopf 1994; Kennedy 1995; Kinis 2000; LaPrade 1992; Marchesi 1997; Marsh 2003; Matsumura 1997; Mehdian 1989; Miller 1992; Mubarak 1993; Nakazawa 2010; Rice 1998; Rideau 1984; Sakai 1977; Sengupta 2002; Stricker 1996; Sussman 1984; Takaso 2010; Thacker 2002; Weimann 1983), while some studies reported peri-operative mortality ranging from 1.4% to 5% (Modi 2009; Gaine 2004; Cervellari 2004; Granata 1996; Hahn 2008; Harper 2004; Heller 2001; Shapiro 1992).

### Outcomes on respiratory function

Galasko found that forced vital capacity could be stabilized for three years and peak expiratory flow rate maintained for up to five years after spinal fusion (Galasko 1992; Galasko 1995). Rideau also found that vital capacity could be maintained static for two years (Rideau 1984); and three participants in Matsumura’s study had increased forced vital capacity after operation (Matsumura 1997). Velasco found that the average rate of decline of FVC reduced from 4% per year to 1.75% per year after surgery (Velasco 2007). However, most studies did not demonstrate obvious benefits of scoliosis surgery in terms of respiratory function (Brook 1996; Chataigner 1998; Cervellari 2004; Eagle 2007; Gayet 1999; Granata 1996; Hahn 2008; Kennedy 1995; Kinali 2006; Mehdian 1989; Miller 1992; Mubarak 1993; Nakazawa 2010; Rice 1998; Rideau 1984; Sakai 1977; Sengupta 2002; Stricker 1996; Sussman 1984). However, most studies did not demonstrate obvious benefits of scoliosis surgery in terms of respiratory function (Brook 1996; Chataigner 1998; Cervellari 2004; Eagle 2007; Gayet 1999; Granata 1996; Hahn 2008; Kennedy 1995; Kinali 2006; Mehdian 1989; Miller 1992; Mubarak 1993; Nakazawa 2010; Rice 1998; Rideau 1984; Sakai 1977; Sengupta 2002).

### Functional outcome and quality of life

In general, previous descriptive studies suggested that surgical correction of scoliosis resulted in better sitting position, quality of life and patient satisfaction (Bentley 2001; Bridwell 1999; Cambridge 1987; Granata 1996; Marchesi 1997; Matsumura 1997; Miller 1991; Miller 1992; Rice 1998; Rideau 1984; Sakai 1977; Shapiro 1992).

### Complications of spinal surgery

Severe complications after spinal surgery are not infrequent and occur in up to 68% of patients (Modi 2009). These include cardiac arrest (Bentley 2001), cardiac arrhythmia (Harper 2004), heart block (Galasko 1992), respiratory failure requiring tracheostomy (Chataigner 1998; Galasko 1992; Galasko 1995; Harper 2004; Heller 2001; Marsh 2003) or mechanical ventilation post-operatively (Bentley 2001; Brook 1996; Heller 2001; Modi 2009), massive bleeding (Heller 2001; Modi 2008a), pneumonia (Bentley 2001; Galasko 1992; Harper 2004; Heller 2001; Modi 2009; Rideau 1984), pleural effusion (Harper 2004; Modi 2009), hemothorax or pneumothorax (Bentley 2001; Heller 2001; Modi 2009), spinal cord injury (Modi 2009), colonic perforation (Bentley 2001), bladder dysfunction (Bentley 2001; Hopf 1994), urinary tract infection (Modi 2009), deep wound infection (Arun 2010; Modi 2008a; Modi 2009; Sengupta 2002), infection necessitating removal or revision of surgical implants (Eagle 2007; Heller 2001), failure of implants (Arun 2010; Bentley 2001; Gaine 2004; Stricker 1996), dislodgement or displacement of implants (Heller 2001; LaPrade 1992; Matsumura 1997), loosening of implants (Arun 2010; Modi 2009; Sengupta 2002), mechanical problems requiring revision surgery (Bentley 2001; Galasko 1992; Gayet 1999; Granata 1996; Sengupta 2002), pseudarthrosis (Gaine 2004; Thacker 2002), bone fracture (Alman 1999), pressure sores (Granata 1996; Modi 2009; Modi 2010), dural leak (LaPrade 1992) and deep vein thrombosis (Heller 2001). Several studies reported that postoperative complications were more frequent in patients with greater severity of scoliosis (Bentley 2001; Sakai 1977; Sussman 1984).

### Comparisons of different operative methods

In general, fusion to sacrum does not offer benefits over fusion to a more proximal level (Gaine 2004; Mubarak 1993; Rice 1998; Sengupta 2002), unless scoliosis is severe and pelvic obliquity is significant (Alman 1999; LaPrade 1992; Modi 2010). Although none of the surgical methods was uniformly better than others, Isola system (Gaine 2004) or segmental spinal fusion (Miller 1991; Miller 1992) might achieve better correction of deformity, and intraspinal wiring might result in shorter operative time and less blood loss compared to sublaminar wiring (LaPrade 1992). Pedicle screw system might also result in shorter operative time and less blood loss compared to sublaminar instrumentation system (Arun 2010).

No meta-analysis of these available data was performed because the retrospective non-randomized, uncontrolled studies were observational in nature and were prone to bias and confounding. There is currently an absence of high level evidence supporting scoliosis surgery in patients with Duchenne muscular dystrophy. There is also a lack of evidence for or against a particular modality of surgical approach. Controlled clinical trials with random allocation into treatment and control groups are needed before firm conclusions on the benefits and risks of scoliosis surgery in patient with DMD can be made.

In the absence of evidence it is our view that clinicians might need to consider anecdotal evidence and their personal experience as well as expert opinions as guidance for their decision on the best care for individual patient. Potential benefits on quality of life and

**Surgery for scoliosis in Duchenne muscular dystrophy (Review)**

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functional status as well as risks of morbidity and mortality should be fully discussed with the patients before embarking on surgery for scoliosis. Patients should also be informed about the uncertainty of benefits on long-term survival and respiratory function after scoliosis surgery.

**AUTHORS’ CONCLUSIONS**

**Implications for practice**

Since there were no RCTs available to evaluate the effectiveness of scoliosis surgery in people with Duchenne muscular dystrophy, no recommendation can be made for clinical practice.

**Implications for research**

RCTs are needed to investigate the effectiveness of scoliosis surgery, in terms of patients’ satisfaction, quality of life, functional status, respiratory function (forced vital capacity, forced expiratory volume in one second, peak expiratory flow) and survival. It should be feasible to randomize patients into surgery versus non-surgical management. Although placebo control treatment might not be feasible, random allocation of patients into different treatment groups is essential to avoid selection bias and ensure baseline comparability of different groups. Although blinding of patients and clinicians is almost impossible, blinding of outcome assessors is important and probably feasible. Quality of life and functional status should be assessed by validated questionnaires and instruments. The relative benefits and risks of different surgical treatment modalities and different extents of spinal fusion should also be investigated by RCTs. Stratifications by potentially important prognostic factors such as age, baseline respiratory function and severity of scoliosis should be considered.

**ACKNOWLEDGEMENTS**

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Arun 2010 [published data only]

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Bentley 2001 [published data only]

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Brook 1996 [published data only]

Cambridge 1987 [published data only]

Cervellati 2004 [published data only]

Chataigner 1998 [published data only]
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**Modi 2009  [published data only]**

**Modi 2010  [published data only]**

**Mubarak 1993  [published data only]**

**Nakazawa 2010  [published data only]**

**Rice 1998  [published data only]**

**Rideau 1984  [published data only]**

**Sakai 1977  [published data only]**

**Sengupta 2002  [published data only]**

**Shapiro 1992  [published data only]**

**Stricker 1996  [published data only]**

**Sussman 1984  [published data only]**

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**Thacker 2002  [published data only]**

**Velasco 2007  [published data only]**

**Weimann 1983  [published data only]**

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**Colbert 1987**

**Dooley 2010**

**Egger 1997**

**Forst 1997**

**Galasko 1977**
Heller 1997

Higgins 2002

Higgins 2011

Hsu 1983

Kurz 1983

Manzur 1992

Manzur 2008

McCarty 1999

Miller 1985

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Van Essen 1997

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Cheuk 2010

* Indicates the major publication for the study
### Characteristics of excluded studies [ordered by study ID]

<table>
<thead>
<tr>
<th>Study</th>
<th>Reason for exclusion</th>
</tr>
</thead>
<tbody>
<tr>
<td>Alman 1999</td>
<td>Retrospective case series, not clinical trial.</td>
</tr>
<tr>
<td>Arun 2010</td>
<td>Retrospective case series, not clinical trial.</td>
</tr>
<tr>
<td>Bellen 1993</td>
<td>Retrospective case series, not clinical trial.</td>
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<tr>
<td>Bentley 2001</td>
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<tr>
<td>Bridwell 1999</td>
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<tr>
<td>Brook 1996</td>
<td>Retrospective case series, not clinical trial.</td>
</tr>
<tr>
<td>Cambridge 1987</td>
<td>Retrospective case series, not clinical trial.</td>
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<tr>
<td>Cervellati 2004</td>
<td>Retrospective case series, not clinical trial.</td>
</tr>
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<td>Chataigner 1998</td>
<td>Retrospective case series, not clinical trial.</td>
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<tr>
<td>Dubousset 1983</td>
<td>Retrospective case series, not clinical trial.</td>
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<td>Eagle 2007</td>
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<td>Gaine 2004</td>
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<td>Galasko 1992</td>
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<td>Granata 1996</td>
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<tr>
<td>Hahn 2008</td>
<td>Retrospective case series, not clinical trial.</td>
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<td>Harper 2004</td>
<td>Prospective case series, not clinical trial.</td>
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<td>Heller 2001</td>
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<tr>
<td>Hopf 1994</td>
<td>Retrospective case series, not clinical trial.</td>
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<tr>
<td>Kennedy 1995</td>
<td>Retrospective case series, not clinical trial.</td>
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<td>Reference</td>
<td>Study Type</td>
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<tr>
<td>Kinali 2006</td>
<td>Retrospective case series, not clinical trial.</td>
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<tr>
<td>LaPrade 1992</td>
<td>Retrospective case series, not clinical trial.</td>
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<tr>
<td>Marchesi 1997</td>
<td>Retrospective case series, not clinical trial.</td>
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<tr>
<td>Marsh 2003</td>
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<td>Matsumura 1997</td>
<td>Retrospective case series, not clinical trial.</td>
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<td>Mehdian 1989</td>
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<td>Miller 1988</td>
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<td>Modi 2008a</td>
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<tr>
<td>Modi 2008b</td>
<td>Retrospective case series, not clinical trial.</td>
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<td>Mubarak 1993</td>
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<td>Nakazawa 2010</td>
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<td>Rice 1998</td>
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<td>Rideau 1984</td>
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<td>Sakai 1977</td>
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<td>Sengupta 2002</td>
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<td>Shapiro 1992</td>
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<td>Stricker 1996</td>
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<tr>
<td>Sussman 1984</td>
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<tr>
<td>Takas 2010</td>
<td>Prospective case series, not clinical trial.</td>
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<td>Reference</td>
<td>Study Design</td>
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<td>Thacker 2002</td>
<td>Retrospective case series, not clinical trial.</td>
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<tr>
<td>Velasco 2007</td>
<td>Retrospective case series, not clinical trial.</td>
</tr>
<tr>
<td>Weimann 1983</td>
<td>Retrospective case series, not clinical trial.</td>
</tr>
</tbody>
</table>
DATA AND ANALYSES

This review has no analyses.

ADDITIONAL TABLES

Table 1. Characteristics of excluded studies

<table>
<thead>
<tr>
<th>Study reference</th>
<th>Number of patients</th>
<th>Treatments</th>
<th>Outcome measures</th>
<th>Findings</th>
<th>Remarks</th>
</tr>
</thead>
<tbody>
<tr>
<td>Arun 2010</td>
<td>43</td>
<td>Sublaminar instrumentation (19) or hybrid sublaminar and pedicle screw (13) or pedical screw (11)</td>
<td>Cobb angle, flexibility index, blood loss, operating time, complications</td>
<td>Percentage correction of Cobb angle was 72.5 +/- 14.5% (Group A), 82 +/- 6% (Group B) and 82 +/- 8% (Group C). Flexibility indices were 60 +/- 6.33% (Group A), 70 +/- 4.65% (Group B) and 67 +/- 6.79% (Group C). Mean blood loss was 4.1 L (Group A), 3.2 L (Group B) and 2.5 L (Group C). Mean operating times were 300 min (Group A), 274 min (Group B) and 234 min (Group C). Complications: 3 wound infections and 2 implant failure (Group A), 1 implant failure (Group B), 1 wound infection and 1 partial screw pull out (Group C)</td>
<td>Concluded that pedicle screw system might be favored because of the lesser blood loss and surgical time</td>
</tr>
<tr>
<td>Alman 1999</td>
<td>48</td>
<td>Spinal fusion to L5 (38) or spinal fusion to sacrum (10) using multiple level sublaminar wires with either a modified unit rod with Galveston extensions to the pelvis cut-off, a modified rod with a cross-link placed at</td>
<td>Cobb angle, torso decompensation, sitting obliquity, spinal obliquity, need for revision surgery, mortality</td>
<td>Sitting obliquity and spinal obliquity increased in patients fused to L5. 2 patients had fracture of L5 lamina. 2 patients required revision surgery</td>
<td></td>
</tr>
<tr>
<td>Study</td>
<td>Patients</td>
<td>Methodology</td>
<td>Outcomes</td>
<td>Findings</td>
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<tr>
<td>Bellen 1993</td>
<td>47</td>
<td>Segmental spinal instrumentation according to Luque's technique</td>
<td>Mortality, complications. Many patients have general and pulmonary and mechanical complications</td>
<td>Concluded that a total spinal arthrodesis could probably be avoided in these patients, which often demonstrate a satisfying spontaneous fusion after instrumentation</td>
<td></td>
</tr>
<tr>
<td>Bentley 2001</td>
<td>101</td>
<td>Modified Luque (87), Harrington-Luque (14)</td>
<td>Cobb angle, pelvic obliquity, mortality, complications, patient satisfaction</td>
<td>Incidence of minor or temporary complications was high, but chiefly occurred in patients with very severe curves and considerable pre-existing immobility</td>
<td></td>
</tr>
<tr>
<td>Bridwell 1999</td>
<td>33</td>
<td>Posterior segmental spinal instrumentation applied from the upper thoracic spine (T2, T3, T4, T5) down to L5 or the sacrum and pelvis. Early in the series, patients with DMD with smaller curves (&lt; 40º) were fixed to L5. All had bilateral segmental fixation with Wisconsin or sublaminar wires at each level and at times with hook supplementation. All patients fused to the sacrum had Galveston or Galveston-Questionnaires to evaluate function, self-image, cosmesis, pain, pulmonary status, patient care, quality of life, satisfaction, radiographic data.</td>
<td>All patients seemed to have benefited from the surgery. Cosmesis, quality of life, and overall satisfaction rated the highest</td>
<td></td>
<td></td>
</tr>
<tr>
<td>Study</td>
<td>n</td>
<td>Procedure and Fixation</td>
<td>Parameter(s)</td>
<td>Outcome</td>
<td>Additional Information</td>
</tr>
<tr>
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<tr>
<td>Brook 1996</td>
<td>17</td>
<td>L-rod instrumentation (10), distal instrumentation with Galveston construct and rigid cross-linking (7)</td>
<td>Cobb angle and pelvic obliquity, %FVC, mortality, complications</td>
<td>Correction of Cobb angle better in the Galveston group (63% versus 51%). No pseudoarthroses or instrument failures in the Galveston group. Totally 4 patients had FVC &lt; 25%, 2 required ventilation postoperatively. No other respiratory complications. No peri-operative mortality</td>
<td>The effect of surgery on respiratory function remains uncertain</td>
</tr>
<tr>
<td>Cambridge 1987</td>
<td>14</td>
<td>Segmental spinal instrumentation (13), Harrington distraction rods (1)</td>
<td>Mortality, complications, sitting tolerance.</td>
<td>No peri-operative mortality. 1 required repeated re-intubation. All achieved excellent long-term sitting tolerance</td>
<td>Recommended posterior spinal fusion with segmental instrumentation when scoliosis &gt; 30º. Spinal fusion did not increase life expectancy or pulmonary function</td>
</tr>
<tr>
<td>Cervellati 2004</td>
<td>20</td>
<td>Modified Luque technique (19) or Cotrel-Dubousset instrumentation (1)</td>
<td>Cobb angle, vital capacity, mortality.</td>
<td>Mean correction at follow-up was 28º. Mean loss of correction was 6º. Vital capacity showed a slow progression, slightly inferior to its natural evolution in untreated patients. Death in 1 patient</td>
<td></td>
</tr>
<tr>
<td>Chataigner 1998</td>
<td>27</td>
<td>Sublaminar wiring with Luque rods (5) or Hartshill rectangle (22) Sacral fixation with ilio-sacral screws linked to the rectangle by Cotrel-Dubousset rods and dominos (15)</td>
<td>Cobb angle, pelvic obliquity, coronal imbalance, sagittal imbalance, vital capacity, mortality, complications</td>
<td>Scoliosis reduced to 10º after surgery and 13º after 30 months' follow-up. Pelvic obliquity was reduced to 4º after surgery and 7º after 30 months. A good spinal balance was present in 20 pa-</td>
<td>Concluded that surgery did not result in respiratory improvement nor in life duration lengthening</td>
</tr>
</tbody>
</table>
Table 1. Characteristics of excluded studies  

<table>
<thead>
<tr>
<th>Study</th>
<th>n</th>
<th>Instrumentation</th>
<th>Outcome Measures</th>
<th>Results</th>
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</thead>
<tbody>
<tr>
<td>Dubouset 1983</td>
<td>37</td>
<td>Luque rods, Harrington rods, segmental instrumentation.</td>
<td>Cobb angle, vital capacity, mortality.</td>
<td>Scoliosis reduced from 80 to 24°. No effect on decline of vital capacity. No clear benefit in length of survival</td>
</tr>
<tr>
<td>Eagle 2007</td>
<td>75</td>
<td>Surgery and nocturnal ventilation (27), nocturnal ventilation only (13), no surgery or ventilation (35)</td>
<td>Survival, complications, FVC</td>
<td>No peri-operative deaths. Complications: GIB (2), postoperative ileus (1), spinal infection requiring removal of surgical rods (1), pressure sores (1), chronic pain due to prominence of metal prosthesis (2). Mean FVC reduced significantly (mean 1.4 L to 1.13 L) after 1 year. Median survival longer in surgery with ventilation group compared to ventilation alone (30 versus 22.2 years). Survival at 24 years higher in surgery with ventilation group compared to ventilation</td>
</tr>
</tbody>
</table>

Surgery for scoliosis in Duchenne muscular dystrophy (Review)  
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<table>
<thead>
<tr>
<th>Study</th>
<th>No.</th>
<th>Intervention</th>
<th>Outcomes</th>
<th>Findings</th>
</tr>
</thead>
<tbody>
<tr>
<td>Gaine 2004</td>
<td>74</td>
<td>Luque rod (55), Isola pedicle screw (19).</td>
<td>Cobb angle, pelvic obliquity, mortality, complications</td>
<td>Fusion to S1 did not offer benefit over fusion to more proximal level. Isola system appears to maintain a slightly better Cobb angle. 1 perioperative mortality due to cardiorespiratory failure. Complications: Failure of implants (3), wound infection (2), pseudarthrosis (2), metal implant prominence requiring removal (1)</td>
</tr>
<tr>
<td>Galasko 1992</td>
<td>55</td>
<td>Surgery (32), refused surgery (23).</td>
<td>Mortality, complications, FVC, PEFR, Cobb angle.</td>
<td>In surgery group, FVC static for 3 years then slightly decreased. Improved PEFR maintained for up to 5 years. Cobb angle improved from 47 to 34º at 5 years. Slightly improved survival with surgery. Complications: respiratory failure requiring tracheostomy (1), pneumonia (1), heart block (1), superficial wound infection (1)</td>
</tr>
<tr>
<td>Galasko 1995</td>
<td>76</td>
<td>Surgery (48), refused surgery (28).</td>
<td>Mortality, complications, FVC, PEFR, Cobb angle.</td>
<td>No pseudarthrosis or post-operative failures. Annual decrease of FVC lower in surgery group (0.07)</td>
</tr>
</tbody>
</table>

Patients with surgery have better lung function and improved survival.
Table 1. Characteristics of excluded studies  (Continued)

<table>
<thead>
<tr>
<th>Study</th>
<th>Method of Surgery</th>
<th>Vital Parameters</th>
<th>Complications</th>
<th>Other Notes</th>
</tr>
</thead>
<tbody>
<tr>
<td>Gayet 1999</td>
<td>Pedicular screwing system in the lumbo-sacral area and transversal attachments with steel threads at the thoracic level. A sub-laminar fastening was placed at L1</td>
<td>Vital capacity, mortality, complications, Cobb angle, pelvic obliquity</td>
<td>Cobb angle decreased from 19 to 5.2º, and 9.5% at the latest measurement. Pelvic balancing was corrected and results have held over time. Vital capacity was reduced by 3.6% per year. Complications: stem rupture (1), superficial infection (4)</td>
<td>Cardiorespiratory function and life expectancy were not improved, but most patients and families were very satisfied by the comfort brought about by the surgical operation</td>
</tr>
<tr>
<td>Granata 1996</td>
<td>Segmental spinal instrumentation and fusion.</td>
<td>Cobb angle, mortality, complications, vital capacity, quality of life, sitting position, aesthetic improvement</td>
<td>29 had a mean 59% correction of scoliosis. Very limited loss of correction over time. One died after cardiac arrest. Complications: pressure sore (1), metal prominence requiring trimming (1). Mean vital capacity decreased from 57 +/- 17% to 34 +/- 13% at 3.9 +/- 2 years after surgery</td>
<td></td>
</tr>
<tr>
<td>Study</td>
<td>Year</td>
<td>Study Design</td>
<td>Outcome Measures</td>
<td>Results</td>
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<tr>
<td>Hahn 2008</td>
<td>20</td>
<td>Spinal fixation with pedicle-screw-alone constructs</td>
<td>%FVC Cobb angle, degree of pelvic tilt, lumbar lordosis, and thoracic kyphosis, mortality, complications</td>
<td>Cobb angle improved from 44 to 10º, pelvic tilt improved from 14 to 3º. Lumbar lordosis improved from 20 to 49º, thoracic kyphosis remained unchanged. No problems related to iliac fixation, no pseudarthrosis or implant failures. No pulmonary complications. %FVC decreased from 55% preoperatively to 44% at the last follow-up. One patient died intraoperatively due to a sudden cardiac arrest. The rigid primary stability with pedicle screws allowed early mobilisation of the patients, which helped to avoid pulmonary complications.</td>
</tr>
<tr>
<td>Harper 2004</td>
<td>45</td>
<td>AO Universal Spinal System inserted through a posterior approach</td>
<td>Mortality, complications, hospital stay.</td>
<td>No significant difference in operative and postoperative outcomes between patients with pre-operative forced vital capacity ≥ 30% and ≤ 30%. Complications in 9 patients: pneumonia, respiratory failure requiring tracheostomy, ARDS, pleural effusion, cardiac arrhythmia. Concluded that routine postoperative use of mask ventilation to facilitate early tracheal extubation was vital.</td>
</tr>
<tr>
<td>Study</td>
<td>N</td>
<td>Method</td>
<td>Outcome Measures</td>
<td>Results</td>
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<tr>
<td>Hopf 1994</td>
<td>20</td>
<td>Multi-segmental instrumentation.</td>
<td>Mortality, complications, Cobb angle.</td>
<td>Mean Cobb angle decreased from 70.6 to 31.2° (mean correction 39.4° or 55.8%). Lordosis of the lumbar spine corrected from 4.1 to 17.8°. No peri-operative mortality. Complication: bladder dysfunction in 1 patient</td>
</tr>
<tr>
<td>Kennedy 1995</td>
<td>38</td>
<td>Surgery (17), no surgery (21).</td>
<td>Cobb angle, forced vital capacity (FVC), mortality.</td>
<td>Mean Cobb angle of the surgical group at 14.9 years was 57 +/- 16.4°, and of the non-surgical group at 15 years was 45 +/- 9.9°. No difference in the rate of deterioration of % FVC which was 3 to 5% per year. No difference in survival in either group</td>
</tr>
<tr>
<td>Study</td>
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<td>Procedure</td>
<td>Outcome Measures</td>
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<tr>
<td>Kinali 2006</td>
<td>123</td>
<td>Surgery (43), no surgery (80)</td>
<td>Survival, FVC, sitting comfort</td>
<td>No difference in survival, respiratory impairment, or sitting comfort among patients managed conservatively or with surgery</td>
</tr>
<tr>
<td>Laprade 1992</td>
<td>9</td>
<td>Sublaminar wiring (4), intraspinous segmental wiring (5)</td>
<td>Mortality, complications, operative time, blood loss, Cobb angle</td>
<td>Operative time and blood loss lower in sublaminar compared to intraspinous wiring. Allogeneic bone grafts to supplement the autogenous bone graft allowed for extensive fusion. Cobb angle decreased by a mean of 32º. Complications: dural leak (1), transient numbness of left foot (1), dislodgement of sacral alar hooks (2)</td>
</tr>
<tr>
<td>Marchesi 1997</td>
<td>25</td>
<td>Modified Luque: sacral screws in each S-1 pedicle and a device for transverse traction between the caudal right-angle bends of the L-rods</td>
<td>Cobb angle, pelvic obliquity, mortality, instrumental failure, sitting balance</td>
<td>Cobb angle decreased from 68 to 18º and pelvic obliquity decreased from 21 to &lt;15º with mean correction of 75%. No instrumentation failure or loss of correction &gt;3º. In every patient, a good sitting balance could be restored. No perioperative mortality</td>
</tr>
</tbody>
</table>
| Marsh 2003    | 30 | Posterior spinal fusion                               | Cobb angle, mortality, complications, hospital stay | Mean correction of Cobb angle 36º. Two subgroups of patients were compared: those with surgery | Concluded that spinal fusion could be offered to patients with DMD even in the presence of a low
Table 1. Characteristics of excluded studies (Continued)

<table>
<thead>
<tr>
<th>Study</th>
<th>Patients</th>
<th>Procedures</th>
<th>Outcomes</th>
<th>Results</th>
</tr>
</thead>
<tbody>
<tr>
<td>Matsumura 1997</td>
<td>8</td>
<td>Luque rod (2), Cotrel-Dubousset rod (6)</td>
<td>Cobb angle corrected from 58.8 to 28.6º with the mean corrective rate of 51.3%. FVC increased in 3 patients with moderate scoliosis (Cobb angle: 50 to 80º). Two cases with low % FVC (16.9% and 30.4%, respectively) had poor prognosis in respiratory status. One died of pneumonia at 17 months after the surgery and the other required mechanical ventilation. Sitting balance improved in all patients.</td>
<td>Recommended spinal fusion for patients with Cobb angle more than 30º and with % FVC more than 35%. Although the impact of spinal fusion upon the life expectancy remained unclear, favorable effect on respiratory function and quality of life could be expected for carefully selected patients with DMD</td>
</tr>
</tbody>
</table>

FVC was used as an indicator of respiratory function: more than 30% pre-operative FVC (17 patients) and those with less than 30% pre-operative FVC (13 patients). One patient in each group required a temporary tracheotomy and there were nine complications in total. The post-operative stay for patients in each group was similar (24 days in the >30% group, 20 days in the <30% group) and the complication rate was comparable with other published series. No peri-operative mortality.
Table 1. Characteristics of excluded studies  (Continued)

<table>
<thead>
<tr>
<th>Study</th>
<th>No.</th>
<th>Intervention</th>
<th>Outcome measures</th>
</tr>
</thead>
<tbody>
<tr>
<td>Mehdian 1989</td>
<td>17</td>
<td>Luque rods secured by conventional sub-laminar wires (9), Luque rods secured by sublaminar nylon straps (4), 2 L-shaped rods connected by H-bars secured by closed wire loops (3), Hartshill rectangle and sub-laminar wires (1)</td>
<td>Cobb angle, respiratory function. Significant loss of correction in Luque rods secured by sublaminar nylon straps and Hartshill system. Strong correlation between advance of scoliosis and respiratory function</td>
</tr>
<tr>
<td>Miller 1988</td>
<td>67</td>
<td>Surgery (21), no surgery (46).</td>
<td>FVC.</td>
</tr>
<tr>
<td>Miller 1991</td>
<td>39</td>
<td>Surgery (17), no surgery (22).</td>
<td>Respiratory function, sitting comfort, sitting appearance.</td>
</tr>
<tr>
<td>Miller 1992</td>
<td>183</td>
<td>Surgery (68), no surgery (115).</td>
<td>Survival, patient comfort, ease of care, respiratory function, quality of life</td>
</tr>
</tbody>
</table>

Surgery for scoliosis in Duchenne muscular dystrophy (Review)
Table 1. Characteristics of excluded studies  

<table>
<thead>
<tr>
<th>Study</th>
<th>Patient Characteristics</th>
<th>Instrumentation</th>
<th>Outcome Measures</th>
<th>Results</th>
</tr>
</thead>
<tbody>
<tr>
<td>Modi 2008a</td>
<td>26 (including 7 cerebral palsy, 5 SMA, 4 others)</td>
<td>Posterior pelvic screw fixation</td>
<td>Cobb angle, pelvic obliquity, complications</td>
<td>Mean Cobb angle: 78.53º (before surgery), 30.7º (after surgery), 33.06º (final follow-up). There was no difference in the percentage correction between the groups with &gt;90º or &lt;90º. Complications: 1 transient loss of lower limb power, 1 deep wound infection</td>
</tr>
<tr>
<td>Modi 2008b</td>
<td>24 patients (including 6 cerebral palsy, 5 SMA, 4 others) and 12 controls (adolescent idiopathic scoliosis)</td>
<td>Posteriod pedicle screw</td>
<td>Cobb angle, pelvic obliquity, apical rotation</td>
<td>Mean Cobb angle decreased from 74 to 32º. Mean pelvic obliquity decreased from 14 to 6º. Mean apical rotation decreased from 42 to 33º. There was no significant difference between different patient groups or between patients and controls</td>
</tr>
<tr>
<td>Modi 2009</td>
<td>50 (including 18 patients with cerebral palsy, 8 patients with SMA and 6 others)</td>
<td>Posterior spinal fusion with segmental spinal instrumentation using pedicle screw fixation</td>
<td>Mortality, complications, Cobb angle, pelvic obliquity</td>
<td>Cobb angle decreased from 79.3+/−30.3º to 31.3+/−21.6º. Pelvic obliquity decreased from 14.6+/−9.4º to 6.8+/−6.3º. 2 deaths (1 due to cardiac arrest, 1 due to hypovolemic shock. 34 patients had at least 1 periop-DMD patients had higher risk of post-operative coccygodynia.</td>
</tr>
</tbody>
</table>
Table 1. Characteristics of excluded studies  

<table>
<thead>
<tr>
<th>Study</th>
<th>Patients (inclusion criteria)</th>
<th>Spinal fixation</th>
<th>Cobb angle, pelvic obliquity, complications</th>
<th>Outcomes</th>
<th>Conclusions</th>
</tr>
</thead>
<tbody>
<tr>
<td>Modi 2010</td>
<td>55 (including 28 patients with cerebral palsy and 10 patients with SMA)</td>
<td>Spinal fixation from T2/T3/T4 to L4/L5 with or without pelvic fixation. Group 1: pelvic obliquity &gt;15° with pelvic fixation; group 2: pelvic obliquity &gt;15° without pelvic fixation; group 3: pelvic obliquity &lt;15° without pelvic fixation</td>
<td>Mean correction of Cobb angle after operation: group 1: 43.8°; group 2: 40°; group 3: 48.7°. Mean loss of correction of Cobb angle at last follow-up: group 1: 0.6°; group 2: 2.3°; group 3: 3°. Mean correction of pelvic obliquity: group 1: 14.4°; group 2: 10.7°; group 3: 5°. Mean loss of correction of pelvic obliquity at last follow-up: group 1: -0.6°; group 2: 6.5°; group 3: 0.8°. Group 2 showed significant loss of pelvic obliquity compared to group 1. Complications: 3 patients had sacral sores in group 1</td>
<td>Outcomes similar between the 2 groups.</td>
<td>Patients who have pelvic obliquity &gt;15 degrees require pelvic fixation to maintain correction</td>
</tr>
</tbody>
</table>
| Mubarak 1993  | 22                            | Luque segmental instrumentation and fusion instrumented to the sacropelvis (12), instrumented to L5 (10) | Cobb angle, pelvic obliquity. | Concluded that if treatment is initiated early, Luque instrumentation and fusion from high thoracic (T2 or T3) to the fifth lumbar vertebra should be suf-
Table 1. Characteristics of excluded studies  

<table>
<thead>
<tr>
<th>Study</th>
<th>N</th>
<th>Intervention</th>
<th>Outcomes</th>
<th>Conclusion</th>
</tr>
</thead>
<tbody>
<tr>
<td>Nakazawa 2010</td>
<td>36</td>
<td>Autogenous bone graft (20), allogeneic bone graft (16)</td>
<td>Cobb angle, operating time, blood loss</td>
<td>No difference in Cobb angle between the 2 groups. Mean operating time longer in autogenous group (253 min) compared to allogeneic group (233 min). Mean blood loss higher in autogenous group (850 ml) compared to allogeneic group (775 ml) 90% and 50% of patients in autogenous group reported donor site pain after 1 week and 3 months respectively. Concluded against autogenous bone graft for scoliosis surgery in DMD patients</td>
</tr>
<tr>
<td>Rice 1998</td>
<td>19</td>
<td>Long spinal fusion to L5 and ongoing wheelchair seating attention</td>
<td>Sitting position</td>
<td>At long-term follow-up 15 patients continued to sit in a well-balanced position</td>
</tr>
<tr>
<td>Rideau 1984</td>
<td>5</td>
<td>Luque segmental spinal stabilization without bone fusion</td>
<td>Cobb angle, vital capacity, mortality, complications, hospital stay, pelvic obliquity, patient comfort</td>
<td>Cobb angle decreased from 27 to 11º. Pelvic obliquity partially reduced. Static vital capacity after 2 years. No peri-operative mortality. 1 bronchopneumonia. All patients more comfortable during wheelchair activities</td>
</tr>
<tr>
<td>Sakai 1977</td>
<td>41</td>
<td>Surgery (10), no surgery (31)</td>
<td>Sitting stability, mortality, complications</td>
<td>Pulmonary complications were minimized by performing preoperative tracheostomy on all patients who had vital capacities less than 40% and or non-func-</td>
</tr>
<tr>
<td>Study</td>
<td>Sample Size</td>
<td>Technique</td>
<td>Measurements</td>
<td>Description</td>
</tr>
<tr>
<td>---------------</td>
<td>-------------</td>
<td>----------------------------------------------</td>
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<td>-------------</td>
</tr>
<tr>
<td>Sengupta 2002</td>
<td>50</td>
<td>Galveston technique (9), L-rod (22), pedicle screw + sub-laminar wires (19)</td>
<td>Cobb angle, pelvic obliquity, mortality, complications, hospital stay</td>
<td>In the pelvic fixation group, the mean Cobb angle and pelvic obliquity were 48° and 19.8° at the time of surgery, 16.7° and 7.2° immediately after surgery, and 22° and 11.6° at the final follow-up (mean 4.6 years). The mean hospital stay was 17 days. 5 major complications: deep wound infection (1), revision of instrumentation prominence at the proximal end (2), loosening of pelvic fixation (2). In the lumbar fixation group, the mean Cobb angle and pelvic obliquity were 19.8° and 9° at the time of surgery, 3.2° and 2.2° immediately after surgery, and 5.2° and 2.9° at the final follow-up (mean 3.5 years). The mean hospital stay (7.7 days) was much less compared with the pelvic fixation group. Pelvic obliquity was corrected and main-</td>
</tr>
<tr>
<td>Study</td>
<td>Patients</td>
<td>Procedure</td>
<td>Cobb Angle</td>
<td>Complications</td>
</tr>
<tr>
<td>------------------</td>
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<td>---------------</td>
</tr>
<tr>
<td>Shapiro 1992</td>
<td>27</td>
<td>Harrington rod (2), Harrington rod with sublaminar wires (7), Harrington rod, Luque rod and 2 double sublaminar wires at each level (17)</td>
<td>Cobb angle, FVC, mortality, complications.</td>
<td>1 sudden cardiac arrest and died intraoperatively. 3 intraoperative complications reversed without sequelae. Mean post-operative correction 13.1 +/- 11.9º, with mean loss of correction 5.1 +/- 3.1º at 2.4 +/- 1.8 years. Mean FVC preoperatively was 45.3 +/- 15.9% with continuing diminution to 28.7 +/- 14.9% at 3.3 +/- 2.2 years after surgery</td>
</tr>
<tr>
<td>Stricker 1996</td>
<td>46 (included other neuromuscular diseases)</td>
<td>Modified Luque technique.</td>
<td>Cobb angle, complications.</td>
<td>Cobb angle decreased from 63 to 24º (correction of about 62%). Failure of implants, pseudarthroses and major losses of correction in purely neuromuscular scolioses could be avoided by using rigid segmental fixation and a dorsolateral fusion with a mixture of autologous and allogeneous bone</td>
</tr>
<tr>
<td>Study</td>
<td>Patients</td>
<td>Description</td>
<td>Complications</td>
<td>Cobb Angle Correction</td>
</tr>
<tr>
<td>-----------------</td>
<td>----------</td>
<td>-------------------------------------------------------------------------------</td>
<td>---------------</td>
<td>------------------------</td>
</tr>
<tr>
<td>Sussman 1984</td>
<td>11</td>
<td>Harrington instrumentation (group I) (3), Luque instrumentation (group II) (3), segmental spinal instrumentation with fusion (group III) (5)</td>
<td>Complications, Cobb angle, hospital stay.</td>
<td>Mean Cobb angle correction: 40% (I), 35% (II), 60% (III). When surgery to stabilize spinal deformity is done in younger patients in whom pulmonary function is better and curves are milder, complication rate and length of hospital stay are diminished, correction and balance are improved, and patients rapidly return to their normal lifestyle. Concluded that segmental spinal instrumentation had advantage of allowing rapid mobilization without need of a cast or body jacket. Recommended stabilization of the collapsing spine surgically with segmental instrumentation and fusion when scoliosis reached 30 to 40º.</td>
</tr>
<tr>
<td>Takas 2010</td>
<td>20</td>
<td>Segmental pedicle screws instrumentation and fusion to L5.</td>
<td>Complications</td>
<td>Mean Cobb angle decreased from 70º to 15º. Mean pelvic obliquity decreased from 13º to 6º. The mean intraoperative blood loss was 890 ml (range: 660 to 1260 ml). The mean total blood loss was 2100 ml (range: 1250 to 2880 ml). There was no major complication.</td>
</tr>
<tr>
<td>Thacker 2002</td>
<td>5</td>
<td>Not detailed in DMD patients.</td>
<td>FEV1, FVC, mortality, complications.</td>
<td>FVC and FEV1 maintained, pseudarthrosis in 1 patient, no peri-operative mortality. Included 7 SMA, 6 spastic cerebral palsy, 3 congenital myopathy, 2 spina bifida, 1 paraspinal neuroblastoma in the series.</td>
</tr>
</tbody>
</table>
| Velasco 2007    | 56       | Posterior spinal fusion                                                       | Percent normal FVC | The rates of FVC decline were 4% per year presurgery, which decreased to 1.75% per year post-

Surgery for scoliosis in Duchenne muscular dystrophy (Review)  
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Table 1. Characteristics of excluded studies (Continued)

<table>
<thead>
<tr>
<th>Study</th>
<th>Patients</th>
<th>Description</th>
<th>Mortality, complications</th>
<th>Conclusions</th>
</tr>
</thead>
<tbody>
<tr>
<td>Weimann 1983</td>
<td>24</td>
<td>Long Harrington instrumentations and spinal fusions from S1 up to the upper thoracic spine (T4, 5, or 6)</td>
<td>One patient died 2 years after his operation from dystrophic cardiomyopathy</td>
<td>Concluded that prophylactic spinal fusion deserved consideration in the care planned for these patients</td>
</tr>
</tbody>
</table>

ARDS: adult respiratory distress syndrome; DMD: Duchenne muscular dystrophy; FVC: forced vital capacity; FEV1: forced expiratory volume in 1 second; PEFR: peak expiratory flow rate; SMA: spinal muscular atrophy

APPENDICES

Appendix 1. MEDLINE strategy

Database: Ovid MEDLINE(R) <1946 to July Week 3 2012>

Search Strategy:

1 randomized controlled trial.pt. (332315)
2 controlled clinical trial.pt. (84684)
3 randomized.ab. (235702)
4 placebo.ab. (133040)
5 drug therapy.fs. (1552464)
6 randomly.ab. (169810)
7 trial.ab. (244167)
8 groups.ab. (1114025)
9 or/1-8 (2885687)
10 exp animals/ not humans.sh. (3757814)
11 9 not 10 (2450652)
12 surg$.mp. or surgery/ (1335297)
13 spine$.mp. (82686)
14 spinal.mp. (269459)
15 vertebra$.mp. (166510)
16 or/13-15 (412217)
17 12 and 16 (56524)
18 spinal fusion/ or spinal fusion.mp. (15911)
19 17 or 18 (62847)
20 scoliosis.mp. or Scoliosis/ (15574)
21 duchenne.mp. or Muscular Dystrophy, Duchenne/ (7902)
22 11 and 19 and 20 and 21 (18)
23 remove duplicates from 22 (17)
Appendix 2. EMBASE search strategy

Database: Embase <1980 to 2012 Week 30>
Search Strategy:

1 crossover-procedure.sh. (34521)
2 double-blind procedure.sh. (109963)
3 single-blind procedure.sh. (16165)
4 randomized controlled trial.sh. (326003)
5 (random$ or crossover$ or cross over$ or placebo$ or (doubl$ adj blind$) or allocat$).tw,ot. (885002)
6 trial.ti. (133129)
7 clinical trial/ (869205)
8 or/1-7 (1482353)
9 (animal/ or nonhuman/ or animal experiment/) and human/ (1194751)
10 animal/ or nonanimal/ or animal experiment/ (3291877)
11 10 not 9 (2727149)
12 8 not 11 (1395248)
13 limit 12 to embase (1081020)
14 Surgery/ or surg$.mp. (1965351)
15 (spine or spinal or vertebra$).mp. (474719)
16 14 and 15 (86443)
17 exp Spine Fusion/ (16226)
18 (spinal fusion or spine fusion).mp. (16755)
19 16 or 17 or 18 (92142)
20 exp Scoliosis/ or scoliosis.mp. (20307)
21 Duchenne Muscular Dystrophy/ or duchenne.mp. (11023)
22 13 and 19 and 20 and 21 (11)

Appendix 3. CENTRAL search strategy

#1 MeSH descriptor General Surgery explode all trees
#2 surgery
#3 (#1 OR #2)
#4 (spine or spinal or vertebra*)
#5 (#3 AND #4)
#6 MeSH descriptor Spinal Fusion, this term only
#7 spinal fusion or spine fusion
#8 (( #5 AND #6 ) OR #7)
#9 scoliosis
#10 duchenne
#11(#8 AND #9 AND #10)

Appendix 4. CINAHL search strategy

Tuesday, July 31, 2012 11:29:22 AM

S29 S18 and S28 13
S28 S25 and S26 and S27 35
S27 ("scoliosis") or (MH "Scoliosis") 3652
S26 ("duchenne") or (MH "Duchenne Muscular Dystrophy") 793
S25 S22 or S24 13207
S24 S23 or spinal fusion or spine fusion 3727
S23 (MH "Spinal Fusion") 3397
S22 S20 and S21 12713
Appendix 5. Proquest Dissertation & Thesis Database search strategy

Duchenne and surgery and scoliosis

Appendix 6. NIH Clinical Trials Database

Duchenne and surgery and scoliosis

WHAT'S NEW

Last assessed as up-to-date: 31 July 2012.

<table>
<thead>
<tr>
<th>Date</th>
<th>Event</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>4 January 2013</td>
<td>New citation required but conclusions have not changed</td>
<td>Review updated with search update to July 31 2012 but no new studies found. Two of the original authors withdrawn</td>
</tr>
<tr>
<td>7 November 2012</td>
<td>New search has been performed</td>
<td>Two studies added to excluded studies tables. Minor editorial revisions</td>
</tr>
</tbody>
</table>
HISTORY

Protocol first published: Issue 3, 2005
Review first published: Issue 1, 2007

<table>
<thead>
<tr>
<th>Date</th>
<th>Event</th>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>22 August 2010</td>
<td>New search has been performed</td>
<td>Review updated with search update but no new studies found</td>
</tr>
<tr>
<td>13 May 2009</td>
<td>Amended</td>
<td>Acknowledgement added.</td>
</tr>
<tr>
<td>2 October 2008</td>
<td>New search has been performed</td>
<td>updated review</td>
</tr>
<tr>
<td>23 October 2006</td>
<td>New citation required and conclusions have changed</td>
<td>Substantive amendment</td>
</tr>
</tbody>
</table>

CONTRIBUTIONS OF AUTHORS

Cheuk DKL: protocol development, searching for trials, quality assessment of trials, data extraction, data input, data analyses, development of final review, corresponding author.

Wong V: protocol development, searching for trials, quality assessment of trials, data extraction, data analyses, development of final review.

Wraige E: protocol development, searching for trials, quality assessment of trials, data extraction, data analyses, development of final review.

Baxter P: protocol development, searching for trials, quality assessment of trials, data extraction, data analyses, development of final review.

Cole A: protocol development, searching for trials, quality assessment of trials, data extraction, data analyses, development of final review.

DECLARATIONS OF INTEREST

No potential conflict of interest is known.

SOURCES OF SUPPORT
Internal sources
  • None, Not specified.

External sources
  • None, Not specified.

DIFFERENCES BETWEEN PROTOCOL AND REVIEW
Risk of bias methodology updated in accordance with the Cochrane Handbook for Systematic Reviews of Interventions (Higgins 2011). Change in authorship: we were unable to contact original authors N’Diaye T and Mayowe V for this update.

INDEX TERMS

Medical Subject Headings (MeSH)
Muscular Dystrophy, Duchenne [*complications]; Scoliosis [complications; *surgery]; Spine [surgery]

MeSH check words
Humans