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# Triage tools for detecting cervical spine injury in paediatric trauma patients (Review)

Tavender E, Eapen N, Wang J, Rausa VC, Babl FE, Phillips N

Tavender E, Eapen N, Wang J, Rausa VC, Babl FE, Phillips N. Triage tools for detecting cervical spine injury in paediatric trauma patients. *Cochrane Database of Systematic Reviews* 2024, Issue 3. Art. No.: CD011686. DOI: 10.1002/14651858.CD011686.pub3.

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# [Diagnostic Test Accuracy Review]

# Triage tools for detecting cervical spine injury in paediatric trauma patients

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**Editorial group:** Cochrane Back and Neck Group. **Publication status and date:** New search for studies and content updated (conclusions changed), published in Issue 3, 2024.

**Citation:** Tavender E, Eapen N, Wang J, Rausa VC, Babl FE, Phillips N. Triage tools for detecting cervical spine injury in paediatric trauma patients. *Cochrane Database of Systematic Reviews* 2024, Issue 3. Art. No.: CD011686. DOI: 10.1002/14651858.CD011686.pub3.

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# ABSTRACT

### Background

Paediatric cervical spine injury (CSI) after blunt trauma is rare but can have severe consequences. Clinical decision rules (CDRs) have been developed to guide clinical decision-making, minimise unnecessary tests and associated risks, whilst detecting all significant CSIs. Several validated CDRs are used to guide imaging decision-making in adults following blunt trauma and clinical criteria have been proposed as possible paediatric-specific CDRs. Little information is known about their accuracy.

### Objectives

To assess and compare the diagnostic accuracy of CDRs or sets of clinical criteria, alone or in comparison with each other, for the evaluation of CSI following blunt trauma in children.

### Search methods

For this update, we searched CENTRAL, MEDLINE, Embase, and six other databases from 1 January 2015 to 13 December 2022. As we expanded the index test eligibility for this review update, we searched the excluded studies from the previous version of the review for eligibility. We contacted field experts to identify ongoing studies and studies potentially missed by the search. There were no language restrictions.

### **Selection criteria**

We included cross-sectional or cohort designs (retrospective and prospective) and randomised controlled trials that compared the diagnostic accuracy of any CDR or clinical criteria compared with a reference standard for the evaluation of paediatric CSI following blunt trauma. We included studies evaluating one CDR or comparing two or more CDRs (directly and indirectly). We considered X-ray, computed tomography (CT) or magnetic resonance imaging (MRI) of the cervical spine, and clinical clearance/follow-up as adequate reference standards.

### Data collection and analysis

Two review authors independently screened titles and abstracts for relevance, and carried out eligibility, data extraction and quality assessment. A third review author arbitrated. We extracted data on study design, participant characteristics, inclusion/exclusion criteria, index test, target condition, reference standard and data (diagnostic two-by-two tables) and calculated and plotted sensitivity and



specificity on forest plots for visual examination of variation in test accuracy. We assessed methodological quality using the Quality Assessment of Diagnostic Accuracy Studies Version 2 tool. We graded the certainty of the evidence using the GRADE approach.

### **Main results**

We included five studies with 21,379 enrolled participants, published between 2001 and 2021. Prevalence of CSI ranged from 0.5% to 1.85%. Seven CDRs were evaluated.

Three studies reported on direct comparisons of CDRs. One study (973 participants) directly compared the accuracy of three index tests with the sensitivities of NEXUS, Canadian C-Spine Rule and the PECARN retrospective criteria being 1.00 (95% confidence interval (CI) 0.48 to 1.00), 1.00 (95% CI 0.48 to 1.00) and 1.00 (95% CI 0.48 to 1.00), respectively. The specificities were 0.56 (95% CI 0.53 to 0.59), 0.52 (95% CI 0.49 to 0.55) and 0.32 (95% CI 0.29 to 0.35), respectively (moderate-certainty evidence). One study (4091 participants) compared the accuracy of the PECARN retrospective criteria with the Leonard de novo model; the sensitivities were 0.91 (95% CI 0.81 to 0.96) and 0.92 (95% CI 0.83 to 0.97), respectively. The specificities were 0.46 (95% CI 0.44 to 0.47) and 0.50 (95% CI 0.49 to 0.52) (moderate- and low-certainty evidence, respectively). One study (270 participants) compared the accuracy of two NICE (National Institute for Health and Care Excellence) head injury guidelines; the sensitivity of the CG56 guideline was 1.00 (95% CI 0.48 to 1.00) compared to 1.00 (95% CI 0.48 to 1.00) with the CG176 guideline. The specificities were 0.46 (95% CI 0.40 to 0.52) and 0.07 (95% CI 0.04 to 0.11), respectively (very low-certainty evidence).

Two additional studies were indirect comparison studies. One study (3065 participants) tested the accuracy of the NEXUS criteria; the sensitivity was 1.00 (95% CI 0.88 to 1.00) and specificity was 0.20 (95% CI 0.18 to 0.21) (low-certainty evidence). One retrospective study (12,537 participants) evaluated the PEDSPINE criteria and found a sensitivity of 0.93 (95% CI 0.78 to 0.99) and specificity of 0.70 (95% CI 0.69 to 0.72) (very low-certainty evidence).

We did not pool data within the broader CDR categories or investigate heterogeneity due to the small quantity of data and the clinical heterogeneity of studies. Two studies were at high risk of bias.

We identified two studies that are awaiting classification pending further information and two ongoing studies.

### **Authors' conclusions**

There is insufficient evidence to determine the diagnostic test accuracy of CDRs to detect CSIs in children following blunt trauma, particularly for children under eight years of age. Although most studies had a high sensitivity, this was often achieved at the expense of low specificity and should be interpreted with caution due to a small number of CSIs and wide CIs. Well-designed, large studies are required to evaluate the accuracy of CDRs for the cervical spine clearance in children following blunt trauma, ideally in direct comparison with each other.

### PLAIN LANGUAGE SUMMARY

### Clinical tools for detecting cervical spine injury (CSI) in children with injuries

### Key message

- There is currently insufficient evidence to determine which clinical decision tools should be used to assist in deciding whether children with potential cervical spine injuries (CSI) require imaging tests to aid diagnosis.

### What is a cervical spine injury and how is it detected?

The cervical spine is the upper part of the spine between the head and shoulders (the neck). The incidence of traumatic CSI in children is very low. However, it is very important not to miss this injury as the consequences can be devastating, including death or lifelong disability. To detect CSI, several types of imaging techniques can be used: computed tomography (CT), magnetic resonance imaging (MRI) and X-rays. A CT scan uses detailed X-rays to produce cross-sectional images of the body and MRI uses radio waves and a powerful magnet to generate the images. While CT scans and X-rays are useful in detecting bone injuries, they do use radiation that can increase the risk of developing cancer, especially in children. To avoid exposing children to unnecessary radiation, it is important to find clinical tests that can determine whether children are at risk for CSI, how accurate they are (called diagnostic accuracy) and whether radiographic imaging is needed.

#### What was the aim of this review?

Clinical decision rules (CDRs) are tools that clinicians use to decide whether a diagnostic test is needed or another clinical action should be taken. We wanted to find out which CDRs are useful in determining which children are at risk for CSI after blunt trauma (for example, in motor vehicle-related accidents and falls), and whether radiographic imaging should be used to help diagnosis. Tools that have been developed for adults are also often used for children, but little information is known about their accuracy in children. The aim of this review was to evaluate all CDRs and tools used in this decision-making process and if they can be used safely and effectively in children.

#### What did we do?



We searched medical databases for studies that compared the diagnostic accuracy of any CDR with another CDR for the evaluation of CSI following blunt trauma in children.

#### What did we find?

We included five studies recruiting 21,379 children, published between 2001 and 2021, that assessed the accuracy of seven CDRs (NEXUS, Canadian C-Spine Rule, PECARN retrospective criteria, NICE guidelines CG56 and CG176, Leonard de novo model and PEDSPINE) to evaluate CSIs following blunt trauma in children.

#### Main results

There is currently insufficient evidence to determine which CDRs are most effective at detecting CSIs following blunt trauma in children, particularly for those younger than eight years of age. Although most CDRs accurately identified children who had a CSI (called high sensitivity), they frequently did not correctly identify children who did not have a CSI (called low specificity). If these CDRs were applied as a rule, a large proportion of children without CSI attending the emergency department for a blunt trauma assessment would receive imaging potentially exposing them to unnecessary radiation. These CDRs are at best a guide to clinical assessment with current evidence not supporting strict use of CDRs in trauma care for children. More research is needed to evaluate the accuracy of CDRs for use in cervical spine assessment in children.

# What are the limitations of the evidence?

The quality of the studies was variable as there were differences in the children recruited, the number of CSIs, and the methods used making us uncertain about the results. There are currently two large ongoing studies that should contribute to the evidence of the accuracy of CDRs in children.

#### How up to date is the evidence?

The evidence is up to date to 13 December 2022.

# SUMMARY OF FINDINGS

# Summary of findings 1. Direct comparisons of clinical decision rules (CDRs) or sets of clinical criteria

Question: what is the diagnostic accuracy of clinical decision rules or sets of clinical criteria used to evaluate for cervical spine injury in the emergency department following blunt trauma in children?

Population: children (aged 0 to < 18 years) who underwent blunt trauma evaluation in the emergency department

**Index test:** CDRs or sets of clinical criteria that compared the diagnostic accuracy of the test for cervical spine injury with the reference standard

**Comparator:** studies comparing ≥ 2 CDRs (directly)

Reference standard: X-ray, CT, MRI or clinical clearance/follow-up in low-risk children

Study types: diagnostic studies with cross-sectional or cohort designs (retrospective or prospective) and randomised controlled trials

Study	Participants (CSI, %)	Sensitivity (95% CI)	Specificity (95% CI)	Certainty of the evidence	
Phillips 2021 973 (0.5%)		NEXUS: 1.00 (0.48 to 1.00)	NEXUS: 0.56 (0.53 to 0.59)	<b>Moderate</b> <sup>a</sup>	
	(0.070)	Canadian C-Spine Rule: 1.00 (0.48 to 1.00)	Canadian C-Spine Rule: 0.52 (0.49 to 0.55)	_	
		PECARN retrospective: 1.00 (0.48 to 1.00)	PECARN retrospective: 0.32 (0.29 to 0.35)	-	
Leonard 2019	4091 (1.8%)	PECARN retrospective: 0.91 (0.81 to 0.96)	PECARN retrospective: 0.46 (0.44 to 0.47)	Moderate <sup>a</sup>	
		Leonard de novo: 0.92 (0.83 to 0.97)	Leonard de novo: 0.50 (0.49 to 0.52)	Low <sup>b</sup>	
Davies 2016	270 (1.85%)	NICE CG56: 1.00 (0.48 to 1.00)	NICE CG56: 0.46 (0.40 to 0.52)	Very low <sup>c</sup>	
(1.6570)		NICE CG176: 1.00 (0.48 to 1.00)	NICE CG176: 0.07 (0.04 to 0.11)	-	

**CDR:** clinical decision rule; **CI:** confidence interval; **CSI:** cervical spine injury; **CT:** computed tomography; **MRI:** magnetic resonance imaging; **NEXUS:** National Emergency X-Radiography Utilization Study; **NICE:** National Institute for Health and Care Excellence; **PECARN:** Pediatric Emergency Care Applied Research Network.

#### **GRADE Working Group grades of evidence**

High certainty: we are very confident that the true effect lies close to that of the estimate of the effect.

**Moderate certainty:** we are moderately confident in the effect estimate; the true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different.

Low certainty: our confidence in the effect estimate is limited; the true effect may be substantially different from the estimate of the effect.

Very low certainty: we have very little confidence in the effect estimate; the true effect is likely to be substantially different from the estimate of effect.

<sup>a</sup> Downgraded one level due to study limitations (unclear risk of bias for the index test and reference standard).

<sup>b</sup> Downgraded one level due to study limitations (unclear risk of bias for the index test and reference standard) and one level due to indirectness (no existing validation data).

<sup>c</sup> Downgraded two levels due to study limitations (retrospective study design) and one level due to indirectness (only children with cervical spine imaging were included).

# Summary of findings 2. Indirect comparisons of clinical decision rules (CDRs) or sets of clinical criteria

Question: what is the diagnostic accuracy of clinical decision rules or sets of clinical criteria used to evaluate for cervical spine injury in the emergency department following blunt trauma in children?

Population: children (aged 0 to < 18 years) who underwent blunt trauma evaluation in the emergency department

Index test: CDRs that compared the diagnostic accuracy of the test for cervical spine injury with the reference standard

Comparator: studies evaluating 1 single CDR or ≥ 2 CDRs (indirectly)

Reference standard: X-ray, CT, MRI or clinical clearance/follow-up in low-risk children

Study types: diagnostic studies with cross-sectional or cohort designs (retrospective or prospective) and randomised controlled trials

Study	Participants (CSI, %)	Sensitivity (95% CI)	Specificity (95% CI)	Certainty of the Evidence	
NEXUS 2 studies		Phillips: 1.00 (0.48 to 1.00)	Phillips: 0.56 (0.53 to 0.59)	Low <sup>a</sup>	
	4038 (0.5% and 0.98%)	Viccellio: 1.00 (0.88 to 1.00)	Viccellio: 0.20 (0.18 to 0.21)	-	
Canadian C-	1 study	Phillips: 1.00 (0.48 to 1.00)	Phillips: 0.52 (0.49 to 0.55)	Moderate <sup>b</sup>	
Spine Rule	973 (0.5%)				
PECARN retro-	2 studies	Phillips: 1.00 (0.48 to 1.00)	Phillips: 0.32 (0.29 to 0.35)	Moderate <sup>b</sup>	
spective	5064 (0.5% and 1.8%)	Leonard: 0.91 (0.81 to 0.96)	Leonard: 0.46 (0.44 to 0.47)	-	
Leonard de no-	1 study	Leonard: 0.92 (0.83 to 0.97)	Leonard: 0.50 (0.49 to 0.52)	Low <sup>c</sup>	
vo	4091 (1.8%)				
PEDSPINE crite- ria	1 study	Pierretti: 0.93 (0.78 to 0.99)	Pierretti: 0.70 (0.69 to 0.72)	Very low <sup>d</sup>	
	12,537 (0.66%)				
NICE CG56	1 study	NICE CG56: 1.00 (95% CI 0.48 to	NICE CG56: 0.46 (95% CI 0.40 to	Very low <sup>e</sup>	
	270 (1.85%)	1.00)	0.52)		
NICE CG176	1 study	NICE CG176: 1.00 (95% CI 0.48 to	NICE CG176: 0.07 (95% CI 0.04 to	Very low <sup>e</sup>	
	270 (1.85%)	1.00)	0.11)		

**CDR:** clinical decision rule; **CI:** confidence interval; **CSI:** cervical spine injury; **CT:** computed tomography; **MRI:** magnetic resonance imaging; **NEXUS:** National Emergency X-Radiography Utilization Study; **NICE:** National Institute for Health and Care Excellence; **PECARN:** Pediatric Emergency Care Applied Research Network.

#### **GRADE Working Group grades of evidence**

High certainty: we are very confident that the true effect lies close to that of the estimate of the effect.

**Moderate certainty:** we are moderately confident in the effect estimate; the true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different.

Low certainty: our confidence in the effect estimate is limited; the true effect may be substantially different from the estimate of the effect.

Very low certainty: we have very little confidence in the effect estimate; the true effect is likely to be substantially different from the estimate of effect.



<sup>a</sup>Downgraded one level due to study limitations (unclear risk of bias for the index test in two studies and reference standard in one study) and one level due to inconsistency.

<sup>b</sup>Downgraded one level due to study limitations (unclear risk of bias for the index test and reference standard).

<sup>c</sup>Downgraded one level due to study limitations (unclear risk of bias for the index test and reference standard) and one level due to indirectness (no existing validation data).

<sup>d</sup>Downgraded three levels due to study limitations (retrospective study design and children with no imaging were not followed up, increasing the risk of missed cervical spine injuries) and one level due to indirectness (no existing validation data).

<sup>e</sup>Downgraded two levels due to study limitations (retrospective study design) and one level due to indirectness (only children with cervical spine imaging were included).



# BACKGROUND

### **Target condition being diagnosed**

Paediatric cervical spine injury (CSI) after blunt trauma is rare, accounting for an estimated 1% to 2% of trauma presentations in children (Garton 2008; Leonard 2019; Mohseni 2011; Patel 2001; Shin 2016; Viccellio 2001). However, the consequences of CSI can be devastating and include death or life-changing neurological damage (Cirak 2004; Hutchings 2009; Kokoska 2001; Leonard 2014; Mohseni 2011; Parent 2011; Patel 2001; Platzer 2007). Concern also exists that undiagnosed CSIs may lead to worsening of neurological symptoms and outcomes (Mortazavi 2011; Ravichandran 1982; Schuster 2005). The long-term prognosis for children who sustain cervical spinal cord injury and survive the first 24 hours is poor; life expectancy is reduced by eight to 53 years, depending on the anatomical level of injury and the degree of spinal cord involvement (NSCISC 2019). Due to prolonged hospital stay, the large number of treatments and associated long-term assistance required, severe paediatric CSI is associated with very high medical, psychological and societal costs (Shavelle 2007; Vogel 2002a; Vogel 2002b; Vogel 2002c). Therefore, in children presenting after blunt trauma, physicians seek to identify all CSIs promptly through a combination of clinical features and imaging tests.

Paediatric CSIs differ from the adult pattern of injury, particularly at younger ages (Junewick 2010; Kreykes 2010; Mohseni 2011; Mortazavi 2011; Parent 2011; Viccellio 2001). Anatomical and behavioural differences account for this. A higher proportion of injuries occur in the upper or axial cervical spine (Occiput to C2) at the younger ages (Kokoska 2001; Leonard 2014; Mohseni 2011; Patel 2001). Injuries of the upper cervical spine are associated with higher morbidity and mortality than those of the lower cervical spine (Leonard 2014; Patel 2001). A more adult pattern of injury is established by late childhood/early teenage years with equal proportions or slightly higher numbers of subaxial CSIs described at older ages (Leonard 2014; Mohseni 2011). A cut-off around eight years of age has often been used to separate injury patterns in older and younger children; some series also separate children under two or three years of age (Kokoska 2001; Leonard 2014; Leonard 2015; Viccellio 2001). At younger ages, the flexibility of the vertebral column also considerably outweighs the capacity for stretch of the spinal cord proper, and spinal cord injury may occur without bony injury. Reported spinal cord injury incidence rates in paediatric CSI vary between 17% and 35% (Gargas 2013; Leonard 2014; Patel 2001).

Several blunt trauma mechanisms can cause paediatric CSI. At all ages, motor vehicle-related accidents account for the largest proportion of injuries, with falls generally described as the second most common mechanism (Leonard 2014; Leonard 2019; Mohseni 2011; Nunn 2021; Patel 2001; Polk-Williams 2008; Shin 2016). In older children, sporting and other recreational activities account for a significant proportion of CSIs (Babcock 2018; Cirak 2004; Leonard 2014; Mortazavi 2011); in younger children, pedestrian accidents and inflicted injuries are also described (Leonard 2014; Mortazavi 2011). Specific mechanisms that involve an axial load, or head-first impact, such as diving may predispose to CSI (Leonard 2011; Leonard 2019).

Clinical features associated with CSI have been described in adult and paediatric studies (Hoffman 2000; Leonard 2011; Leonard 2019; Stiell 2001). These factors may include features of examination (e.g. posterior neck tenderness, torticollis, abnormal neurology, altered level of consciousness, significant other injury), and history (e.g. neck pain, previous CSI, underlying predisposing conditions such as Down's syndrome). Clinical assessment in children may be further complicated by the child's ability to both communicate and co-operate with clinical examination and their ability to discriminate symptoms suspicious for CSI, such as midline neck tenderness from discomfort, anxiety and other injury complaints. Neck pain rather than specific posterior midline tenderness has been described as a factor associated with CSI in children (Leonard 2011). Age-related differences in factors associated within children have also been described (Leonard 2015).

Given the consequences of CSIs, physicians seek to identify all injuries, generally through the use of imaging modalities such as plain radiography (X-ray), computed tomography (CT) or magnetic resonance imaging (MRI). X-ray and CT are the most common initial imaging tests used by physicians to diagnose or rule out paediatric CSI. In contrast to adult practice where CT is considered the gold standard (National Clinical Guideline Centre 2014; Ryken 2013), X-ray is often advocated as the first-line investigation in children (Browne 2003; Burns 2011; Chaudhry 2016; Chung 2011; Hannon 2015; Herman 2019; National Clinical Guideline Centre 2014; Nigrovic 2012; Slaar 2017). Some guidelines and publications suggest a combination of X-ray and targeted upper cervical spine CT (Chung 2011; Garton 2008; Sun 2013); others recommend CT in children deemed to be at higher risk, such as those with altered levels of consciousness, although this definition of "higher risk" often varies (Easter 2011; Hannon 2015; Herman 2019; Mortazavi 2011; National Clinical Guideline Centre 2014).

X-rays are associated with significantly lower radiation doses than CT scans (Booth 2012; Jimenez 2008); however, missed CSI rates of 10% to 25% have been reported with plain X-rays in children (Chung 2011; Garton 2008; Nigrovic 2012; Rana 2009). CT, while superior to X-ray in detecting bony CSI (Parizel 2010; Ryken 2013), may also not show all paediatric CSIs or their extent (Rozzelle 2013). One study reported a sensitivity of 23% (specificity 100%) in detecting soft tissue abnormalities in children (Henry 2013); other retrospective reviews have found 17% (Gargas 2013) and 20% (Nunn 2021) of paediatric CSIs not to be apparent on CT.

MRI is often used as a second- or third-line test, particularly where neurological symptoms are present, the patient is unable to be clinically assessed or there are ongoing concerns of CSI. While a superior modality for spinal cord and spinal soft tissue abnormalities (Parizel 2010) and free of ionising radiation, it has several limitations including availability, cost, time to perform the scan and possible need for sedation or general anaesthesia due to the prolonged immobilisation required in young or unco-operative children. It may also be less ideal than CT scan for bony injuries. These factors have limited consideration of MRI use in screening after blunt trauma to date. In addition, the clinical significance of the injuries detected on MRI is also sometimes unclear (Booth 2012).

Spinal cord injury without radiographic abnormalities (SCIWORA) has been described in children with spinal cord injury, with incidence varying greatly between studies (Bosch 2002; Brown 2001; Cirak 2004; Farrell 2017; Gore 2009; Kreykes 2010; Leonard 2014; Mahajan 2013; Mortazavi 2011; Pang 2004; Polk-Williams 2008; Yucesoy 2008). One review reported rates between 5% and 67%, with an overall incidence estimated at around 35% (Pang



2004). The SCIWORA terminology predates the widespread use and availability of MRI (Pang 1982), and there is ongoing definitional debate as to whether the term should include MRI detectable injury or not (Farrell 2017; Yucesoy 2008). Some more recent studies have divided SCIWORA into two categories, those with MRI abnormalities and those with a normal MRI, with more favourable clinical outcomes described in the latter group (Farrell 2017; Mahajan 2013).

It is unethical and unfeasible to image all children presenting with blunt trauma for possible CSI given concerns about unnecessary exposure to ionising radiation (X-ray, CT) and increased lifetime cancer risk (Brenner 2007; Chen 2014; Mathews 2013; Miglioretti 2013; Pearce 2012), the risks of sedation (CT, MRI) (Cutler 2007; Goldwasser 2015; Hoyle 2014), resource implications (cost, time, bed space) and patient discomfort with prolonged assessments (Chan 1994; Leonard 2012; March 2002; Sundstrom 2014). Physicians are thus faced with the decision of which children require imaging and in whom it can be safely avoided (i.e. which children are considered at very low risk of CSI and can be "clinically cleared" without imaging). Ideally, a well-evidenced clinical decision rule (CDR) or tool would be administered during the initial clinical assessment to guide this clinical decision-making process, and minimise unnecessary tests and their associated risks, whilst detecting all significant CSIs.

#### Index test(s)

The tools under evaluation are any CDRs or sets of clinical criteria used to evaluate CSI in children and adolescents following blunt trauma that provide guidance on whether imaging is required, or whether it can be safely avoided. CDRs are composed of at least three variables of history, examination findings or simple tests, and are applied during the initial clinical assessment, prior to any imaging. The definition of a positive result is dependent on whether the rule aims to identify children at high risk of CSI, who require further imaging, or at very low risk of CSI, where imaging is not required. These tools may be prospectively or retrospectively derived.

The development of a CDR is a three-step process involving derivation, validation and impact analysis (assessing the impact of the rule on clinician behaviour) (Laupacis 1997; McGinn 2000; Stiell 1999). The thresholds for a positive result are dependent on the nature of the CDR, whether the CDR is intended to identify children at high risk of CSI, who require further imaging, or at very low risk of CSI, where imaging is not required.

Tools derived for adults that are commonly used for children, such as the National Emergency X-Radiography Utilization Study (NEXUS) (Viccellio 2001) and the Canadian C-Spine Rule (Stiell 2001), were also evaluated providing their use was assessed in children. Tools evaluated were not limited by primary imaging modality recommended (if any).

# **Clinical pathway**

The clinical pathway for assessment of possible CSIs is part of the standard trauma workup for children presenting with blunt trauma to the emergency department (ED). Clinical features such as history, mechanism of injury and examination findings are considered to determine whether radiographic imaging of the cervical spine is indicated. A well-evidenced CDR or set of clinical criteria guiding

clinicians on whether a child is at higher or lower risk of CSI would assist in this decision-making process. It would allow those at higher risk of CSI to be more accurately identified and unnecessary tests with their incumbent risks, such as exposure to ionising radiation, and costs, to be avoided. Children are also usually immobilised during assessment for CSI with their movements heavily restricted (often confined to lying still on a bed) and they may wear a cervical spine collar or neck brace (Chan 1994; Leonard 2012; March 2002; Sundstrom 2014). Early identification of children at low risk and suitable for clinical clearance (i.e. clearance without imaging) would also minimise these discomforts and free hospital resources, benefiting both the patient and healthcare system.

#### Rationale

Paediatric CSIs, while rare, can have devastating consequences, including death and long-term disability. Radiographic imaging is used to identify CSIs but has significant risks and costs including exposure to ionising radiation and increased lifetime cancer risks. Physicians aim to identify all CSIs in children whilst minimising unnecessary imaging in children deemed low risk for CSI. CDRs or tools can assist in this decision-making process, but evidence to support their use in paediatric populations has been limited.

Well-established adult rules to guide the decision to image in possible CSI exist (Hoffman 2000; Stiell 2001); however, the validity of their use in children, particularly at younger ages, has been questioned (Garton 2008; Slaar 2017; Viccellio 2001). In more recent years, clinical criteria or risk factors derived from specific paediatric cohorts have been proposed as possible CDRs or tools specifically for children (Leonard 2011; Leonard 2019).

This review seeks to expand the previous Cochrane review (Slaar 2017) beyond examining NEXUS and Canadian C-Spine Rule tests to include all CDRs or sets of clinical criteria used to evaluate CSI (and guide the decision to image) in children and adolescents following blunt trauma. It will also ascertain whether any new evidence exists to inform the use of these well-established adult rules in children.

#### OBJECTIVES

To assess and compare the diagnostic accuracy of clinical decision rules (CDRs) or sets of clinical criteria, alone or in comparison with each other, for the evaluation of CSI following blunt trauma in children.

#### Secondary objectives

We were unable to explore heterogeneity in this review due to the small number of studies for each CDR. However, we will explore the following types of heterogeneity in the estimates of diagnostic test accuracy in subsequent reviews if the number of studies increases, including:

- differences in the healthcare setting and study design: previous studies stated that CSI is seen less often in general EDs than in paediatric trauma hospitals, and that a CT of the neck is more common in general EDs (Adelgais 2014);
- study quality, as assessed by the Quality Assessment of Diagnostic Accuracy Studies Version 2 (QUADAS-2) checklist (Whiting 2011);
- age-related differences: we hypothesised that both the applicability of the CDRs and the type of injury would differ



according to age in children younger than eight years and eight years or older (Leonard 2014).

# METHODS

#### Criteria for considering studies for this review

#### **Types of studies**

We included studies that compared the diagnostic accuracy of any CDR or clinical criteria compared with a reference standard for the evaluation of CSI in children presenting to EDs following blunt trauma. De novo CDRs and validated CDRs were included; however, de novo CDRs were considered at high risk of bias due to the lack of validation data. Eligible study designs included diagnostic studies with cross-sectional or cohort designs (retrospective or prospective) and randomised controlled trials. We included studies evaluating one CDR or comparing two or more CDRs (directly and indirectly). We only included results from full reports. We excluded case-control studies because of the bias they might introduce and reports that evaluated predictor finding models.

#### Participants

We included children (aged 0 to less than 18 years) who underwent blunt trauma evaluation in the ED. We excluded studies specifically focused on children with a history of previous surgery of the cervical spine or congenital cervical spine anomalies, or both.

In studies with mixed populations where data related to participants aged less than 18 years could not be separated from older participants, we attempted to contact the study authors for more information. If we were unable to contact study authors to request additional data, we listed them in the Studies awaiting classification table. However, if study authors confirmed that age disaggregated data were unavailable, we excluded the studies.

#### Index tests

The tests under evaluation were any CDR or set of clinical criteria that compared the diagnostic accuracy of the test for the target condition with the reference standard.

#### **Target conditions**

The target condition was clinically important CSI, defined as any fracture, dislocation, ligamentous injury or spinal cord injury (either detectable by diagnostic imaging or spinal cord injury without radiographic association) involving the cervical region and attached ligamentous structures.

#### **Reference standards**

Evaluation for CSI in the ED requires expert clinical assessment and often diagnostic imaging. X-ray, CT and MRI can be used in the assessment of CSI with each modality having different strengths and weaknesses. Therefore, we included studies in which participants were diagnosed with CSI using any reference standard, that is, X-ray, CT or MRI, following presentation to the ED.

In children who are low risk for CSI, it may not be feasible or ethical to perform diagnostic imaging for reasons described in the Background. As such, we also included studies where children received follow-up in the ED to clinically clear the cervical spine in individuals deemed at lower-risk by the treating clinician. Clinical follow-up in the ED was defined as clinical evaluation of the neck after removal of the neck collar (if worn) in these children.

To reduce the risk of incorrectly classifying children as positive or negative for CSI, children who received only clinical follow-up in the ED and no imaging should have had an additional follow-up some time after discharge. This additional follow-up may have involved clinical evaluation of the neck by a treating clinician, review of the medical record for additional imaging of the cervical spine, telephone follow-up to verify the absence of CSI or other systems to ensure initially missed CSI were diagnosed.

We included studies with participants who did not receive imaging or an additional clinical follow-up after discharge from the ED; however, this was considered a source of bias in the QUADAS-2 assessment.

We included children who underwent an eligible reference standard or obtained clinical follow-up within 72 hours of presentation at the ED following blunt trauma.

### Search methods for identification of studies

#### **Electronic searches**

The Information Specialist of the Cochrane Back and Neck Review Group developed the initial search strategy for the previous version of the review (Slaar 2017). For the update, a specialist librarian reviewed and modified the search strategy when necessary, based on updates to thesaurus terms for each database listed below. The search strategies for identifying diagnostic test accuracy studies consisted of controlled vocabulary and keyword terms for each of the following concepts: the index or reference test, the target condition and the patient description.

We searched the following databases from 1 January 2015 to 13 December 2022 to capture any studies published since the previous version of the review (Slaar 2017).

- Cochrane Central Register of Controlled Trials (CENTRAL, in the Cochrane Library) Issue 12, 2022
- MEDLINE Ovid
- Embase Ovid
- ProQuest Dissertations & Theses Database for relevant conference proceedings, dissertations, and theses
- PubMed (www.ncbi.nlm.nih.gov/pubmed)
- OpenGrey for 'grey literature' (www.opengrey.eu/)
- ClinicalTrials.gov (clinicaltrials.gov/)
- Science Citation Index (Web of Science, Core Collection)
- World Health Organization International Clinical Trials Registry Platform (ICTRP) (www.who.int/clinical-trials-registryplatform)

The search strategies can be found in Appendix 1.

#### Searching other resources

We re-examined all studies excluded from the previous version of the review to determine if any studies should now be included based on our expanded index test eligibility criteria to include all CDRs. We sought to identify additional studies through searching reference lists of primary studies and relevant systematic reviews. We also contacted authors to request information of

any unpublished or ongoing studies. There were no language restrictions. In non-English full-text studies, we first examined the English abstract and title for eligibility. If the abstract was potentially relevant, we planned to translate the full text.

#### Data collection and analysis

#### **Selection of studies**

For this review update, we uploaded and screened references in Covidence. Pairs of review authors (NE, ET, VR, JW) independently screened each title and abstract identified in the search. We retrieved full texts for potentially relevant references, and two review authors independently screened them. We resolved disagreements by recourse to a third review author. Two review authors were authors of one of the included studies. They were not involved in the screening and selection of papers.

#### Data extraction and management

Two review authors (ET, NE) independently extracted data using a piloted data extraction form, resolving disagreements by consultation with a third review author if necessary. We contacted study authors to request additional information.

Data collected included:

- study ID (year of publication, author, citation);
- study design (consecutive/random, retrospective/prospective, cohort, cross-sectional, randomised controlled trials);
- sample characteristics (number of participants and children enrolled and analysed, age, sex);
- setting (type of acute care setting(s), location);
- inclusion and exclusion criteria;
- index test(s) used (name, clinical criteria, interpretation);
- target condition (definition, prevalence in sample);
- reference standard (definition, any information related to execution or interpretation);
- results (data to populate a two-by-two table).

This information was documented for each study in the Characteristics of included studies table.

Two review authors were authors of one of the included studies. They were not involved in the data extraction.

### Assessment of methodological quality

Two review authors (ET, NE), independently and in duplicate, assessed the methodological quality of each study, using the QUADAS-2 tool (Whiting 2011). We resolved disagreements by consensus.

The QUADAS-2 tool consists of four domains: patient selection, index tests, reference standard, and flow and timing. For this update, we modified the patient selection domains by removing one core signalling question that was addressed by the eligibility criteria; "was a case-control design avoided?" and added an additional signalling question to check if the data were collected prospectively. Retrospective data are prone to selective and incomplete recording. We clarified the meaning of the flow and timing domain signalling question "did all patients receive the same reference standard?" to be more clinically appropriate. The same reference standard could be either all patients underwent the same type of imaging or all patients underwent follow-up after discharge. The tailored version of the tool is provided in Table 1.

Studies that evaluated de novo CDRs were rated as high risk of bias in the index test domain because of the lack of existing validation data. In the current review update, the reference standards were radiographic imaging. However, the decision to obtain imaging was at the treating clinicians' discretion, and some children received clinical clearance of the cervical spine without imaging. We preferred if children who did not receive imaging underwent a follow-up some time after discharge to ensure no CSIs were missed. Therefore, we rated studies with children who did not receive imaging or follow-up after discharge as high risk of bias in the reference standard domain. If studies only included those children that received imaging, excluding those who presented with blunt trauma and were cleared clinically, this was recorded as high risk for the applicability question in the patient selection domain.

The risk of bias judgement ('high', 'low' or 'unclear') for each domain was dependent on the signalling questions. If the answers to all signalling questions within a domain were judged as 'yes' (indicating low risk of bias for each question), then the domain was judged at low risk of bias. If any signalling question was judged as 'no' (indicating a high risk of bias), the overall domain was also categorised at high risk of bias.

#### Statistical analysis and data synthesis

We extracted indices of the diagnostic performance of all clinical tools from data presented in each study. We generated diagnostic two-by-two tables, from which we calculated sensitivities and specificities for each index test with 95% confidence intervals (CI), and presented them in forest plots and also in a receiver operator curve (ROC) space. If data presented in trials were uninterpretable to generate two-by-two tables, we contacted the authors of the study requesting clarification.

We planned to perform meta-analyses of sensitivity and specificity employing a bivariate logistic normal model using a hierarchical approach (Reitsma 2005). This approach would enable us to calculate summary estimates of sensitivity and specificity while dealing with sources of variation within and between studies and any correlation that might exist between sensitivity and specificity. With the model estimates, we aimed to plot sensitivities and specificities in forest plots and in ROC space. If we had identified a sufficient number of studies with direct or indirect comparison between two or more index tests, we planned to use the methods proposed by Nyaga 2018 for meta-analysis. These methods will be used for randomised controlled trials and studies that used both direct and indirect comparisons.

We also planned to compare the different index tests and tried to find whether these tests had different sensitivities or specificities, employing a bivariate model.

We planned to compare tests by adding covariates for different types of index tests into the bivariate model and testing the significance (P = 0.05) of the parameters of covariates. If almost none of the primary studies directly compared these tools, we would have included all studies that evaluated at least one of the index tests into the test comparison. In other words, test comparison would not be limited to direct comparisons, but would have used all the evidence available. We planned to compare them

qualitatively if data were insufficient for comparison by statistical tests. Verification bias is to be expected, since we included the use of different types of reference standards in test-positive (X-ray, CT scan, or MRI) and test-negative (clinical follow-up). All the statistical analyses were performed using the analysis functions of Review Manager 5 (Review Manager 2014).

#### Investigations of heterogeneity

We intended to investigate sources of heterogeneity in terms of differences in the healthcare setting, study design, study quality and age-related differences. However, we were unable to formally explore heterogeneity, due to a lack of relevant studies. If sufficient data becomes available in future updates, we will use forest plots and sensitivities and specificities plotted in ROC space for visual examination of heterogeneity between studies. We will add covariates, for example, age groups (less than eight years of age versus eight years and older) and QUADAS-2 items bivariate model to investigate the heterogeneity between studies in the meta-analysis (Whiting 2011). We could only have investigated heterogeneity if there was a sufficient number of studies providing adequate information on the factor of interest.

#### Sensitivity analyses

We planned to undertake sensitivity analysis by removing studies at high risk of bias. However, we were unable to conduct sensitivity analyses because there were too few studies.

#### Assessment of reporting bias

As yet there are no quantitative methods for reporting bias in diagnostic test accuracy studies; therefore, we did not assess reporting bias.

# Summary of findings and assessment of the certainty of the evidence

We summarised key findings in Summary of findings 1 and Summary of findings 2. We assessed the certainty of evidence using the GRADE approach, which evaluates five domains: risk of bias, indirectness, inconsistency, imprecision and publication bias (Schünemann 2020). We explained our decisions to downgrade the certainty of evidence in the footnotes of the summary of findings tables.

# RESULTS

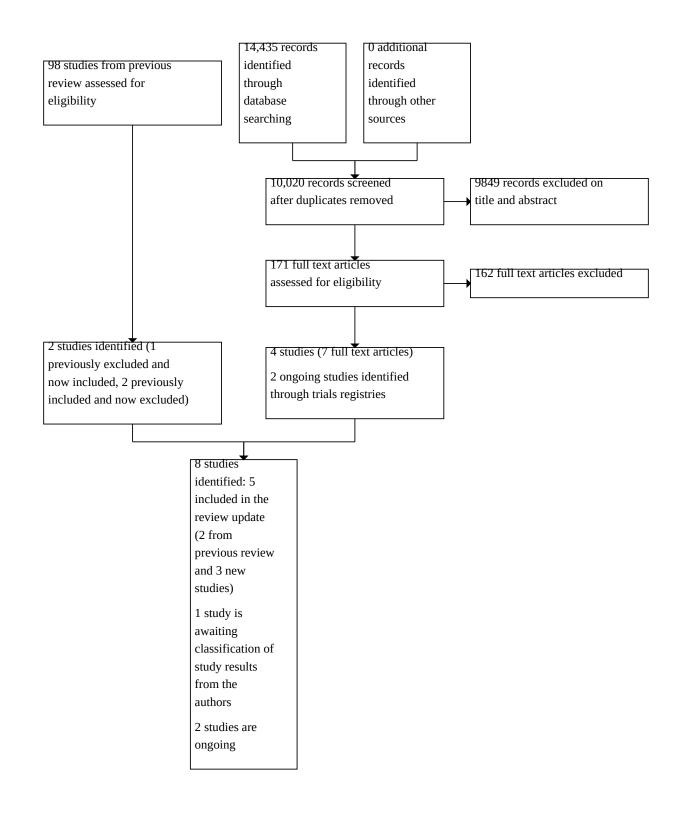
#### **Results of the search**

We identified 14,435 citations in the search update on 13 December 2022. The number of citations by search engine is shown in Table 2. After removal of duplicates, we screened 10,020 records by title and abstract and excluded 9849. Due to changes in the inclusion criteria for this review update, we evaluated the 98 fulltext articles assessed for eligibility in the previous version of the review (included and excluded studies), in addition to the 171 fulltext articles identified in the search update. We excluded 162/171 full-text articles identified through the updated search. The main reasons for exclusion were irrelevant index tests or different study population (e.g. prehospital or adult study sample). We excluded two studies evaluated in the previous review (Ehrlich 2009; Jaffe 1987). We excluded Jaffe 1987 because it included a second nonconsecutive cohort of participants with CSI, and Ehrlich 2009 because of the case-matched study design. Further details are provided in the Characteristics of excluded studies table.

This update includes five studies: one study evaluated in the previous review (Viccellio 2001); one study excluded from the previous review and now included (Pieretti-Vanmarcke 2009); and three new studies identified in the updated search (Davies 2016; Leonard 2019; Phillips 2021). Two studies are awaiting classification, with authors contacted for further eligibility information (Arbuthnot 2017; Vargas 2022), and we identified two ongoing studies (ACTRN12621001050842; NCT05049330). Refer to Figure 1 for the PRISMA flow diagram of search results and screening results.



# Figure 1. Study flow diagram.





#### **Included studies**

We reported the main characteristics of the five studies in the Characteristics of included studies table. All studies were reported in full-text publications.

Included studies reported on the diagnostic accuracy of seven CDRs, summarised in Table 3; five previously described CDRs (NEXUS, Canadian C-Spine Rule, PECARN retrospective criteria, National Institute for Health and Care Excellence (NICE) clinical guideline 56 (CG56) and 176 (CG176)) and two derived CDRs (Leonard de novo (Leonard 2019), PEDSPINE (Pieretti-Vanmarcke 2009)). One study directly compared the accuracy of three index tests (Phillips 2021), two studies directly compared the accuracy of two studies reported the accuracy of one index test (Pieretti-Vanmarcke 2009; Viccellio 2001). The inclusion criteria for age of children were: less than 10 years of age following blunt trauma (Davies 2016), younger than 18 years of age (Leonard 2019; Viccellio 2001), under 16 years of age (Phillips 2021), and three years of age and younger (Pieretti-Vanmarcke 2009).

The total number of children enrolled in the included studies was 21,379 and ranged from 278 (Davies 2016) to 12,882 (Pieretti-Vanmarcke 2009). The prevalence of CSI ranged from 0.5% (Phillips 2021) to 1.85% (Davies 2016). Studies were conducted in the USA

(Leonard 2019; Viccellio 2001); the UK (Davies 2016); Australia (Phillips 2021); and across the USA, Canada and Brazil (Pieretti-Vanmarcke 2009). The studies were published between 2001 and 2021.

#### Excluded studies

We excluded 164 reports with reasons (Characteristics of excluded studies table).

#### Studies awaiting classification

Two studies are awaiting classification while we await replies from authors (Arbuthnot 2017; Vargas 2022; Characteristics of studies awaiting classification table).

#### **Ongoing studies**

We identified two ongoing studies (ACTRN12621001050842; NCT05049330; Characteristics of ongoing studies table).

#### Methodological quality of included studies

We reported the results of the methodological quality assessment of included studies in the Characteristics of included studies table. Figure 2 summarises the results of the quality assessment of the included studies and an individual assessment for each study is provided in Figure 3.

# Figure 2. Risk of bias and applicability concerns graph: review authors' judgements about each domain presented as percentages across included studies.

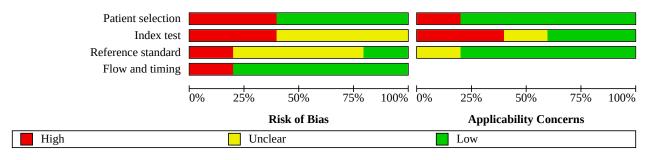
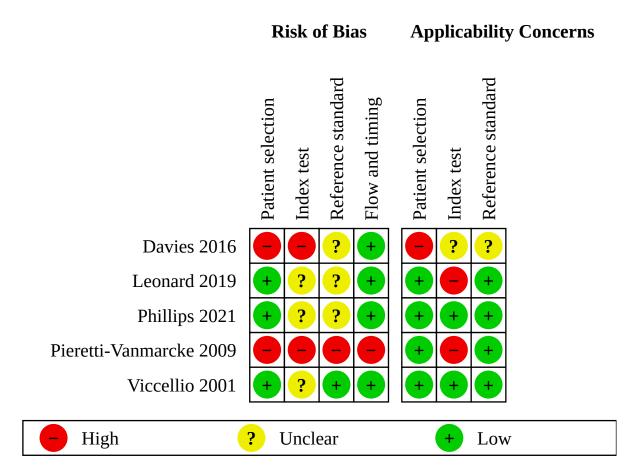




Figure 3. Risk of bias and applicability concerns summary: review authors' judgements about each domain for each included study.



#### **Patient selection**

We considered two studies at high risk of bias in the patient selection domain as they were retrospective studies of medical records or trauma registries (Davies 2016; Pieretti-Vanmarcke 2009), with the remaining studies rated at low risk of bias. Davies 2016 was rated as high concern of applicability as they included only those children who presented and had imaging of the cervical spine. Retrospective studies rely on routinely collected data and are, therefore, susceptible to selective and incomplete recording. Davies 2016 excluded eight children due to incomplete data. Four studies were at low concern of applicability because they were of prospective cohort design, included a consecutive or random sample of participants and avoided inappropriate exclusions.

#### Index test

Two studies applied the index test retrospectively, after the reference standard was interpreted and were rated at high risk of bias (Davies 2016; Pieretti-Vanmarcke 2009). Three studies of prospective design collected data on rule predictor variables prior to radiographic imaging (Leonard 2019; Phillips 2021; Viccellio 2001). However, there were insufficient details on the blinding of imaging results prior to index tests being interpreted. Therefore, they were assessed at unclear risk of bias. All studies reported the use of a prespecified threshold.

Two studies derived a new CDR (Leonard de novo and PEDSPINE) and were, therefore, rated as high concern for applicability due to an absence of existing validation data (Leonard 2019; Pieretti-Vanmarcke 2009). However, Leonard 2019 also validated an existing CDR, the PECARN retrospective criteria, and, for this rule, we had low concerns for its applicability. Davies 2016 was rated as unclear applicability concerns because of their retrospective study design. We had low concern for applicability in the index test domain for Phillips 2021 and Viccellio 2001 because they applied validated CDRs prospectively.

#### **Reference standard**

In two studies, the reference standard was radiographic imaging (X-ray, CT or MRI) in all children (Davies 2016; Viccellio 2001). Two studies had a reference standard of radiographic imaging or for those who did not receive imaging, clinical clearance in the ED with subsequent telephone follow-up after discharge for all children (Leonard 2019; Phillips 2021). The final study had radiographic imaging for some children and clinical clearance in the ED for others (Pieretti-Vanmarcke 2009).

The reference standard domain was at low risk of bias in one study (Viccellio 2001), unclear risk of bias in three studies (Davies 2016; Leonard 2019; Phillips 2021), and high risk of bias in one study (Pieretti-Vanmarcke 2009). Pieretti-Vanmarcke 2009 was at high risk of bias because it did not follow up after discharge for children



who did not receive imaging. Therefore, there was a risk that the target condition may have been incorrectly classified for these children. Although Leonard 2019 and Phillips 2021 both included follow-up after discharge for children who did not receive imaging, there was insufficient information provided on whether those who interpreted the reference standard (radiologists) had knowledge of the index test results or data collection forms. Davies 2016 was at unclear risk of bias and unclear concern for applicability because the target condition, CSI, was not defined. All other studies were rated at low concern for applicability as the target condition was clearly defined.

#### Flow and timing

NEXUS

One study was at high risk of bias in the flow and timing domain (Pieretti-Vanmarcke 2009). There was an appropriate interval between presentation to the ED and conduct of the index test/ reference standard and children included in the analysis were clearly defined and any exclusions were for appropriate reasons (e.g. child did not meet inclusion criteria or data were missing). However, children did not receive the same reference standard and those not receiving imaging were not followed up after discharge, increasing the risk of verification bias of the index test.

#### Findings

We evaluated five studies, which enrolled 21,379 children, for the presence of CSI using the NEXUS criteria, Canadian C-Spine Rule, PECARN retrospective criteria, NICE CG56 and CG176 and assessment of the certainty of the evidence (Davies 2016; Leonard 2019; Phillips 2021; Pieretti-Vanmarcke 2009; Viccellio 2001), and two derived CDRs (Leonard de novo (Leonard 2019) and PEDSPINE (Pieretti-Vanmarcke 2009)). The average incidence of CSI in these five studies was 1.16% with a median prevalence of CSI of 0.98% (interquartile range (IQR) 0.50% to 1.85%). Since the number of eligible studies for each CDR was fewer than four, and the inclusion criteria and outcomes of those studies were too diverse, we did not conduct a meta-analysis and therefore presented no summary estimates in this review. Instead, we interpreted sensitivity and specificity from each primary study separately. Since there were few data, we were unable to investigate heterogeneity. We generated diagnostic two-by-two tables, from which we calculated sensitivities and specificities for each index test with 95% CIs, and presented them in forest plots (see Figure 4 and Summary of findings 1). We could not perform sensitivity analyses because there were too few studies.

# Figure 4. Forest plot of NEXUS, Canadian C-spine Rule, NICE Guidelines (CG56 and CG176), PECARN Retrospective, Leonard de novo and PEDSPINE.

NEXUS							
Study	ТР	FP	FN	TN	Sensitivity (95% CI)	Specificity (95% CI)	Sensitivity (95% CI) Specificity (95% CI)
Phillips 2021	5	425	0	543	1.00 [0.48, 1.00]	0.56 [0.53, 0.59]	
Viccellio 2001	30	2432	0	603	1.00 [0.88, 1.00]	0.20 [0.18, 0.21]	
Canadian C-Spine Rule							0 0.2 0.4 0.6 0.8 1 0 0.2 0.4 0.6 0.8
Study	ТР	FP	FN	TN	Sensitivity (95% CI)	Specificity (95% CI)	Sensitivity (95% CI) Specificity (95% CI)
Phillips 2021	5	467	0	501	1.00 [0.48, 1.00]	0.52 [0.49, 0.55]	
PECARN Retrospective							0 0.2 0.4 0.6 0.8 i 0 0.2 0.4 0.6 0.8
Study	ТР	FP	FN	TN	Sensitivity (95% CI)	Specificity (95% CI)	Sensitivity (95% CI) Specificity (95% CI)
Leonard 2019	67	2186	7	1831	0.91 [0.81, 0.96]	0.46 [0.44, 0.47]	-+ -
Phillips 2021	5	658	0	310	1.00 [0.48, 1.00]	0.32 [0.29, 0.35]	
Leonard de novo							0 0.2 0.4 0.6 0.8 1 0 0.2 0.4 0.6 0.8
Study	ТР	FP	FN	TN	Sensitivity (95% CI)	Specificity (95% CI)	Sensitivity (95% CI) Specificity (95% CI)
Leonard 2019	68	1998	6	2019	0.92 [0.83, 0.97]	0.50 [0.49, 0.52]	
PEDSPINE							0 0.2 0.4 0.6 0.8 1 0 0.2 0.4 0.6 0.8
Study	ТР	FP	FN	TN	Sensitivity (95% CI)	Specificity (95% CI)	Sensitivity (95% CI) Specificity (95% CI)
Pieretti-Vanmarcke 2009	28	1224	2	2925	0.93 [0.78, 0.99]	0.70 [0.69, 0.72]	
NICE guideline (CG56)							0 0.2 0.4 0.6 0.8 1 0 0.2 0.4 0.6 0.8
Study	ТР	FP	FN	TN	Sensitivity (95% CI)	Specificity (95% CI)	Sensitivity (95% CI) Specificity (95% CI)
Davies 2016	5	144	0	121	1.00 [0.48, 1.00]	0.46 [0.40, 0.52]	·····
NICE guideline (CG176)							0 0.2 0.4 0.6 0.8 i 0 0.2 0.4 0.6 0.8
Study	ТР	FP	FN	TN	Sensitivity (95% CI)	Specificity (95% CI)	Sensitivity (95% CI) Specificity (95% CI)
Davies 2016	5	246	0	19	1.00 [0.48, 1.00]	0.07 [0.04, 0.11]	
							0 0.2 0.4 0.6 0.8 1 0 0.2 0.4 0.6 0.8



#### **Direct comparisons**

# NEXUS, Canadian C-Spine Rule and PECARN retrospective criteria

Phillips 2021, a prospective single-centre study, directly compared the accuracy of NEXUS, Canadian C-Spine Rule and the PECARN retrospective criteria in an enrolled cohort of 1010 children aged under 16 years who were immobilised before arrival to the ED for possible CSI, presented with neck pain in the context of trauma or considered at risk of neck injury by the ED team. The reference standard was imaging (X-ray, CT or MRI) or clinical clearance with all children followed up by telephone to ensure no CSIs were missed. The CSI prevalence in the children included in the analysis (973 children) was 0.5%. The sensitivities of NEXUS, Canadian C-Spine Rule and the PECARN retrospective criteria were 1.00 (95% CI 0.48 to 1.00), 1.00 (95% CI 0.48 to 1.00) and 1.00 (95% CI 0.48 to 1.00), respectively. The sensitivities of the three CDRs were not provided in the paper, with the authors deeming a formal validation inappropriate due to the low incidence of CSI (0.5%). The specificities for NEXUS, Canadian C-Spine Rule and the PECARN retrospective criteria were 0.56 (95% CI 0.53 to 0.59), 0.52 (95% CI 0.49 to 0.55) and 0.32 (95% CI 0.29 to 0.35), respectively. We considered the evidence for the estimates of sensitivity and specificity to be moderate certainty, downgraded one level for risk of bias.

#### Pediatric Emergency Care Applied Research Network (PECARN) retrospective criteria and Leonard de novo

Leonard 2019, a prospective multicentre study, evaluated the accuracy of the PECARN retrospective criteria in comparison to a de novo model (Leonard de novo) in 4091 children younger than 18 years of age, who presented to the ED for blunt trauma and were transported from the scene of the injury by emergency medical services in spinal motion restriction devices, underwent trauma team evaluation, had cervical spine imaging ordered in the ED, or combinations of these. The reference standard was imaging (X-ray, CT or MRI), medical record review 21 days later for subsequent imaging and a follow-up call if no imaging was noted to ensure no CSIs were missed. The prevalence of CSI was 1.8%.

The PECARN retrospective criteria were derived from a multicentre retrospective case-control study in children with blunt trauma and described eight variables associated with paediatric CSI which, if applied as a CDR, would have detected 98% of CSIs in their retrospective derivation cohort (Leonard 2011). The Leonard de novo model included CSI risk factors from the prospective study with good test accuracy in identifying CSIs (see Table 3 for details) (Leonard 2019). The sensitivities for the PECARN retrospective criteria and the Leonard de novo model were 0.91 (95% CI 0.81 to 0.96) and 0.92 (95% CI 0.83 to 0.97), respectively. The specificities for the PECARN retrospective criteria and the Leonard de novo model were 0.46 (95% CI 0.44 to 0.47) and 0.50 (95% CI 0.49 to 0.52), respectively. Although the sensitivity and specificity of the Leonard de novo was greater than the PECARN retrospective criteria (moderate-certainty evidence), the certainty of evidence was rated as low, downgraded one level due to risk of bias and one level due to indirectness/high concern for applicability due to an absence of existing validation data.

#### National Institute for Health and Clinical Excellence (NICE) Head Injury clinical guideline 56 and NICE Head Injury clinical guideline 176

Davies 2016 evaluated the diagnostic accuracy of the NICE Head Injury CG56 in comparison with the NICE Head Injury CG176 in the detection of paediatric CSI. A retrospective review was undertaken of all children under 10 years of age who underwent emergency cervical spine imaging following blunt trauma at a Level 1 trauma centre in the UK. The total number of enrolled participants was 278 and the prevalence of CSI in the children included in the analysis (270 children) was 1.85%. Davies 2016 was at high risk of bias in the patient selection domain as it was a retrospective study of medical records and rated at high concerns of applicability as only those children who presented and had imaging of the cervical spine were included. As the index test was applied retrospectively, after the reference standard was interpreted, it was also rated as high risk of bias in the index test domain.

The sensitivity of the CG56 guideline was 1.00 (95% CI 0.48 to 1.00) and specificity was 0.46 (95% CI 0.40 to 0.52). The sensitivity of the CG176 guideline was 1.00 (95% CI 0.48 to 1.00) and specificity was 0.07 (95% CI 0.04 to 0.11). We considered the evidence for the estimates of sensitivity and specificity to be very low certainty, downgraded two levels for risk of bias and one level for indirectness.

#### Indirect comparisons

#### National Emergency X-Radiography Utilization Study (NEXUS)

The NEXUS study was validated in a prospective, observational study in 21 EDs of 34,069 children who underwent radiography of the cervical spine after blunt trauma (2.5% were eight years old or younger) (Hoffman 2000). Two studies evaluated the diagnostic accuracy of the NEXUS criteria to evaluate for CSI in children following blunt trauma (Phillips 2021; Viccellio 2001). The total number of participants was 4038. The prevalences of CSI in the target populations were 0.5% (Phillips 2021) and 0.98% (Viccellio 2001).

Viccellio 2001, a prospective multicentre study and the paediatric cohort of the original NEXUS cohort, tested the accuracy of the NEXUS criteria in 3065 blunt trauma patients younger than 18 years who received cervical spine imaging (X-ray, CT, MRI or a combination of these). The number of children for whom a CT was obtained was unclear. Phillips 2021, a prospective single-centre study, tested the accuracy of NEXUS in direct comparison to the Canadian C-Spine Rule and to the PECARN retrospective criteria in an enrolled cohort of 1010 children aged under 16 years who were immobilised before arrival in the ED for possible CSI, presented with neck pain in the context of trauma or considered at risk of neck injury by the ED team. The reference standard was imaging (X-ray, CT or MRI) or clinical clearance with all children followed up by telephone to ensure no CSIs were missed. The sensitivity of the NEXUS criteria was calculated for Phillips 2021, as it was not provided in the paper. The authors deemed a formal validation inappropriate due to the low incidence of CSI (0.5%) in the 973 children included in the analysis.

The sensitivity of the NEXUS criteria in the studies was 1.00 (95% CI 0.88 to 1.00) (Viccellio 2001) and 1.00 (95% CI 0.48 to 1.00) (Phillips 2021). The specificity of the NEXUS criteria varied in the studies and was 0.20 (95% CI 0.18 to 0.21) (Viccellio 2001)

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and 0.56 (95% CI 0.53 to 0.59) (Phillips 2021). We considered the evidence for the estimates of sensitivity and specificity to be low certainty, downgraded one level for risk of bias and one level for inconsistency.

#### **Canadian C-Spine Rule**

While the original Canadian C-Spine Rule study was never validated in children (Stiell 2001), one study evaluated its diagnostic accuracy for CSI in children following blunt trauma (Phillips 2021). Phillips 2021, a prospective single-centre study, tested the accuracy of the Canadian C-Spine Rule in direct comparison to the NEXUS criteria and the PECARN retrospective criteria in 973 children aged under 16 years who were immobilised before arrival in the ED for possible CSI, presented with neck pain in the context of trauma or considered at risk of neck injury by the ED team. The reference standard was imaging (X-ray, CT or MRI) or clinical clearance with all children followed up by telephone to ensure no CSIs were missed.

The sensitivity of the Canadian C-Spine Rule in the study was 1.00 (95% CI 0.48 to 1.00) and the specificity was 0.52 (95% CI 0.49 to 0.55) (Phillips 2021). We calculated the sensitivity as it was not provided in the paper. The CIs for sensitivity were wide due to the small number of injuries (5/973 children had confirmed CSI (0.5%)). We considered the evidence for the estimates of sensitivity and specificity to be moderate certainty, downgraded one level for risk of bias.

# Pediatric Emergency Care Applied Research Network (PECARN) retrospective criteria

The PECARN criteria were derived from a retrospective case-control study and described eight variables associated with paediatric CSI which, if applied as a CDR, would have detected 98% of CSIs in their retrospective derivation cohort (Leonard 2011). Two studies evaluated the diagnostic accuracy of the PECARN retrospective criteria to evaluate for CSI in children following blunt trauma (Phillips 2021; Leonard 2019). The total number of participants was 5064 and the prevalence of CSI was 1.8% in Leonard 2019 and 0.5% in Phillips 2021. Leonard 2019, a prospective multicentre study, evaluated the accuracy of the PECARN retrospective criteria in 4091 children younger than 18 years of age, that presented to the ED for blunt trauma and were transported from the scene of the injury by emergency medical services in spinal motion restriction devices, underwent trauma team evaluation, had cervical spine imaging ordered in the ED, or a combination of these. The reference standard was imaging (X-ray, CT or MRI), medical record review 21 days later for subsequent imaging and a follow-up call if no imaging was noted to ensure no CSIs were missed. Phillips 2021, a prospective single-centre study, tested the accuracy of PECARN retrospective criteria in direct comparison to the Canadian C-Spine Rule and to the NEXUS criteria in 973 children aged under 16 years who were either immobilised before arrival in the ED for possible CSI, presented with neck pain in the context of trauma or considered at risk of neck injury by the ED team. The reference standard was imaging (X-ray, CT or MRI) or clinical clearance with all children followed up by telephone to ensure no CSIs were missed. The sensitivity of the PECARN retrospective criteria was calculated for Phillips 2021, as it was not provided in the paper. The authors deemed a formal validation not appropriate due to the low incidence of CSI (0.5%).

The sensitivities of the PECARN retrospective criteria in the studies were 0.91 (95% Cl 0.81 to 0.96) (Leonard 2019) and 1.00 (95%

CI 0.48 to 1.00) (Phillips 2021). The specificities of the PECARN retrospective criteria were 0.46 (95% CI 0.44 to 0.47) (Leonard 2019) and 0.32 (95% CI 0.29 to 0.35) (Phillips 2021). We considered the evidence for the estimates of sensitivity and specificity to be moderate certainty, downgraded one level for risk of bias.

#### **PEDSPINE criteria**

Pieretti-Vanmarcke 2009 was a multicentre retrospective study developed and validated the PEDSPINE CSI criteria for children aged three years and younger who sustained blunt trauma. Trauma registries from 22 institutions located in the USA, Canada and Brazil were reviewed over a 10-year period. The total number of participants analysed was 12,537 and the prevalence of CSI was 0.66%. The reference standard was X-rays, CT, MRI or clinical clearance. X-rays were obtained in 32.3% of children, CT in 30.6% and MRI in 3.8%. Imaging was not undertaken in 33.3% of children. Two-thirds of the sample (8354 children) was used to evaluate and develop the PEDSPINE clinical predictors of CSI and one-third (4179 children) was used to validate the criteria. The validation set results were reported. Pieretti-Vanmarcke 2009 was at high risk of bias in the patient selection domain as it was a retrospective study and as the index test was applied retrospectively, after the reference standard was interpreted. It was also rated at high risk of bias in the index test domain.

The sensitivity of the PEDSPINE criteria was 0.93 (95% CI 0.78 to 0.99) and specificity was 0.70 (95% CI 0.69 to 0.72). We considered the evidence for the estimates of sensitivity and specificity to be very low certainty, downgraded three levels for risk of bias and one level for indirectness.

### DISCUSSION

#### Summary of main results

We included five studies evaluating the diagnostic accuracy of seven CDRs (NEXUS, Canadian C-Spine Rule, PECARN retrospective criteria, NICE CG56 and CG176, Leonard de novo and PEDSPINE) to evaluate children with blunt trauma for CSI. One study reported on three index tests (Phillips 2021), two studies reported on two index tests (Davies 2016; Leonard 2019), and two studies reported on one index test (Pieretti-Vanmarcke 2009; Viccellio 2001). The inclusion criteria for age of children were: less than 10 years following blunt trauma (Davies 2016), younger than 18 years (Leonard 2019; Viccellio 2001), under 16 years (Phillips 2021), and three years and younger (Pieretti-Vanmarcke 2009). The total number of enrolled participants in the included studies was 21,379 and ranged from 278 (Davies 2016) to 12,882 (Pieretti-Vanmarcke 2009). The incidence of CSI ranged from 0.5% (Phillips 2021) to 1.85% (Davies 2016). Studies were conducted in the USA (Leonard 2019; Viccellio 2001); the UK (Davies 2016); Australia (Phillips 2021); and across the USA, Canada and Brazil (Pieretti-Vanmarcke 2009). The studies were published between 2001 and 2021.

We assessed the five studies using the four QUADAS-2 risk of bias domains, and assessed the certainty of evidence using the GRADE approach.

For those studies that assessed direct comparisons of CDRs, the evidence for the estimates of sensitivity and specificity for Phillips 2021 (NEXUS, Canadian C-Spine Rule and PECARN retrospective criteria) and Leonard 2019 (PECARN retrospective) were considered moderate certainty, as both studies were downgraded for risk of

bias (unclear risk of bias for the index test and reference standard). The evidence was of low certainty for the new CDR (Leonard de novo) due to risk of bias (unclear risk of bias for the index test and reference standard) and indirectness (no existing validation data). Davies 2016 (NICE guidelines CG56 and CG176) was assessed as very low certainty of evidence, downgraded due to high risk of bias (retrospective study design) and indirectness (only included those children who had imaging of the cervical spine and excluded eight children due to incomplete data).

The additional two indirect comparison studies assessed NEXUS (Viccellio 2001) and PEDSPINE (Pieretti-Vanmarcke 2009). The evidence for the estimates of sensitivity and specificity of NEXUS were low certainty due to risk of bias and inconsistency. Pieretti-Vanmarcke 2009 (PEDSPINE) was assessed as very low-certainty evidence, downgraded due to high risk of bias (retrospective study design and no follow-up) and indirectness (absence of existing validation data).

Since the number of eligible studies for each CDR was fewer than four, and the inclusion criteria and outcomes of those studies were too diverse, we did not conduct a meta-analysis and, therefore, presented no summary estimates in this review. Instead, we interpreted sensitivity and specificity from each primary study separately.

All studies demonstrated high CDR sensitivity in identifying children at risk of CSI (greater than 90%), albeit with relatively wide and varied CIs. However, there is debate about what is considered an acceptable sensitivity, given the potential consequences of missing CSI such as death and lifetime disability. Any CDR considered for clinical use should have a very high sensitivity in detecting people at risk, and ideally narrow CIs. Wider CIs suggest that studies may be underpowered, and in fact two studies are in progress with much larger sample sizes to address this question more accurately (ACTRN12621001050842; NCT05049330). Furthermore, the question remains of what lower sensitivity limit is acceptable for such a potentially devastating condition. The original NEXUS cohort study demonstrated a sensitivity of 1.00, with narrower CIs (95% CI 0.88 to 1.00) (Viccellio 2001); however, other concerns such as the median age of the paediatric population, the small number of injuries under the age of nine years and the potential for missed injuries at younger ages described in some retrospective studies caution interpretation of these findings (Ehrlich 2009; Garton 2008).

Sensitivity and specificity are generally paired outcomes and often inversely associated (i.e. choosing a threshold with higher sensitivity will result in lower specificity). Most included studies described low specificity of the cervical spine CDRs. CDRs for use in the assessment of possible CSI aim to either identify children at higher risk of injury and thus in need of imaging, or children at very low risk for whom imaging can be safely avoided. Imaging itself is not without risks and costs, including exposure to ionising radiation. If CDRs are applied to guide the use of imaging, the lower the specificity, the higher the imaging rate may be, and thus CDRs could actually increase baseline imaging rates without necessarily improving injury detection. This unintended consequence of CDRs has been previously described (Weber 2019). The impact of CDR/ tool use (and CDR specificity) may differ depending on baseline population imaging rates; one Australian study suggested that strict tool use could increase imaging rates from a baseline imaging rate of 41% to between 44% and 68% (Phillips 2021), whereas US data suggest the reverse may apply with Leonard 2019 describing a baseline imaging rate of 78%, and Leonard 2011 calculating that the PECARN retrospective criteria could potentially decrease imaging rates by up to 25%.

Several CDRs have been proposed for use in children to aid in the assessment of possible CSI. While high sensitivity has been reported, findings should be interpreted with caution given the wide CIs. The potential impacts on baseline imaging rates also warrants consideration.

#### Strengths and weaknesses of the review

This is an update of a systematic review on the diagnostic accuracy of CDRs or sets of clinical criteria used to evaluate for CSI following blunt trauma in children. One strength of this review update was the expanded inclusion criterion to include all CDRs compared to the previous review that assessed only NEXUS and the Canadian C-Spine Rule. We performed an extensive search in numerous databases and selected articles using clear inclusion and exclusion criteria. Another strength of this review was that we evaluated the evidence using the QUADAS-2 tool (Whiting 2011). This tool provides important information about potential sources of bias and enables a simple and clear presentation of the assessment.

One of the limitations of our review was that only a few studies were eligible for inclusion. Therefore, we could not conduct sensitivity analyses or formally investigate potential sources of heterogeneity. Only one study tested the accuracy of the NEXUS rule, Canadian C-Spine Rule and PECARN retrospective criteria by direct comparison (Phillips 2021), and the authors identified that low numbers of CSI in a single-centre precluded formal validation of any rule. Therefore, there is limited evidence for which CDR was superior to determine if imaging is indicated in detecting CSI in children following blunt trauma.

Another weakness of this review was that the results were based on a relatively low number of children diagnosed with CSI. A larger sample size would be desirable to better evaluate the accuracy of the CDRs, which should indirectly lead to a higher number of events (children with CSI).

#### Applicability of findings to the review question

The aim of this review was to evaluate if any CDRs are accurate decision tools for detecting CSI in children following blunt trauma.

All the included studies involved a paediatric cohort of participants, with all having a median age under 11 years except the earliest published (Viccellio 2001), which had a median age of 15 years. Two studies focused on the younger ages; Pieretti-Vanmarcke 2009 only included children aged under three years and Davies 2016 included children under 10 years. The inclusion of younger children is important because adolescent injuries are generally considered to follow a more adult pattern, and the greatest clinical concern of the applicability and accuracy of adult-derived CDRs such as Canadian C-Spine Rule and NEXUS, and thus the potential for missed injuries, exists at younger ages (Ehrlich 2009;Garton 2008).

All studies were conducted in largely well-resourced trauma or tertiary EDs across several countries. Resource access constraints may influence the threshold for imaging and CDR application.



Inclusion criteria also varied between studies; two included only those already considered at higher risk by ED clinicians (receiving imaging) (Davies 2016; Viccellio 2001); others included a broader population of ED participants. This is an important consideration when applying study results outside their original populations, as imaging rates vary across both time and healthcare system, depending on perceived risks of the prevalence of the condition, radiation exposure and medicolegal concerns (Babl 2017; Leonard 2019; Phillips 2021).

On the strength of currently available evidence, caution is advised with strictly applying the considered CDRs in practice to children. Future studies may offer better clarity. There are two ongoing studies; NCT05049330, which will validate a Pediatric CSI Risk Assessment Tool in more than 20,000 children younger than 18 years of age and ACTRN12621001050842 will validate three CDRs (NEXUS, Canadian C-Spine Rule and latest published PECARN criteria) in children aged less than 16 years with possible CSI after known or suspected blunt trauma in a large multicentre population.

# AUTHORS' CONCLUSIONS

#### Implications for practice

Currently, there is insufficient evidence to determine which clinical decision rule (CDR) is the most accurate in detecting cervical spine injuries (CSIs) in children following blunt trauma and those available are at best a guide to clinical assessment. Current evidence does not support strict or protocolised adoption of any of the CDRs in paediatric trauma care. Although most studies had a high sensitivity, this was often achieved at the expense of specificity (and resulting high imaging rates). The specificities of the CDRs were generally low, ranging from 0.07 to 0.70. The main goal of CDRs is to identify all CSIs whilst minimising unnecessary imaging tests. Therefore, sensitivity needs to be high; the challenge is in improving specificity and maintaining high sensitivity. Data on children under the age of eight years of age are particularly sparse; therefore, there is currently no strong evidence to support the use of these CDRs in this age group. Although Pieretti-Vanmarcke 2009 assessed the accuracy of PEDSPINE for children aged three years and younger who sustained blunt trauma, as a retrospective study it

was at high risk of bias in the patient selection domain as the index test was applied retrospectively, after the reference standard was interpreted and it was at high risk of bias in the index test domain.

#### Implications for research

Since the incidence of CSIs in children is low, a large cohort is needed to test the accuracy of CDRs. Hence, future research should focus on large adequately powered multicentre prospective trials to assess the accuracy of CDRs in children. Only then can we determine whether they are sufficiently sensitive and specific to be applied as a decision tool following blunt trauma. It would be important to include enough children younger than eight years of age to ensure the decision tools could be used in children of all ages. Also, there should be an adequate number of CSI events. Although PEDSPINE focuses on children aged three years and younger, this will need to be assessed prospectively. A study should optimally evaluate CDRs in direct comparison to others in all paediatric trauma populations. It is important that children are clinically followed up if radiographic imaging was not conducted to reduce the likelihood of missing CSIs. In planning the study, trialists should conduct a power analysis to determine how many children younger and older than eight years of age should be included, and how many events (children with CSI) would be required. There are currently two large ongoing multicentre prospective studies that should contribute to the evidence base of the accuracy of CDRs in children. NCT05049330 will validate a Pediatric CSI Risk Assessment Tool (derived in Leonard 2019) in children younger than 18 years of age and ACTRN12621001050842 will validate three CDRs (National Emergency X-Radiography Utilization Study (NEXUS), Canadian C-Spine Rule and the latest published Pediatric Emergency Care Applied Research Network (PECARN) criteria) in children aged less than 16 years.

# ACKNOWLEDGEMENTS

We thank Poh Chua, Librarian at the Royal Children's Hospital Melbourne, for her help with the modification of the search strategy and literature search for the review update. We thank Anne Lawson, Cochrane Central Production Service, for copy-editing the final review and the review author team from the previous version of the review for their contribution to the protocol development and review findings.

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Davies 2016

Slaar A, Fockens MM, Wang J, Maas M, Wilson DJ, Goslings JC, et al. Triage tools for detecting cervical spine injury

# CHARACTERISTICS OF STUDIES

# **Characteristics of included studies** [ordered by study ID]

in pediatric trauma patients. *Cochrane Database of Systematic Reviews* 2017, Issue 12. Art. No: CD011686. [DOI: 10.1002/14651858.CD011686.pub2]

\* Indicates the major publication for the study

Study characteristics			
Patient Sampling	Level 1 trauma centre, Royal London Hospital, UK		
	Study dates: 1 October 2008 to 1 October 2013		
	<b>Sampling:</b> retrospective cohort (medical record review) – children aged < 10 years who underwent cervical spine imaging following blun trauma		
Patient characteristics and setting	<b>Inclusion criteria:</b> children aged < 10 years presenting following bl trauma and who had any imaging of the cervical spine (X-ray, CT, MI or a combination of these) within 24 hours of admission to exclude CSI. Children with incomplete data were excluded (8/278)		
	Participant characteristics		
	Children enrolled: 278		
	Children included in analysis: 270 (8 excluded due to incomplete data		
	Mean age: 5.2 years (range 0.2–9.9 years)		
	Sex: 180 boys, 90 girls		
	Children with CSI: 5 (1.85%)		
	Setting: level 1 trauma centre, London UK		
Index tests	<b>Index tests:</b> NICE guideline 56 (CG56) and NICE guideline 176 (CG176) – see Table 3 for details		
	Test administrator/training: not stated		
	Blinding of examiners: not stated		
Target condition and reference standard(s)	Target condition: CSI (no definition provided)		
	<b>Reference standard:</b> X-ray, MRI, CT, or a combination of these within 24 hours of admission		
Flow and timing	Time between presentation to ED with blunt trauma and imaging was assumed to be < 1 day. 8 children were excluded due to incomplete data. A total of 68 children had a cervical spine X-ray of which 44 (64% were reported as technically inadequate; of these 5 had a subsequent CT. 6 cases with adequate X-rays underwent subsequent CT and 1 had a subsequent MRI		



#### Davies 2016 (Continued)

Notes

Funding: not stated

# Methodological quality

ltem	Authors' judgement	Risk of bias	Applicability con- cerns
DOMAIN 1: Patient selection			
Was a consecutive or random sample of patients enrolled?	Yes		
Did the study avoid inappropriate exclusions?	Yes		
Prospective design	No		
Could the selection of patients have introduced bias?		High risk	
Are there concerns that the included patients and set- ting do not match the review question?			High
DOMAIN 2: Index test (All tests)			
Were the index test results interpreted without knowledge of the results of the reference standard?	No		
If a threshold was used, was it pre-specified?	Yes		
Could the conduct or interpretation of the index test have introduced bias?		High risk	
Are there concerns that the index test, its conduct, or in- terpretation differ from the review question?			Unclear
DOMAIN 3: Reference standard			
Is the reference standard likely to correctly classify the tar- get condition?	Unclear		
Were the reference standard results interpreted without knowledge of the results of the index tests?	Yes		
Could the reference standard, its conduct, or its inter- pretation have introduced bias?		Unclear risk	
Are there concerns that the target condition as defined by the reference standard does not match the question?			Unclear
DOMAIN 4: Flow and timing			
Did all patients receive the same reference standard?	Yes		
Were all patients included in the analysis?	Yes		
Was there an appropriate interval between presentation to ED with blunt trauma and conduct of the index test and ref- erence standard?	Yes		



# Davies 2016 (Continued)

Could the patient flow have introduced bias?

Low risk

Study characteristics			
Patient Sampling	4 tertiary care children's hospitals in the USA		
	Study dates: March 2014 to November 2016		
	Sampling: prospective, consecutive sampling		
Patient characteristics and setting	<b>Inclusion criteria:</b> children aged < 18 years who presented to the ED for blunt trauma and were transported from the scene of injury by emergency medical services in spinal motion restriction devices (cervical collars and rigid longboards), underwent trauma team evaluation, had cervical spine imaging ordered in the ED or a combination of these		
	<b>Exclusion criteria:</b> children whose injury mechanism was solely penetrating trauma, whose legal guardian had a substantial language barrier, who were in state's custody, or who were transferred from the study site for definitive care		
	Participant characteristics		
	Children enrolled: 4144		
	Children included in analysis: 4091 (legal guardians withdrew 53 children)		
	Mean age of cohort: 9.4 years		
	Mean age of those with CSI: 10.7 years		
	Sex: 2373 boys, 1718 girls		
	Children with CSI: 74 (1.8%)		
	Children aged < 8 years: 1609 (39.3%); 23 (1.4%) had CSI		
	Setting: tertiary care children's hospitals, USA		
Index tests	<b>Index tests:</b> Pediatric Emergency Care Applied Research Network (PECARN) retrospectiv criteria and Leonard de novo model – see Table 3 for details		
	<b>Test administrator/training:</b> trained research personnel administered electronic branch-logic questionnaires to treating ED providers		
	<b>Blinding of examiners:</b> observations were gained regarding CSI risk factors before knowledge of cervical spine imaging results, if ordered or if obtained at a transferring hospital, before knowledge of their institutional radiologist's interpretation		
Target condition and reference stan- dard(s)	<b>Target condition:</b> CSI defined as vertebral fractures, ligamentous injury, intraspinal haemorrhage, or spinal cord injury (either on MRI or spinal cord injury without radiographic association) involving the cervical region (occiput to seventh cervical vertebra, including ligamentous structures attaching the seventh cervical vertebra to first thoracia vertebra)		
	<b>Reference standard:</b> cervical spine imaging reports and if applicable, spine surgeon consultation notes. If the imaging report conflicted with the spine surgeon consultation, cla ification was sought. For children who did not undergo imaging, the medical record was reviewed 21 days later for subsequent imaging and if no imaging was noted, a follow-up		



Leonard 2019 (Continued)	call with a legal guardian v sence of CSI	was conducted 21–28 days	after the ED visit to verify the ab-
Flow and timing	Time between presentation with blunt trauma to ED and imaging was assumed to be < 1 day		
Comparative			
Notes		Development grant R21HDC	nnedy Shriver National Institute of 076108-02. Funded by the National
Methodological quality			
Item	Authors' judgement	Risk of bias	Applicability concerns
DOMAIN 1: Patient selection			
Was a consecutive or random sample of patients enrolled?	Yes		
Did the study avoid inappropriate exclu- sions?	Yes		
Prospective design	Yes		
Could the selection of patients have introduced bias?		Low risk	
Are there concerns that the included patients and setting do not match the review question?			Low concern
DOMAIN 2: Index test (All tests)			
Were the index test results interpreted without knowledge of the results of the reference standard?	Unclear		
If a threshold was used, was it pre-spec- ified?	Yes		
Could the conduct or interpretation of the index test have introduced bias?		Unclear risk	
Are there concerns that the index test, its conduct, or interpretation differ from the review question?			High
DOMAIN 3: Reference standard			
Is the reference standard likely to cor- rectly classify the target condition?	Yes		
Were the reference standard results in- terpreted without knowledge of the re- sults of the index tests?	Unclear		



Leonard 2019 (Continued)	
Could the reference standard, its con- duct, or its interpretation have intro- duced bias?	Unclear risk
Are there concerns that the target condition as defined by the reference standard does not match the ques- tion?	Low concern
DOMAIN 4: Flow and timing	
Did all patients receive the same refer- ence standard?	Yes
Were all patients included in the analy- sis?	Yes
Was there an appropriate interval be- tween presentation to ED with blunt trauma and conduct of the index test and reference standard?	Yes
Could the patient flow have intro- duced bias?	Low risk

# Phillips 2021

Study characteristics	
Patient Sampling	Single-centre tertiary paediatric ED in Brisbane, Australia
	Study dates: September 2015 to September 2016
	Sampling: prospective, consecutive sampling
Patient characteristics and setting	<b>Inclusion criteria:</b> aged < 16 years and met ≥ 1 of the following: immobilisation prearrival for possible CSI; presentation with neck pain in the context of trauma; otherwise considered at risk of neck injury by the ED team (e.g. multitrauma patient or trauma patient with abnormal neurology, posturing or altered consciousness level)
	<b>Exclusion criteria:</b> declined to participate, did not wait to be seen or a successful fol- low-up telephone call was viewed unlikely (e.g. overseas resident, no easily identifiable guardian, transient living situation, insufficient English language). Children assessed by ED clinicians as having had their cervical spines fully assessed and cleared at another hospital prior to transfer for the definitive management of other injuries were excluded
	Participant characteristics
	Children enrolled: 1010 (37 excluded as had initial imaging prior to arrival of which 9 had confirmed CSI)
	Children included in analysis: 973
	Median age: 10.9 years (IQR 7.1–13.6 years)
	Sex: 643 boys, 330 girls
	Children with CSI: 5 (0.5%)



Phillips 2021 (Continued)	Children aged < 9 years 20	ne (20, 204)	
	Children aged < 8 years: 29		
	Children aged < 2 years: 46		
	Setting: tertiary paediatri	c ED, Australia	
Index tests	Index tests: NEXUS criteria, Canadian C-Spine Rule and PECARN rule – see Table 3 for de- tails		
	"clinician interpreted crite points: prospectively by cl	ria" – no training reported inicians based on informat	prospectively by clinicians using Variables were collected at 2 time ion available during initial assessmen lete clinical notes were available to as
		diologists at each site inter vere used for assessing inde	rpreted all radiographic images and ex test criteria
Target condition and reference stan- dard(s)	Target condition: CSI defi specialist paediatric radio		I on X-ray, CT or MRI as reported by
		y, CT or MRI or clinical clear children to ensure no CSIs v	rance if no imaging sought. Telephone were missed
Flow and timing	day. 396 (40.7%) children l ceived X-ray, 130 (13.5%) C	nad their cervical spine ima T and 29 (3%) MRI. 577 (59 telephone follow-up and v	and imaging was assumed to be < 1 Iged after ED arrival; 315 (32.4%) re- .3%) children did not receive imaging; vere included in the analysis. There
	X-ray and sometimes a CT	scan were obtained as the	primary imaging modality
Comparative			
Notes		mergency Medicine Found . NP, JA, RB, GA and MW ob	ation (Australasia) Queensland Pro- tained grant funding
Methodological quality			
Item	Authors' judgement	Risk of bias	Applicability concerns
DOMAIN 1: Patient selection			
Was a consecutive or random sample of patients enrolled?	Yes		
Did the study avoid inappropriate ex- clusions?	Yes		
Prospective design	Yes		
Could the selection of patients have introduced bias?		Low risk	
Are there concerns that the includ- ed patients and setting do not match the review question?			Low concern



Phillips 2021 (Continued)			
Were the index test results interpret- ed without knowledge of the results of the reference standard?	Unclear		
If a threshold was used, was it pre- specified?	Yes		
Could the conduct or interpretation of the index test have introduced bias?		Unclear risk	
Are there concerns that the index test, its conduct, or interpretation differ from the review question?			Low concern
DOMAIN 3: Reference standard			
Is the reference standard likely to cor- rectly classify the target condition?	Yes		
Were the reference standard results in- terpreted without knowledge of the re- sults of the index tests?	Unclear		
Could the reference standard, its conduct, or its interpretation have introduced bias?		Unclear risk	
Are there concerns that the target condition as defined by the refer- ence standard does not match the question?			Low concern
DOMAIN 4: Flow and timing			
Did all patients receive the same refer- ence standard?	Yes		
Were all patients included in the analy- sis?	Yes		
Was there an appropriate interval be- tween presentation to ED with blunt trauma and conduct of the index test and reference standard?	Yes		
Could the patient flow have intro- duced bias?		Low risk	

Pieretti-Vanmarcke 2009

Study characteristics

Patient Sampling

Trauma registries of 22 institutions in the USA, Canada and Brazil. There were 15 paediatric level I, 6 adult level I, and 1 adult level II trauma centres

Study dates: 1 January 1995 to 1 January 2005		
<b>Sampling:</b> retrospective sampling of children aged ≤ 3 years after trauma identified in the paediatric trauma registries. Sample rand split into 2 data sets: 2/3 of the sample was used to identify clinic. tors of CSI to develop a scoring algorithm (evaluation set) and 1/3 date the algorithm (validation set)	domly al predic-	
<b>Inclusion criteria:</b> children aged < 3 years who sustained blunt tr	auma	
Participant characteristics		
Total children enrolled: 12,882		
Total children included in analysis: 12,537 (345 children were exc they died immediately upon presentation to the ED)	luded as	
Mean age: 1.4 (SD 0.8) years		
Sex: 7463 boys, 5074 girls		
Children with CSI: 83 (0.66%)		
Validation set		
Total children in analysis: 4179		
Children with CSI: 30 (0.72%)		
<b>Setting:</b> paediatric and adult, level I and level II hospitals in the U Canada and Brazil	ISA,	
Index tests: PEDSPINE – see Table 3 for details		
Test administrator/training: not stated		
Blinding of examiners: not stated		
<b>Target condition:</b> CSI defined by any osseous or ligamentous injucervical spine seen on CT, X-ray or MRI	ury to the	
Reference standard: X-ray, CT or MRI or clinical clearance		
Time between presentation to ED with blunt trauma and imaging was as- sumed to be < 1 day. X-rays (2 or 3 views) were obtained in 4046 children (32.3%), CT in 3358 (30.6%) and MRI in 478 (3.8%)		
<b>Funding:</b> grants from the American Association for the Surgery of Foundation and Anthem Blue Cross/Blue Shield of Connecticut	Trauma	
We contacted the authors to request additional data on participa acteristics	nt char-	
_	Sampling: retrospective sampling of children aged ≤ 3 years after         trauma identified in the paediatric trauma registries. Sample ransplit into 2 data sets: 2/3 of the sample was used to identify clinic         tors of CSI to develop a scoring algorithm (evaluation set) and 1/3         date the algorithm (validation set)         Inclusion criteria: children aged < 3 years who sustained blunt tr	

Pieretti-Vanmarcke 2009 (Continued)			
Was a consecutive or random sample of patients en- rolled?	Yes		
Did the study avoid inappropriate exclusions?	Yes		
Prospective design	No		
Could the selection of patients have introduced bias?		High risk	
Are there concerns that the included patients and setting do not match the review question?			Low concern
DOMAIN 2: Index test (All tests)			
Were the index test results interpreted without knowl- edge of the results of the reference standard?	No		
If a threshold was used, was it pre-specified?	Yes		
Could the conduct or interpretation of the index test have introduced bias?		High risk	
Are there concerns that the index test, its conduct, or interpretation differ from the review question?			High
DOMAIN 3: Reference standard			
Is the reference standard likely to correctly classify the target condition?	No		
Were the reference standard results interpreted with- out knowledge of the results of the index tests?	Yes		
Could the reference standard, its conduct, or its in- terpretation have introduced bias?		High risk	
Are there concerns that the target condition as de- fined by the reference standard does not match the question?			Low concern
DOMAIN 4: Flow and timing			
Did all patients receive the same reference standard?	No		
Were all patients included in the analysis?	Yes		
Was there an appropriate interval between presenta- tion to ED with blunt trauma and conduct of the index test and reference standard?	Yes		
Could the patient flow have introduced bias?		High risk	



Study characteristics			
Patient Sampling	UCLA Emergency Medicine Center (USA) and 20 participating centres. Study sites comprise a range of acute care facilities, including academic trauma centres, community trauma centres, and community EDs		
	Study dates: 1990–2000 (estimated)		
	Sampling: prospective multicentre cohort study		
Patient characteristics and setting	<b>Inclusion criteria:</b> children aged < 18 years with blunt trauma injure received cervical spine imaging	ıries wl	
	Participant characteristics		
	Total adults and children enrolled: 34,069		
	Children aged < 18 years: 3065 (all included in analysis)		
	Age: 0–2 years (88 children, 2.9%), 2–8 years (817 children, 26.7%) years (2160 children, 70.5%)	9–17	
	Sex: not stated		
	Children with CSI: 30 (0.98%)		
	Children aged < 8 years with CSI: 4		
	Setting: acute care facilities in the USA		
Index tests	Index tests: NEXUS criteria – see Table 3 for details		
	<b>Test administrator/training:</b> physicians at participating centres took brief training programmes	under-	
	<b>Blinding of examiners:</b> study radiologists at each site interpreted ographic studies. Neither the official radiology interpretation nor ing of injuries was performed with knowledge of the findings on th US data form	the coo	
Target condition and reference standard(s)	Target condition: CSI defined as cervical spine fracture or dislocation	tion	
	Reference standard: X-ray, CT, MRI, or a combination of these		
Flow and timing	Time between presentation to ED with blunt trauma and imaging was assumed to be < 1 day. Only those children who were selected for radi- ographic imaging were included. X-rays and CT scan were the primary imaging modalities		
Comparative			
Notes	Analysis of the paediatric population included in the original Hoff 2000, the seminal publication of the NEXUS tool	man	
	<b>Funding:</b> Grant R01 HS08239 from the Agency for Healthcare Rese and Quality	arch	
Methodological quality			
Item	Authors' judgement Risk of bias Applicability cerns	con-	

DOMAIN 1: Patient selection			
Was a consecutive or random sample of patients en- rolled?	Yes		
Did the study avoid inappropriate exclusions?	Yes		
Prospective design	Yes		
Could the selection of patients have introduced bias?		Low risk	
Are there concerns that the included patients and setting do not match the review question?			Low concern
DOMAIN 2: Index test (All tests)			
Were the index test results interpreted without knowl- edge of the results of the reference standard?	Unclear		
If a threshold was used, was it pre-specified?	Yes		
Could the conduct or interpretation of the index test have introduced bias?		Unclear risk	
Are there concerns that the index test, its conduct, or interpretation differ from the review question?			Low concern
DOMAIN 3: Reference standard			
Is the reference standard likely to correctly classify the target condition?	Yes		
Were the reference standard results interpreted with- out knowledge of the results of the index tests?	Yes		
Could the reference standard, its conduct, or its in- terpretation have introduced bias?		Low risk	
Are there concerns that the target condition as de- fined by the reference standard does not match the question?			Low concern
DOMAIN 4: Flow and timing			
Did all patients receive the same reference standard?	Yes		
Were all patients included in the analysis?	Yes		
Was there an appropriate interval between presenta- tion to ED with blunt trauma and conduct of the index test and reference standard?	Yes		
Could the patient flow have introduced bias?		Low risk	



CSI: cervical spine injury; CT: computed tomography; ED: emergency department; IQR: interquartile range; MRI: magnetic resonance imaging; NEXUS: National Emergency X-Radiography Utilization Study; NICE: National Institute for Health and Care Excellence; PECARN: Pediatric Emergency Care Applied Research Network; SD: standard deviation; UCLA: University of California, Los Angeles.

## Characteristics of excluded studies [ordered by study ID]

Study	Reason for exclusion
Adelgais 2014	Index test not relevant
Ahmad 2017	Index test not relevant
Al-Sarheed 2020	Index test not relevant
Alas 2021	Index test not relevant
Anderson 2006	Target condition not relevant
Anderson 2010	Index test not relevant
Atesok 2018	Ineligible study design (review)
Babcock 2018	Index test not relevant
Babu 2016	Different study population (only people with CSI)
Bailey 2022	Ineligible study design (insufficient evidence to present a 2 × 2 table on the accuracy of the CDR)
Baker 1999	Index test not relevant
Bandiera 2003	Different study population (adults)
Banit 2000	Index test not relevant
Bayless 1989	Index test not relevant
Benayoun 2016	Different study population (mostly adults; data on children not reported separately)
Bennett 2015	Index test not relevant
Blacksin 1995	Index test not relevant
Boese 2015	Index test not relevant
Borock 1991	Different study population (only people with CSI)
Boustani 2015	Different study population (adults)
Brockmeyer 2012	Different study population (suspected or confirmed CSI)
Brooks 2001	Index test not relevant
Brown 2001	Index test not relevant
Browne 2003	Target condition not relevant
Browne 2017	Index test not relevant



Study	Reason for exclusion
Browne 2021	Different study population (prehospital)
Burns 2011	Different target condition
Caltili 2017	Different study population (mostly adults). Contacted authors for data on children but they did not respond.
Carter 2017	Different study population (suspected or confirmed CSI)
Chaudhry 2016	Index test not relevant
Clayton 2012	Index test not relevant
Coffey 2011	Different study population (adults)
Como 2011	Index test not relevant
Cui 2016	Index test not relevant
Dahlquist 2013	Index test not relevant
Dalle 2022	Index test not relevant
Dickinson 2004	Different study population (adults)
DiGiacomo 2002	Index test not relevant
Douglas 2022	Ineligible study design (study did not compare the accuracy of the cervical spine clearance guide- line to the reference standard)
Dranoff 2019	Different study population (adults)
Dwyer 2019	Different study population (suspected or confirmed CSI)
Edwards 2001	Index test not relevant
Ehrlich 2009	Ineligible study design; retrospective case-matched design
Ekhator 2022	Ineligible study design
Eren 2020	Index test not relevant
Ersoy 1995	Index test not relevant
Fischer 1984	Index test not relevant
Flynn-O'Brien 2016	Index test not relevant
Gajera 2017	Ineligible study design (review)
Garton 2008	Different study population (only people with CSI)
Gbaanador 1986	Index test not relevant
Ghelichkhani 2021	Different study population (adults)



Study	Reason for exclusion
Gonzalez 1999	Index test not relevant
Gonzalez 2009	Index test not relevant
Gonzalez 2013	Different study population (prehospital evaluation)
Griffen 2003	Index test not relevant
Griffith 2011	Different study population (adults)
Griffith 2013	Different study population (adults)
Griffith 2014	Different study population (adults)
Hale 2015	Index test not relevant
Handler 2018	Index test not relevant
Hannon 2015	Index test not relevant
Hanson 2000a	Different study population (adults)
Hanson 2000b	Different study population (adults)
Hasan 2022	Index test not relevant
Hasler 2011	Different study population (adults)
Hazboun 2021	Index test not relevant
Heffernan 2005	Different study population (adults)
Henry 2016	Index test not relevant
Henry 2021	Index test not relevant
Herman 2019	Ineligible study design (review)
Hoffman 1992	Authors did not separate data for children from adults (only a few children were included)
Hollingshead 2000	Index test not relevant
Hood 2015	Ineligible study design (review)
Hopper 2020	Index test not relevant
Hutchings 2009	Index test not relevant
Ihalainen 2017	Index test not relevant
Inaba 2015	Different study population (adults)
Inaba 2016	Target condition not relevant
Jacob 2016	Index test not relevant



Study	Reason for exclusion							
Jaffe 1987	Ineligible study design; recruited a second non-consecutive cohort of CSI cases							
Jakes 2015	Ineligible study design (review)							
Jarvers 2020	Index test not relevant							
Kadom 2019	Ineligible study design (review)							
Kaminski 2017	Index test not relevant							
Kavuri 2019	Index test not relevant							
Keenan 2001	Index test not relevant							
Kerr 2005	Different study population (adults)							
Khetarpal 2021	Ineligible participant population (prehospital)							
Kokabi 2011	Different study population (adults)							
Lee 2003	Index test not relevant							
Lee 2022	Index test not relevant							
Lemley 2015	Ineligible study design (review)							
Leonard 2011	Ineligible study design (case control study)							
Letica-Kriegel 2022	Ineligible study design (review)							
Liawrungrueang 2020	Ineligible study design (case study)							
Luehmann 2020	Index test not relevant							
Malomo 1995	Index test not relevant							
Mannix 2011	Index test not relevant							
Markuske 1983	Target condition not relevant							
Markuske 1988	Target condition not relevant							
Martin 2004	Index test not relevant							
McLaughlin 2019	Ineligible study design (review)							
McMahon 2015	Index test not relevant							
Meek 2007	Index test not relevant							
Meldon 1998	Different study population (out-of-hospital participants)							
Mitrofan 2016	Ineligible study design (review)							
Morrison 2012	Index test not relevant							



Study	Reason for exclusion							
Mower 2001	Index test not relevant							
NCT05605847	Different study population (adults)							
Neifeld 1988	Different study population (adults)							
Nguyen 2005	Index test not relevant							
Nolte 2022	Different study population (prehospital)							
Novick 2018	Index test not relevant							
Nunn 2021	Index test not relevant							
Omran 2001	Index test not relevant							
Overberger 2018	Different study population (adults)							
Overmann 2020	Index test not relevant							
Pannu 2017	Index test not relevant							
Pennell 2020	Index test not relevant							
Pepin 2015	Ineligible study design (full text unavailable). Contacted authors but they did not reply.							
Platzer 2006a	Index test not relevant							
Platzer 2006b	Index test not relevant							
Poorman 2019	Index test not relevant							
Pulfrey 2002	Different study population (adults)							
Quigley 2014	Index test not relevant							
Raza 2013	Different study population (adults)							
Robinson 2022	Index test not relevant							
Rolfe 2019	Target condition not relevant							
Ropele 2009	Index test not relevant							
Rosati 2015	Index test not relevant							
Rose 2012	Index test not relevant							
Ross 1987	Index test not relevant							
Saddison 1991	Different study population (only people with CSI)							
Sanchez 2005	Index test not relevant							
Scarrow 1999	Different study population (only people with CSI)							



Study	Reason for exclusion								
Schleehauf 1989	Index test not relevant								
Sharma 2023	Ineligible study design (study did not compare the accuracy of a CDR to the reference standard)								
Sheikh 2012	Different study population (adults)								
Shin 2016	Index test not relevant								
Singh 2018	Index test not relevant								
Slaar 2016	Index test not relevant								
Smart 2003	Index test not relevant								
Sokoloff 2022	Different study population								
Songür Kodik 2020	Different study population (mostly adults; data on children not extractable)								
Stanton 2017	Ineligible study design (review)								
Stiell 2003	Different study population (adults)								
Stiell 2009	Different study population (adults)								
Stiell 2010	Different study population (adults)								
Stiell 2018	Different study population (mostly adults; data on children not extractable)								
Stroh 2001	Index test not relevant								
Sun 2013	Target condition not relevant								
Syrmos 2015	Ineligible study design (case study)								
Tahvonen 2013	Index test not relevant								
Ten Brinke 2021	Index test not relevant.								
Tricks 2019	Full text unavailable. Contacted the authors who explained that no full-text studies had been pub- lished								
Vaillancourt 2017	Different study population (prehospital)								
Vaillancourt 2020	Different study population (prehospital)								
Valusek 2010	Index test not relevant								
Velmahos 1996	Index test not relevant								
Vittetoe 2022	Index test not relevant								
Waddell 2018	Target condition not relevant								
Zebracki 2022	Index test not relevant.								



#### Study

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Reason for exclusion

Özkan 2015

Different study population (suspected or confirmed CSI)

CDR: clinical decision rule; CSI: cervical spine injury.

# Characteristics of studies awaiting classification [ordered by study ID]

Arbuthnot 2017						
Patient Sampling	Level 1 paediatric trauma centre (Boston Children's Hospital), USA					
	Retrospective review					
Patient characteristics and setting	Inclusion criteria: people aged ≤ 21 years with blunt trauma injury who underwent cervical spine evaluation					
Index tests	Boston Children's Hospital paediatric cervical spine clearance algorithm					
Target condition and refer-	Target condition: cervical spine injury (no definition provided)					
ence standard(s)	Reference standard: X-ray, MRI, CT, or a combination of these within 24 hours of admission					
Flow and timing	Time between presentation to ED with blunt trauma and imaging was assumed to be < 1 day. 1 missed injury					
Comparative						
Notes	Authors contacted for data for children aged 0 to < 18 years (inclusion criteria for the review)					

Vargas 2022						
Patient Sampling	3 level 1 paediatric trauma centres, USA					
Patient characteristics and setting	Children aged < 8 years with traumatic injury admitted to 1 of 3 level 1 paediatric trauma centres between August 2007 and August 2017					
Index tests	Previously identified 6 risk factors that increased the odds of having a CSI in children aged < 8 years					
Target condition and refer- ence standard(s)	<ul> <li>Target condition: CSI defined as</li> <li>radiographic evidence of CSI or</li> <li>radiographic evidence of CSI or treatment due to clinical concern, or both</li> <li>Reference standard: radiography</li> </ul>					
Flow and timing	Time between presentation to ED with blunt trauma and imaging: not reported (abstract)					
Comparative						
Notes	Authors contacted requesting full-text article to assess study for eligibility.					

CT: computed tomography; ED: emergency department; MRI: magnetic resonance imaging.

# Characteristics of ongoing studies [ordered by study ID]

#### ACTRN12621001050842

Study name	Study Of Neck Injuries In Children (SONIC)					
Target condition and refer- ence standard(s)	Target condition: cervical spine injury Reference standard: imaging or follow-up (or both)					
Index and comparator tests	NEXUS, Canadian C-Spine Rule and PECARN					
Starting date	Ethics: 9 April 2021					
Contact information	Dr Natalie Phillips Email: natalie.phillips@health.qld.gov.au					
Notes	11 sites and estimated completion date: 30 December 2025 Australia and New Zealand Clinical Registry: ACTRN12621001050842					

### NCT05049330

Study name	Development and testing of a pediatric cervical spine injury risk assessment tool (C-Spine)					
Target condition and refer- ence standard(s)	Target condition: cervical spine injury Reference standard: imaging and follow-up					
Index and comparator tests	Pediatric CSI Risk Assessment Tool					
Starting date	12 December 2018					
Contact information	Email: Julie.Leonard@Nationwidechildrens.org					
Notes	18 centres					
	Estimated completion date: 1 September 2023					
	ClinicalTrials.gov Identifier: NCT05049330					

NEXUS: National Emergency X-Radiography Utilization Study; PECARN: Pediatric Emergency Care Applied Research Network.

# DATA

Presented below are all the data for all of the tests entered into the review.

## Table Tests. Data tables by test

Test	No. of studies	No. of participants
1 NEXUS	2	4038



Test	No. of studies	No. of participants
2 Canadian C-Spine Rule	1	973
3 PECARN Retrospective	2	5064
4 Leonard de novo	1	4091
5 PEDSPINE	1	4179
6 NICE guideline (CG56)	1	270
7 NICE guideline (CG176)	1	270

## Test 1. NEXUS

Study	ТР	FP	FN	TN	Sensitivity (95% CI)	Specificity (95% CI)	Sensitivity (95% CI)	Specificity (95% CI)
Phillips 2021	5	425	0	543	1.00 [0.48, 1.00]	0.56 [0.53, 0.59]		
Viccellio 2001	30	2432	0	603	1.00 [0.88, 1.00]	0.20 [0.18, 0.21]		• • • •
							0 0.2 0.4 0.6 0.8 1 0	0.2 0.4 0.6 0.8 1

# Test 2. Canadian C-Spine Rule

Study	ТР	FP	FN	TN	Sensitivity (95% CI)	Specificity (95% CI)		Sen	sitivity	<b>(9</b> 5%	CI)			Spee	cificity	(95%	CI)	
Phillips 2021	5	467	0	501	1.00 [0.48, 1.00]	0.52 [0.49, 0.55]						-				•		
							0	0.2	0.4	0.6	0.8	1	0	0.2	0.4	0.6	0.8	1

# Test 3. PECARN Retrospective

Study	ТР	FP	FN	TN	Sensitivity (95% CI)	Specificity (95% CI)	Sensitivity (95% CI) Specificity (95% CI)
Leonard 2019	67	2186	7	1831	0.91 [0.81, 0.96]	0.46 [0.44, 0.47]	-+ -
Phillips 2021	5	658	0	310	1.00 [0.48, 1.00]	0.32 [0.29, 0.35]	<del></del> <b>.</b>
							0 0.2 0.4 0.6 0.8 1 0 0.2 0.4 0.6 0.8 1

#### Test 4. Leonard de novo

Study	ТР	FP	FN	TN	Sensitivity (95% CI)	Specificity (95% CI)	Sensit	tivity	(95%	CI)			Spe	cificity	y <b>(9</b> 5%	CI)	
Leonard 2019	68	1998	6	2019	0.92 [0.83, 0.97]	0.50 [0.49, 0.52]				-	-						
							0.2	0.4	0.6	0.8	1	0	0.2	0.4	0.6	0.8	

# Test 5. PEDSPINE

Study	ТР	FP	FN	TN	Sensitivity (95% CI)	Specificity (95% CI)		Sens	sitivity	(95%	CI)			Spe	ificity	y <b>(9</b> 5%	GCI)	
Pieretti-Vanmarcke 2009	28	1224	2	2925	0.93 [0.78, 0.99]	0.70 [0.69, 0.72]						•					•	
							0	0.2	0.4	0.6	0.8	1	0	0.2	0.4	0.6	0.8	1

# Test 6. NICE guideline (CG56)

Study	ТР	FP	FN	TN	Sensitivity (95% CI)	Specificity (95% CI)	Sensitivity (95% CI) Specificity (95% CI)
Davies 2016	5	144	0	121	1.00 [0.48, 1.00]	0.46 [0.40, 0.52]	
							0 0.2 0.4 0.6 0.8 1 0 0.2 0.4 0.6 0.8 1

# Test 7. NICE guideline (CG176)

Study	ТР	FP	FN	TN	Sensitivity (95% CI)	Specificity (95% CI)		Sen	sitivity	y <b>(9</b> 5%	CI)			Spe	cificity	y <b>(9</b> 5%	o CI)	
Davies 2016	5	246	0	19	1.00 [0.48, 1.00]	0.07 [0.04, 0.11]			-			-	4	F.				
							-					_						
							Ò	0.2	0.4	0.6	0.8	1	Ò	0.2	0.4	0.6	0.8	1

# ADDITIONAL TABLES

# Table 1. Assessment of methodological quality: QUADAS-2 and additional items

Quality item	Risk of bias		Applicability					
	Quality indicator	Notes	Quality indica- tor	Notes				
Domain 1	Could the selection	of participants have introduced bias?	Are there concerr					
Patient selec- tion	(high/low/unclear)		ed patients and settings do not match the review question? (high/ low/unclear)					
	1. Was a consec- utive or random	<b>Yes:</b> if a consecutive or random sample of pa- tients was enrolled	fpa-					
	sample of patients enrolled?	<b>No:</b> if non-consecutive patients or a non-random sample was enrolled						
		<b>Unclear:</b> if there is insufficient information on enrolled patients						
	2. Did the study avoid inappropri-	This needs to be addressed on a case-to-case ba- sis.						
	ate exclusions?	<b>Yes:</b> if all children who presented with blunt trau- ma were included or if there were appropriate reasons provided for all excluded participants						
		<b>No:</b> if eligible patients were excluded without pro- viding a reason or if exclusions might affect test accuracy (e.g. excluding on the basis of certain clinical features or comorbidities)						
		<b>Unclear:</b> if there is insufficient information on exclusions						
	3. Was the study of	Yes: if study was of prospective study design						
	prospective study design?	<b>No:</b> if the study was of retrospective or cross-sec- tional study design						
		<b>Unclear:</b> if there is insufficient information on study design						

Domain 2 Index test	Could the interpret (high/low/unclear)	ation of the index test have introduced bias?	test, its conduct, tation differ from	Are there concerns that the index test, its conduct, or the interpre- tation differ from the review ques- tion? (high/low/unclear)				
	1. Were the index test results inter-	<b>Yes:</b> if the index test results were always interpreted without knowledge of the reference standard						
	preted without knowledge of the results of the ref- erence standard?	<b>No:</b> if the index test results were interpreted with knowledge of the results of the reference standard						
		<b>Unclear:</b> if there is insufficient information pro- vided on whether the results of the index test were interpreted without knowledge of the refer- ence standard						
	2. Was the thresh-	Yes: if the threshold was prespecified						
	old used prespeci- fied?	No: if the threshold was not prespecified						
		<b>Unclear:</b> if it is unclear whether the threshold was prespecified						
Domain 3	-	ation of the reference standard have introduced	Are there concern					
Reference stan- dard	bias? (high/low/un	clear)	condition as defined by the refer- ence standard does not match the review question? (high/low/un- clear)					
	1. Is the reference standard likely to correctly classify the target condi-	<b>Yes:</b> if the target condition was defined and all pa- tients received either imaging (CT, MRI or X-ray) or clinical evaluation to clear the cervical spine and an additional follow-up after discharge	1. Did the study provide a clear definition of what was con-	<b>Yes:</b> if the target condition was clearly defined				
	tion?	<b>No:</b> if any participant did not receive either imag- ing or clinical evaluation to clear the cervical spine	sidered to be a "positive" result for the reference standard?	<b>No:</b> if the target condition was not defined				
		<b>Unclear:</b> if the target condition definition was unclear or if information on the interpretation or execution of the reference standard was unclear		<b>Unclear:</b> if the definition of the target condition was not clearly reported				
	2. Were the refer- ence standard re- sults interpreted without knowl- edge of the results	<b>Yes:</b> if the interpreter of the reference standard was clearly not aware of the results of the index text <b>No:</b> if the interpreter of the reference standard						
	of the index test?	was aware of the results of the index text <b>Unclear:</b> if insufficient information was provided on independent or blind assessment of the refer- ence test						
Domain 4 Flow & timing	Could the patient f (low/high/unclear)	low have introduced bias?						
Flow & timing	1. Is the time pe- riod between pre- sentation to ED	<b>Yes:</b> if the delay between presentation to ED with blunt trauma, execution of the index test(s) and reference standard was acceptable in most partic-						

Table 1. Assessm	ent of methodolog with blunt trau- ma and execu- tion of the refer- ence standard and index test short enough to be rea- sonably sure that the target con- dition did not change between the two tests?	<ul> <li>ical quality: QUADAS-2 and additional items (Continued)</li> <li>ipants. If imaging was conducted in the ED it was acceptable</li> <li>No: if the delay between presentation to ED with blunt trauma, execution of the index test(s) and reference standard was unacceptable in most participants</li> <li>Unclear: if the time between presentation to ED with blunt trauma, execution of the index test(s) and reference standard was unclear</li> </ul>
	2. Did all patients receive the same reference stan- dard?	<ul> <li>Yes: all patients underwent the same type of imaging or all patients underwent follow-up after discharge</li> <li>No: patients received different reference standards without follow-up after discharge</li> <li>Unclear: if information provided was unclear</li> </ul>
	3. Were all pa- tients included in the analysis?	<ul> <li>Yes: if all participants were included in the analysis or if not all participants were included in the analysis but:</li> <li>the withdrawals did not meet inclusion criteria prior to execution of index test</li> <li>the withdrawals were explained and were appropriate</li> <li>No: if any participant was excluded from the analysis for inappropriate reasons or exclusions were not explained</li> <li>Unclear: if information provided was unclear</li> </ul>

ED: emergency department.

Table 2.	Number of citations by search engine
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Search engine	Number of citations
CENTRAL	154
MEDLINE	3109
Embase	7001
Proquest Dissertation and Theses	35
PubMed	2810
OpenGrey	6
ClinicalTrials.gov	57
WHO ICTRP	3



# Table 2. Number of citations by search engine (Continued)

Web of Science	1260
Subtotal	14,435
Minus duplicates	4415
TOTAL	10,020

# Table 3. Index tests for CSI identification

Test	Clinical predictors	Classification of result
NEXUS	<ul> <li>Focal neurological deficit present</li> <li>Midline spinal tenderness present</li> <li>Altered level of consciousness present</li> <li>Intoxication present</li> <li>Distracting injury present</li> </ul>	A negative test occurs when all predic- tors are absent and suggests imaging can be avoided.
Canadian C-Spine Rule	<ul> <li>High-risk predictors</li> <li>Age ≥ 65 years</li> <li>Dangerous mechanism (fall from ≥ 0.9 m (3 feet), axial load to the head, high-speed motor vehicle collision (e.g. &gt; 100 km/ hour, rollover, ejection), motorised recreational vehicles, bicycle collision)</li> <li>Paraesthesias in extremities</li> <li>Low-risk predictors</li> <li>Simple rear-end motor vehicle collision (exclude hit by bus/large truck, rollover, hit by high-speed vehicle, pushed into traffic)</li> <li>Sitting position in emergency department</li> <li>Ambulatory at any time since the injury</li> <li>Delayed onset of neck pain</li> <li>Absence of midline cervical spine tenderness</li> <li>If patient has any low-risk predictor then a physical examination is needed to ascertain if the patient can rotate their neck 45° left and right.</li> </ul>	A positive test occurs if any high-risk predictor is present or if low-risk predic- tors are absent and suggests imaging is warranted. A positive test also occurs if any low-risk predictor is present but a patient is unable to actively rotate their neck 45° left and right.
NICE clinical guideline 56	<ul> <li>Severe head injury (GCS ≤ 8)</li> <li>Strong clinical suspicion despite normal plain films</li> <li>Plain films are technically difficult or inadequate</li> </ul>	A positive index test occurs if any pre- dictors are present and are considered appropriate indications for CT imaging.
NICE clinical guideline 176	<ul> <li>GCS &lt; 13 on initial assessment</li> <li>Strong clinical suspicion despite normal plain films</li> <li>Plain films are technically difficult or inadequate</li> <li>Patient is intubated</li> <li>Focal peripheral neurological signs</li> <li>Paraesthesia in upper and lower limbs</li> <li>A definitive diagnosis is required (such as before surgery)</li> <li>The patient is having other areas scanned for multiregion trauma</li> </ul>	A positive index test occurs if any pre- dictors are present and are considered appropriate indications for CT imaging.



Table 3. Index tests fo	<ul> <li><b>or CSI identification</b> (Continued)</li> <li>Plain films demonstrate a significant bony injury</li> </ul>	
Pierretti de novo (PEDSPINE)	<ul> <li>GCS ≥ 14</li> <li>Motor vehicle crash</li> <li>GCS<sub>EYE</sub> = 1</li> <li>Age &gt; 2 years (24–36 months)</li> </ul>	Each of the predictors were assigned points as part of an overall weighted score (3 points to GCS < 14, 2 points to GCS <sub>EYE</sub> = 1 and motor vehicle crash and 1 point to age > 2 years). A negative in- dex test occurs if the weighted score is 0 or 1 and suggests imaging can be avoid- ed.
PECARN retrospective criteria	<ul> <li>High-risk motor vehicle crash</li> <li>Diving</li> <li>Conditions predisposing to CSI</li> <li>Substantial torso injury</li> <li>Torticollis (decreased neck mobility by report or examination)</li> <li>Neck pain (child complaint if &gt; 2 years)</li> <li>Focal neurological findings</li> <li>Altered mental status</li> </ul>	Identified an 8 variable model of predic- tive factors for CSI after blunt trauma. These factors are considered in the de- velopment of the Leonard de novo mod- el. A negative test occurs when all pre- dictors are absent and suggests imaging can be avoided.
Leonard de novo	<ul> <li>Altered mental status</li> <li>Focal neurological findings</li> <li>Substantial torso injury</li> <li>Neck pain</li> <li>Torticollis</li> <li>Conditions predisposing to cervical injury</li> <li>Diving</li> <li>High-risk motor vehicle crash</li> </ul>	A negative test occurs when all predic- tors are absent and suggests imaging can be avoided.

CSI: cervical spine injury; CT: computed tomography; GCS: Glasgow Coma Scale; NEXUS: National Emergency X-Radiography Utilization Study; NICE: National Institute for Health and Care Excellence; PECARN: Pediatric Emergency Care Applied Research Network.

## APPENDICES

## Appendix 1. Search strategies - update

## MEDLINE

Search 13 December 2022

- 1. (NEXUS or CCR).mp.
- 2. National-Emergency-X-Radiography.mp.
- 3. (Canadian-C-Spine or Canadian-cervical-spine).mp.
- 4. ((clinical or critical or treatment) adj3 (pathway\* or protocol\*)).mp.
- 5. (algorithm\* or guideline\*).mp.
- 6. (decision adj3 (tree\* or rule\* or tool\*)).mp.
- 7. (triage or protocol\*).mp.
- 8. exp guideline/
- 9. Guideline Adherence/



- 10. exp Guidelines as Topic/
- 11. exp algorithms/
- 12. exp Clinical Protocols/
- 13. Decision Trees/
- 14. exp decision support techniques/
- 15. Critical Pathways/
- 16. Triage/
- 17. ((neurolog\* or physical\* or clinical\*) adj3 (exam\* or assess\* or sign\*)).mp.
- 18. (MRI\* or CT\* or Computed-tomograph\* or CAT-scan\*).mp.
- 19. (x-ray\* or xray\* or radiograph\* or roentgenogra\* or imaging).mp.
- 20. exp Physical Examination/
- 21. exp trauma severity indices/
- 22. "Severity of Illness Index"/
- 23. X-Rays/
- 24. tomography/ or exp tomography, emission-computed/ or exp tomography, x-ray/
- 25. exp Magnetic Resonance Imaging/
- 26. Radiography/
- 27. or/1-26
- 28. ((cervical-spine or c-spine) adj5 clear\*).mp.
- 29. (cervical adj5 (trauma\* or injur\* or fracture\* or sublux\* or dislocat\* or avuls\* or instab\*)).mp.
- 30. (spinal-cord-injury-without-radiographic-abnormalit\* or SCIWORA).mp.
- 31. exp Cervical Vertebrae/
- 32. exp Neck Injuries/
- 33. exp Spinal Injuries/
- 34. exp Spinal Cord Injuries/
- 35. or/28-34

36. (perinatal or perinatology or newborn\* or new-born\* or baby or babies or neonat\* or neo-nat\* or infan\* or toddler\* or pre-schooler\* or preschooler\* or kinder or kinders or kindergarten\* or kinder-aged or boy or boys or girl or girls or child or children or childhood or pediatric\* or paediatric\* or adolescen\* or youth or youths or teen or teens or teenage\* or school-age\* or schoolage\* or school-child\* or schoolchild\* or school-boy\* or schoolboy\* or juvenile\* or preteen\* or pre-teen\*).af.

37. 27 and 35 and 36

38. limit 37 to yr="2015 -Current"

39. limit 38 to english language

### Embase

Search 13 December 2022

1. (NEXUS or CCR).mp.



- 2. National-Emergency-X-Radiography.mp.
- 3. (Canadian-C-Spine or Canadian-cervical-spine).mp.
- 4. ((clinical or critical or treatment) adj3 (pathway\* or protocol\*)).mp.
- 5. (algorithm\* or guideline\*).mp.
- 6. (decision adj3 (tree\* or rule\* or tool\*)).mp.
- 7. (triage or protocol\*).mp.
- 8. practice guideline/ or clinical pathway/ or exp clinical protocol/
- 9. exp algorithm/
- 10. "decision tree"/
- 11. exp decision support system/
- 12. emergency health service/
- 13. ((neurolog\* or physical\* or clinical\*) adj3 (exam\* or assess\* or sign\*)).mp.
- 14. (MRI\* or CT\* or Computed-tomograph\* or CAT-scan\*).mp.
- 15. (x-ray\* or xray\* or radiograph\* or roentgenogra\* or imaging).mp.
- 16. exp physical examination/
- 17. exp neurologic examination/
- 18. exp injury scale/
- 19. X ray/
- 20. exp tomography/
- 21. exp radiography/
- 22. or/1-21
- 23. ((cervical-spine or c-spine) adj5 clear\*).mp.
- 24. (cervical adj5 (trauma\* or injur\* or fracture\* or sublux\* or dislocat\* or avuls\* or instab\*)).mp.
- 25. (spinal-cord-injury-without-radiographic-abnormalit\* or SCIWORA).mp.
- 26. exp cervical spine/
- 27. exp neck injury/
- 28. exp spine injury/
- 29. exp spinal cord injury/
- 30. or/23-29

31. (perinatal or perinatology or newborn\* or new-born\* or baby or babies or neonat\* or neo-nat\* or infan\* or toddler\* or pre-schooler\* or preschooler\* or kinder or kinders or kindergarten\* or kinder-aged or boy or boys or girl or girls or child or children or childhood or pediatric\* or paediatric\* or adolescen\* or youth or youths or teen or teens or teenage\* or school-age\* or schoolage\* or school-child\* or schoolchild\* or schoolchild\* or schoolboy\* or juvenile\* or preteen\* or pre-teen\*).af.

32. 22 and 30 and 31

33. limit 32 to (english language and yr="2015 -Current")

### **CENTRAL (the Cochrane Library)**

Search 13 December 2022

- 1. (NEXUS or CCR) (Word variations have been searched)
- 2. (National-Emergency-X-Radiography) (Word variations have been searched)
- 3. (Canadian-C-Spine or Canadian-cervical-spine) (Word variations have been searched)
- 4. ((clinical or critical or treatment) NEAR/3 (pathway\* or protocol\*)) (Word variations have been searched)
- 5. ((clinical or critical or treatment) NEAR/3 (pathway\* or protocol\*)) (Word variations have been searched)
- 6. (decision NEAR/3 (tree\* or rule\* or tool\*)) (Word variations have been searched)
- 7. (triage or protocol\*) (Word variations have been searched)
- 8. MeSH descriptor: [Guideline] explode all trees
- 9. MeSH descriptor: [Guideline Adherence] this term only
- 10. MeSH descriptor: [Guidelines as Topic] explode all trees
- 11. MeSH descriptor: [Algorithms] explode all trees
- 12. MeSH descriptor: [Clinical Protocols] explode all trees
- 13. MeSH descriptor: [Decision Trees] this term only
- 14. MeSH descriptor: [Decision Support Techniques] explode all trees
- 15. MeSH descriptor: [Critical Pathways] this term only
- 16. MeSH descriptor: [Triage] this term only
- 17. ((neurolog\* or physical\* or clinical\*) NEAR/3 (exam\* or assess\* or sign\*)) (Word variations have been searched)
- 18. (MRI\* or CT\* or Computed-tomograph\* or CAT-scan\*) (Word variations have been searched)
- 19. (x-ray\* or xray\* or radiograph\* or roentgenogra\* or imaging) (Word variations have been searched)
- 20. MeSH descriptor: [Physical Examination] explode all trees
- 21. MeSH descriptor: [Trauma Severity Indices] explode all trees
- 22. MeSH descriptor: [Severity of Illness Index] this term only
- 23. MeSH descriptor: [X-Rays] this term only
- 24. MeSH descriptor: [Tomography] this term only
- 25. MeSH descriptor: [Tomography, Emission-Computed] explode all trees
- 26. MeSH descriptor: [Tomography, X-Ray] explode all trees
- 27. MeSH descriptor: [Magnetic Resonance Imaging] explode all trees
- 28. MeSH descriptor: [Radiography] this term only

29. #1 or #2 or #3 or #4 or #5 or #6 or #7 or #8 or #9 or #10 or #11 or #12 or #13 or #14 or #15 or #16 or #17 or #18 or #19 or #20 or #21 or #22 or #23 or #24 or #25 or #26 or #27 or #28

- 30. ((cervical-spine or c-spine) NEAR/5 clear\*) (Word variations have been searched)
- 31. (cervical NEAR/5 (trauma\* or injur\* or fracture\* or sublux\* or dislocat\* or avuls\* or instab\*)) (Word variations have been searched)
- 32. (spinal-cord-injury-without-radiographic-abnormalit\* or SCIWORA)

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- 33. MeSH descriptor: [Cervical Vertebrae] explode all trees
- 34. MeSH descriptor: [Neck Injuries] explode all trees
- 35. MeSH descriptor: [Spinal Injuries] explode all trees
- 36. MeSH descriptor: [Spinal Cord Injuries] explode all trees
- 37. #30 or #31 or #32 or #33 or #34 or #35 or #36

38. (perinatal or perinatology or newborn\* or new-born\* or baby or babies or neonat\* or neo-nat\* or infan\* or toddler\* or pre-schooler\* or preschooler\* or kinder or kinders or kindergarten\* or kinder-aged or boy or boys or girl or girls or child or children or childhood or pediatric\* or paediatric\* or adolescen\* or youth or youths or teen or teens or teenage\* or school-age\* or schoolage\* or school-child\* or schoolchild\* or school-boy\* or schoolboy\* or juvenile\* or preteen\* or pre-teen\*)

39. #29 and #37 and #38

#### **Proquest Dissertations & Theses database**

Search 13 December 2022

#### 1. Anywhere except full text

"NEXUS" OR "CCR" OR "National Emergency X Radiography" OR "Canadian c spine" OR "Canadian cervical spine" OR "Clinical pathway\*" OR "critical pathway\*" OR "treatment pathway\*" OR "clinical protocol\*" OR "critical protocol\*" OR "treatment protocol\*" OR "algorithm\*" OR "guideline\*" OR "decision tree\*" OR "decision tool\*" OR "decision rule\*" OR "decision support" OR "triage" OR "protocol\*" OR (("neurolog\*" OR "physical\*" OR "clinical\*") AND ("exam\*" OR "assess\*" OR "sign" OR "signs")) OR "MRI" OR "MRIS" OR "CT" OR "CTS" OR "Tomograph\*" OR "CAT scan\*" OR "x-ray\*" OR "xray\*" OR "radiogra\*" OR "injury scale"

AND

#### 2. Anywhere except full text

(("cervical-spine" OR "c-spine") AND "clear") OR "cervical trauma\*" OR "cervical injur\*" OR "cervical fracture\*" OR "cervical sublux\*" OR "cervical disloc\*" OR "cervical avuls\*" OR "cervical instab\*" OR "Spinal cord injury without radiographic abnormalit\*" OR "SCIWORA" OR "cervical-vertebra\*" OR "neck-injur\*" OR "spinal-injur\*" OR "spinal-trauma" OR "spine-injur\*" OR "spine-trauma" OR "spinal-cord-injur\*" OR "spinal-cord-injur\*" OR "spinal-cord-trauma" OR "spinal-trauma" OR "spinal-trauma"

AND

### 3. Anywhere except full text

"perinatal" OR "perinatology" OR "newborn\*" OR "new-born\*" OR "baby" OR "babies" OR "neonat\*" OR "neo-nat\*" OR "infan\*" OR "toddler\*" OR "pre-schooler\*" OR "preschooler\*" OR "kinder" OR "kinders" OR "kinders" OR "kinder-aged" OR "boys" OR "girls" OR "girls" OR "children" OR "childhood" OR "pediatric\*" OR "paediatric\*" OR "adolescen\*" OR "youth" OR "youths" OR "teens" OR "teens" OR "teenage\*" OR "school-age\*" OR "schoolage\*" OR "school-child\*" OR "school-child\*" OR "school-boys" OR "school-boys" OR "juvenile\*" OR "pre-teen\*"

Limit from 2015

#### PubMed

#### Search 13 December 2022

("NEXUS" OR "CCR" OR "National Emergency X Radiography" OR "Canadian c spine" OR "Canadian cervical spine" OR "Clinical pathway\*" OR "treatment pathway\*" OR "clinical protocol\*" OR "critical protocol\*" OR "treatment protocol\*" OR "algorithm\*" OR "guideline\*" OR "decision tree\*" OR "decision tool\*" OR "decision rule\*" OR "decision support" OR "triage" OR "protocol\*" OR (("neurolog\*" OR "physical\*" OR "clinical\*") AND ("exam\*" OR "assess\*" OR "sign" OR "signs")) OR "MRI" OR "MRIS" OR "CT" OR "CT" OR "CT" OR "CT" OR "critical protocol\*" OR "injury scale")) OR "MRI" OR "MRIS" OR "CT" OR "CT" OR "CT" OR "severity of illness" OR "Glasgow-coma-scale" OR "injury-severity-score\*" OR "injury scale") AND((("cervical-spine" OR "cervical avuls\*" OR "cervical instab\*" OR "cervical injury without radiographic abnormalit\*" OR "SCIWORA" OR "cervical-vertebra\*" OR "spinal-trauma" OR "spinal-cord-trauma" OR "spinal-cord-trauma" OR "spinal-cord-trauma" OR "spinal-cord-trauma" OR "spinal-cord-trauma" OR "spinal-fracture\*" OR "whiplash-injur\*" OR "whiplash-injur\*"



"perinatology" OR "newborn\*" OR "new-born\*" OR "baby" OR "babies" OR "neonat\*" OR "neo-nat\*" OR "infan\*" OR "toddler\*" OR "pre-schooler\*" OR "preschooler\*" OR "kinder" OR "kinders" OR "kindergarten\*" OR "kinder-aged" OR "boy" OR "boys" OR "girl" OR "girls" OR "child" OR "children" OR "childhood" OR "pediatric\*" OR "paediatric\*" OR "adolescen\*" OR "youth" OR "youths" OR "teen" OR "teens" OR "teenage\*" OR "school-age\*" OR "schoolage\*" OR "school-child\*" OR "schoolchild\*" OR "school-girl\*" OR "school-girl\*" OR "preteen\*" OR "preteen\*") AND (NOTNLM OR publisher[sb] OR inprocess[sb] OR pubmednotmedline[sb] OR indatareview[sb] OR pubstatusaheadofprint)

Limited to English; 2015-

#### OpenGrey

Search 15 December 2022

(NEXUS OR CCR OR "National Emergency X Radiography" OR "Canadian c spine" OR "Canadian cervical spine" OR Clinical-pathway\* OR critical-protocol\* OR critical-protocol\* OR treatment-protocol\* OR algorithm\* OR guideline\* OR decision-tree\* OR decision-tool\* OR decision-rule\* OR decision-support OR triage OR protocol\* OR ((neurolog\* OR physical\* OR clinical\*) AND (exam\* OR assess\* OR sign OR signs)) OR MRI OR MRIS OR CT OR CTS OR Tomograph\* OR CAT-scan\* OR x-ray\* OR xray\* OR radiogra\* OR roentgenogra\* OR imaging OR trauma-severity OR severity-of-illness OR Glasgow-coma-scale OR injury-severity-score\* OR injury-scale) AND(((cervical-spine OR c-spine) AND clear) OR cervical-trauma\* OR cervical-injur\* OR cervical-fracture\* OR cervical-sublux\* OR cervical-disloc\* OR cervical-avuls\* OR cervical-instab\* OR Spinal-cord-injury-without-radiographic-abnormalit\* OR SCIWORA OR cervical-vertebra\* OR neck-injur\* OR spinal-injur\* OR spinal-trauma OR spinal-cord-injur\* OR s

Limit to English

Limit from 2015

#### ClinicalTrials.gov

Search 14 December 2022

#### **Condition or disease**

SCIWORA OR ((Cervical OR c-spine OR spinal OR neck OR spine OR whiplash OR whip-lash) AND (trauma OR sublux OR dislocat\* OR avuls\* OR instab\* OR injur\* OR abnormalit\* OR compression OR fracture\*))

#### AND

#### **Other terms**

NEXUS OR CCR OR "National Emergency X Radiography" OR "Canadian c spine" OR "Canadian cervical spine" OR pathway\* OR protocol\* OR algorithm\* OR guideline\* OR decision OR triage

### Science Citation Index (Web of Science, Core Collection)

Search 15 December 2022

#1 TS=(NEXUS OR CCR OR National-Emergency-X-Radiography OR Canadian-c-spine OR Canadian-cervical-spine OR Clinical-pathway\* OR critical-pathway\* OR treatment-pathway\* OR clinical-protocol\* OR critical-protocol\* OR treatment-protocol\* OR algorithm\* OR guideline\* OR decision-tree\* OR decision-tool\* OR decision-rule\* OR decision-support OR triage OR protocol\* OR MRI OR MRIS OR CT OR CTS OR Tomograph\* OR CAT-scan\* OR x-ray\* OR xray\* OR radiogra\* OR roentgenogra\* OR imaging OR trauma-severity OR severity-of-illness OR Glasgow-coma-scale OR injury-severity-score\* OR injury-scale)

#2 TS=((neurolog\* OR physical\* OR clinical\*) AND (exam\* OR assess\* OR sign OR signs))

#3 #1 OR #2

#4 TS=((cervical-spine OR c-spine) AND clear\*)

#5 TS=(cervical-trauma\* OR cervical-injur\* OR cervical-fracture\* OR cervical-sublux\* OR cervical-disloc\* OR cervical-avuls\* OR cervicalinstab\* OR Spinal-cord-injury-without-radiographic-abnormalit\* OR SCIWORA OR cervical-vertebra\* OR neck-injur\* OR spinal-injur\* OR spinal-trauma OR spine-injur\* OR spine-trauma OR spinal-cord-injur\* OR spinal-cord-trauma OR spinal-cord-compression OR spinalfracture\* OR spine-fracture\* OR whiplash-injur\* OR whip-lash-injur\*)

#6 #4 OR #5

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#7 TS=(perinatal OR perinatology OR newborn\* OR new-born\* OR baby OR babies OR neonat\* OR neo-nat\* OR infan\* OR toddler\* OR preschooler\* OR preschooler\* OR kinder OR kinders OR kindergarten\* OR kinder-aged OR boy OR boys OR girl OR girls OR child OR children OR childhood OR pediatric\* OR paediatric\* OR adolescen\* OR youth OR youths OR teen OR teenage\* OR school-age\* OR schoolage\* OR school-child\* OR schoolchild\* OR school-girl\* OR schoolgirl\* OR school-boy\* OR schoolboy\* OR juvenile\* OR preteen\* OR pre-teen\*)

#8 #3 AND #6 AND #7

Limited to English; 2015-

#### WHO International Clinical Trials Registry Platform (ICTRP)

Search 15 December 2022

(NEXUS OR National Emergency X-Radiography OR Canadian c-spine OR Canadian Cervical Spine) AND (cervical fracture OR cervical injury OR cervical trauma ORR cervical dislocation OR cervical instability OR cervical avulsion)

#### Appendix 2. Search strategies – original review

MEDLINE

Search 24 February 2015

1 (NEXUS or CCR).mp. (5678)

- 2 National Emergency X-Radiography.mp. (48)
- 3 (Canadian c-spine or Canadian cervical spine).mp. (40)

4 ((Clinical or critical or treatment) adj3 (pathway\* or protocol\*)).mp. (52575)

5 (algorithm\* or guideline\*).mp. (488839)

6 (decision adj3 (tree\* or rule\* or tool\*)).mp. (15657)

7 (triage or protocol\*).mp. (366206)

8 or/1-7 [Triage tool keywords] (856300)

9 exp Guideline/ (25808)

10 Guideline Adherence/ (21997)

11 exp guidelines as topic/ (117788)

12 exp algorithms/ (181301)

13 exp Clinical Protocols/ [includes antineoplastic protocols] (127941)

14 Decision Trees/ (8964)

15 exp decision support techniques/[includes data interpretation, statistical] (61776)

16 Critical Pathways/ (4775)

17 triage/ (8242)

18 or/9-17 [Triage tool MeSH terms] (522506)

198 or 18 [Triage tools] (907178)

20 ((neurolog\* or physical\* or clinical\*) adj3 (exam\* or assess\* or sign\*)).mp. (384485)

21 MRI\*.mp. (359278)

22 (CT\* or Computed Tomography or CAT scan\*).mp. (384167)

23 (X ray\* or x-ray\* or xray\* or radiogra\* or roentgenogra\*).mp. (734670)

24 Imaging.mp. (660363)



25 or/20-24 [Reference standard keywords] (1748610)

- 26 exp physical examination/ or exp neurologic examination/ (1075565)
- 27 exp trauma severity indices/ [includes Glasgow Coma Scale, Injury Severity Score, others] (24531)
- 28 "Severity of Illness Index"/ [not exploded leave out Karnofsky Performance Status cancer ADL measure] (173516)
- 29 X-Rays/ (16413)
- 30 Tomography/ or exp Tomography, Emission-Computed/ or exp Tomography, X-Ray/ [includes tomography, x-ray computed] (386295)
- 31 exp Magnetic Resonance Imaging/ (317050)
- 32 Radiography/ (24883)
- 33 or/26-32 [Reference standard MeSH terms] (1874249)
- 34 25 or 33 [Reference standard] (2858529)
- 35 19 or 34 [Triage tools or reference standard] (3572718)
- 36 ((Cervical spine or c-spine) adj5 clear\*).mp. (241)

37 (cervical adj5 (trauma\* or injur\* or fracture\* or sublux\* or dislocat\* or avuls\* or instab\*)).mp. [cervical spine injury, cervical spine trauma] (10352)

- 38 (Spinal cord injury without radiographic abnormality or SCIWORA).mp. (113)
- 39 or/36-38 [cervical trauma keywords] (10494)
- 40 exp Cervical Vertebrae/ [includes axis and atlas] (30752)
- 41 exp Neck Injuries/ [includes whiplash injuries] (6628)
- 42 exp Spinal Injuries/ [includes spinal fractures] (17863)
- 43 exp Spinal Cord Injuries/ [includes spinal cord compression, others] (38273)
- 44 spinal fractures/ (10537)
- 45 or/40-44 [cervical trauma MeSH terms] (81875)
- 46 39 or 45 [cervical trauma terms] (84419)
- 47 (Pediatric\* or paediatric\* or peadiatric\*).mp. (239652)
- 48 (Child\*).mp. (1864223)
- 49 (neonate\* or newborn\* or new-born\*).mp. (639758)
- 50 (infant\* or baby or babies or toddler\*).mp. (1042256)
- 51 (adolescen\* or juvenile\* or youth\* or teen\* or preteen\*).mp. (1709977)
- 52 or/47-51 [pediatric keywords] (3315420)
- 53 exp Pediatrics/ [includes perinataology, neonatology] (44737)
- 54 exp Child/ [includes child, preschool] (1562070)
- 55 exp Infant/ [includes infant, newborn] (946461)
- 56 Adolescent/ (1632349)
- 57 or/53-56 [pediatric MeSH terms] (2910260)
- 58 52 or 57 [Pediatric terms] (3316163)

### 59 35 and 46 and 58 (7985)

\*.mp. [mp=title, abstract, original title, name of substance word, subject heading word, keyword heading word, protocol supplementary concept word, rare disease supplementary concept word, unique identifier]

### **MEDLINE In-Process & Other Non-Indexed Citations**

Search 24 February 2015

- 1 (NEXUS or CCR).mp. (492)
- 2 National Emergency X-Radiography.mp. (3)
- 3 (Canadian c-spine or Canadian cervical spine).mp. (13)
- 4 ((Clinical or critical or treatment) adj3 (pathway\* or protocol\*)).mp. (3193)
- 5 (algorithm\* or guideline\*).mp. (49484)
- 6 (decision adj3 (tree\* or rule\* or tool\*)).mp. (1156)
- 7 (triage or protocol\*).mp. (29444)
- 8 or/1-7 [Triage tool keywords] (79262)
- 9 exp Guideline/ or / (62)
- 10 Guideline Adherence/ (0)
- 11 exp guidelines as topic//(0)
- 12 exp algorithms/ (0)
- 13 exp Clinical Protocols/ [includes antineoplastic protocols] (0)
- 14 Decision Trees/ (0)
- 15 exp decision support techniques/ [includes data interpretation, statistical] (0)
- 16 Critical Pathways/ (0)
- 17 triage/ (0)
- 18 or/9-17 [Triage tool MeSH terms] (62)
- 198 or 18 [Triage tools] (79262)
- 20 ((neurolog\* or physical\* or clinical\*) adj3 (exam\* or assess\* or sign\*)).mp. (30580)
- 21 MRI\*.mp. (24133)
- 22 (CT\* or Computed Tomography or CAT scan\*).mp. (39230)
- 23 (X ray\* or x-ray\* or xray\* or radiogra\* or roentgenogra\*).mp. (62975)
- 24 Imaging.mp. (61152)
- 25 or/20-24 [Reference standard keywords] (175595)
- 26 exp physical examination/ or exp neurologic examination/ (0)
- 27 exp trauma severity indices/ [includes Glasgow Coma Scale, Injury Severity Score, others] (0)
- 28 "Severity of Illness Index"/ [not exploded leave out Karnofsky Performance Status cancer ADL measure] (0)
- 29 X-Rays/ (0)

30 Tomography/ or exp Tomography, Emission-Computed/ or exp Tomography, X-Ray/ [includes tomography, x-ray computed] (0)

**Triage tools for detecting cervical spine injury in paediatric trauma patients (Review)** Copyright © 2024 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.



31 exp Magnetic Resonance Imaging/(0)

32 Radiography/(0) 33 or/26-32 [Reference standard MeSH terms] (0) 34 25 or 33 [Reference standard] (175595) 35 19 or 34 [Triage tools or reference standard] (244188) 36 ((Cervical spine or c-spine) adj5 clear\*).mp. (20) 37 (cervical adj5 (trauma\* or injur\* or fracture\* or sublux\* or dislocat\* or avuls\* or instab\*)).mp. [cervical spine injury, cervical spine trauma] (833) 38 (Spinal cord injury without radiographic abnormality or SCIWORA).mp. (10) 39 or/36-38 [cervical trauma keywords] (848) 40 exp Cervical Vertebrae/ [includes axis and atlas] (0) 41 exp Neck Injuries/ [includes whiplash injuries] (0) 42 exp Spinal Injuries/ [includes spinal fractures] (0) 43 exp Spinal Cord Injuries/ [includes spinal cord compression, others] (0) 44 spinal fractures/(0) 45 or/40-44 [cervial trauma MeSH terms] (0) 46 39 or 45 [cervical trauma terms] (848) 47 (Pediatric\* or paediatric\* or peadiatric\*).mp. (20264) 48 (Child\* ).mp. (67273) 49 (neonate\* or newborn\* or new-born\*).mp. (9578) 50 (infant\* or baby or babies or toddler\*).mp. (19120) 51 (adolescen\* or juvenile\* or youth\* or teen\* or preteen\*).mp. (24018) 52 or/47-51 [pediatric keywords] (106495) 53 exp Pediatrics/ [includes perinataology, neonatology] (0) 54 exp Child/ [includes child, preschool] (0) 55 exp Infant/ [includes infant, newborn] (0) 56 Adolescent/ (0) 57 or/53-56 [pediatric MeSH terms] (0) 58 52 or 57 [Pediatric terms] (106495) 59 35 and 46 and 58 (39) Embase Search 24 February 2015 1 (NEXUS or CCR).mp. (7884) 2 National Emergency X-Radiography.mp. (59) 3 (Canadian c-spine or Canadian cervical spine).mp. (74)



4 ((Clinical or critical or treatment) adj3 (pathway* or protocol*)).mp. (116085)
5 (algorithm* or guideline*).mp. (665390)
6 (decision adj3 (tree* or rule* or tool*)).mp. (16747)
7 (triage or protocol*).mp. (426356)
8 or/1-7 (1087993)
9 practice guideline/ or clinical pathway/ or clinical protocol/ (315692)
10 exp algorithm/ (188334)
11 "decision tree"/ (6358)
12 exp decision support system/ (14055)
13 emergency health service/ [used for triage] (69092)
14 or/9-13 (575274)
15 8 or 14 (1149485)
16 ((neurolog* or physical* or clinical*) adj3 (exam* or assess* or sign*)).mp. (779457)
17 MRI*.mp. (587063)
18 (CT* or Computed Tomography or CAT scan*).mp. (600538)
19 (X ray* or x-ray* or xray* or radiogra* or roentgenogra*).mp. (876246)
20 Imaging.mp. (1064732)
21 or/16-20 (2762608)
22 exp physical examination/ (160790)
23 exp neurologic examination/ (350398)
24 exp injury scale/ [used for trauma severity indices] (29378)
25 X ray/ (41737)
26 exp tomography/ (728786)
27 exp computer assisted tomography/ (636166)
28 exp nuclear magnetic resonance imaging/ (576997)
29 exp radiography/ (897201)
30 or/22-29 (2073134)
31 21 or 30 (3328617)
32 15 or 31 (4251928)
33 ((Cervical spine or c-spine) adj5 clear*).mp. (344)
34 (cervical adj5 (trauma* or injur* or fracture* or sublux* or dislocat* or avuls* or instab*)).mp. (17235)
35 (Spinal cord injury without radiographic abnormality or SCIWORA).mp. (181)
36 or/33-35 (17402)
37 exp cervical spine/ (27701)
38 exp neck injury/ (10805)



39 exp spine injury/ (30879) 40 exp spinal cord injury/ (54289) 41 exp spine fracture/ (15827) 42 or/37-41 (112517) 43 36 or 42 (115463) 44 (Pediatric\* or paediatric\* or peadiatric\*).mp. (389216) 45 (Child or children or childhood).mp. (2006835) 46 (neonate\* or newborn\* or new-born\*).mp. (567587) 47 (infant\* or baby or babies or toddler\*).mp. (778286) 48 (adolescen\* or juvenile\* or youth\* or teen\* or preteen\*).mp. (1399956) 49 or/44-48 (3309713) 50 exp pediatrics/ (77383) 51 exp child/ (2059816) 52 exp infant/ (857030) 53 exp adolescent/ (1253833) 54 exp juvenile/ (2715056) 55 exp adolescence/ (66747) 56 exp childhood/ (50991) 57 exp childhood injury/ (7203) 58 or/50-57 (2785218) 59 49 or 58 (3349155) 60 32 and 43 and 59 (9187) CENTRAL Search 24 February 2015 #1 NEXUS or CCR 783 #2 National Emergency X-Radiography 2 #3 Canadian c-spine or Canadian cervical spine 59 #4 ((Clinical or critical or treatment) near/3 (pathway\* or protocol\*)) 8326 #5 algorithm\* or guideline\* 24279 #6 (decision near/3 (tree\* or rule\* or tool\*)) 2370 #7 triage or protocol\* 56538 #8 #1 or #2 or #3 or #4 or #5 or #6 or #7 76450 #9 MeSH descriptor: [Guideline] explode all trees 19 #10 MeSH descriptor: [Practice Guideline] 15 #11 MeSH descriptor: [Guideline Adherence] 739

#12 MeSH descriptor: [Guidelines as Topic] explode all trees 2078 #13 MeSH descriptor: [Practice Guidelines as Topic] 1770 #14 MeSH descriptor: [Algorithms] explode all trees 3040 #15 MeSH descriptor: [Clinical Protocols] explode all trees 13095 #16 MeSH descriptor: [Decision Trees] 895 #17 MeSH descriptor: [Decision Support Techniques] explode all trees 3202 #18 MeSH descriptor: [Critical Pathways] 262 #19 MeSH descriptor: [Triage] 258 #20 #9 or #10 or #11 or #12 or #13 or #14 or #15 or #16 or #17 or #18 or #19 22374 #21 #8 or #20 78691 #22 ((neurolog\* or physical\* or clinical\*) near/3 (exam\* or assess\* or sign\*)) 45542 #23 MRI\* 11288 #24 CT\* or Computed Tomography or CAT scan 56407 #25 X ray\* or x-ray\* or xray\* or radiogra\* or roentgenogra\* 23437 #26 Imaging 21622 #27 #22 or #23 or #24 or #25 or #26 123319 #28 MeSH descriptor: [Physical Examination] explode all trees 72073 #29 MeSH descriptor: [Neurologic Examination] explode all trees 16982 #30 MeSH descriptor: [Trauma Severity Indices] explode all trees 993 #31 MeSH descriptor: [Severity of Illness Index] this term only 14375 #32 MeSH descriptor: [X-Rays] 44 #33 MeSH descriptor: [Tomography] explode all trees 11885 #34 MeSH descriptor: [Tomography, Emission-Computed] explode all trees 2630 #35 MeSH descriptor: [Tomography, X-Ray] explode all trees 4107 #36 MeSH descriptor: [Magnetic Resonance Imaging] explode all trees 5716 #37 MeSH descriptor: [Radiography] explode all trees 13863 #38 #28 or #29 or #30 or #31 or #32 or #33 or #34 or #35 or #36 or #37 103001 #39 #27 or #38 199647 #40 #21 or #39 253140 #41 ((Cervical spine or c-spine) near/5 clear\*) 10 #42 (cervical near/5 (trauma\* or injur\* or fracture\* or sublux\* or dislocat\* or avuls\* or instab\*)) 472 #43 Spinal cord injury without radiographic abnormality or SCIWORA 2 #44 #41 or #42 or #43 474 #45 MeSH descriptor: [Cervical Vertebrae] explode all trees 776 #46 MeSH descriptor: [Neck Injuries] explode all trees 205



#47 MeSH descriptor: [Spinal Injuries] explode all trees 720
#48 MeSH descriptor: [Spinal Cord Injuries] explode all trees 906
#49 MeSH descriptor: [Spinal Fractures] explode all trees 636
#50 #45 or #46 or #47 or #48 or #49 2442
#51 #44 or #50 2730
#52 Pediatric\* or paediatric\* or peadiatric\* 41377
#53 Child or children or childhood 94858
#54 neonate\* or newborn\* or new-born\* 19365
#55 infant\* or baby or babies or toddler\* 41412
#56 adolescen\* or juvenile\* or youth\* or teen\* or preteen\* 98728
#57 #52 or #53 or #54 or #55 or #56 176411
#58 MeSH descriptor: [Pediatrics] explode all trees 546
#59 MeSH descriptor: [Child] explode all trees 135

#60 MeSH descriptor: [Infant] explode all trees 13304

#61 MeSH descriptor: [Adolescent] 76925

#62 #58 or #59 or #60 or #61 89391

#63 #57 or #62 176421

#64 40 and 51 and 63 4201

#65 #64 in Trials 651

#### **Science Citation Index**

Search 24 February 2015

# 4 1,220 #3 AND #2 AND #1

Indexes=SCI-EXPANDED Timespan=All years

#31,508,937

TOPIC: (Pediatric\* OR paediatric\* OR peadiatric\* OR Child OR children OR childhood OR neonate\* OR newborn\* OR new-born\* OR infant\* OR baby OR babies OR toddler\* OR adolescen\* OR juvenile\* OR youth\* OR teen\* OR preteen\*)

Indexes=SCI-EXPANDED Timespan=All years

#219,789

TS=(cervical spine clear\* OR c-spine clear\* OR clearing the c-spine OR clearing the cervical spine OR cervical trauma\* OR cervical injur\* OR cervical fracture\* OR cervical sublux\* OR cervical disloc\* OR cervical avuls\* OR cervical instab\* OR Spinal cord injury without radiographic abnormality OR SCIWORA)

Indexes=SCI-EXPANDED Timespan=All years

#14,736,061

TS=(NEXUS OR CCR OR National Emergency X-Radiography OR Canadian c-spine OR Canadian cervical spine OR Clinical pathway\* OR critical pathway\* OR treatment pathway\* OR clinical protocol\* OR treatment protocol\* OR algorithm\* OR guideline\* OR decision tree\* OR decision tool\* OR decision rule\* OR triage OR protocol\* OR neurolog\* assess\* OR neurolog\* exam\* OR neurolog\* sign\* OR physical\* exam\* OR physical\* assess\* OR physical\* sign\* OR clinical\* exam\* OR clinical\* assess\* OR clinical\* sign\* OR MRI\* OR CT OR Computed Tomography OR CAT scan\* OR X ray\* OR x-ray\* OR radiogra\* OR roentgenogra\* OR imaging)

Indexes=SCI-EXPANDED Timespan=All years

#### **Proquest Dissertations & Theses database**

Search 24 February 2015

#### Advanced search :

all(( (NEXUS OR CCR OR National Emergency X-Radiography OR Canadian c-spine OR Canadian cervical spine OR Clinical pathway\* OR critical pathway\* OR treatment pathway\* OR clinical protocol\* OR treatment protocol\* OR algorithm\* OR guideline\* OR decision tree\* OR decision tool\* OR decision rule\* OR triage OR protocol\* OR neurolog\* assess\* OR neurolog\* exam\* OR neurolog\* sign\* OR physical\* exam\* OR physical\* assess\* OR physical\* sign\* OR clinical\* exam\* OR clinical\* assess\* OR clinical\* sign\* OR MRI\* OR CT\* OR Computed Tomography OR CAT scan\*OR X ray\* OR x-ray\* OR xray\* OR radiogra\* OR roentgenogra\* OR imaging) AND (cervical spine clear\* OR c-spine clear\* OR clearing the c-spine OR clearing the cervical spine OR cervical trauma\* OR cervical injur\* OR cervical fracture\* OR cervical sublux\* OR cervical disloc\* OR cervical avuls\* OR cervical instab\* OR Spinal cord injury without radiographic abnormality OR SCIWORA) AND (Pediatric\* OR paediatric\* OR peadiatric\* OR child OR children OR childhood OR neonate\* OR newborn\* OR new-born\* OR infant\* OR baby OR babies OR toddler\* OR adolescen\* OR juvenile\* OR youth\* OR teen\* OR preteen\*)))

Additional limits - Source type: Conference Papers & Proceedings, Dissertations & Theses

#### PubMed

Search 24 February 2015. This search contained population terms.

Searched 5 March 2015. This search did not contain population terms.

((NEXUS OR CCR OR National Emergency X-Radiography OR Canadian c-spine OR Canadian cervical spine OR Clinical pathway\* OR critical pathway\* OR treatment pathway\* OR clinical protocol\* OR treatment protocol\* OR algorithm\* OR guideline\* OR decision tree\* OR decision tool\* OR decision rule\* OR triage OR protocol\* OR neurolog\* assess\* OR neurolog\* exam\* OR neurolog\* sign\* OR physical\* exam\* OR physical\* assess\* OR physical\* sign\* OR clinical\* exam\* OR clinical\* assess\* OR clinical\* sign\* OR MRI\* OR CT\* OR Computed Tomography OR CAT scan\*OR X ray\* OR x-ray\* OR xray\* OR radiogra\* OR roentgenogra\* OR imaging) AND (cervical spine clear\* OR c-spine clear\* OR clearing the c-spine OR clearing the cervical spine OR cervical trauma\* OR cervical injur\* OR cervical fracture\* OR cervical sublux\* OR cervical disloc\* OR cervical avuls\* OR cervical instab\* OR Spinal cord injury without radiographic abnormality or SCIWORA) AND (pubstatusaheadofprint OR publisher[sb] or pubmednotmedline[sb]))

### OpenGrey

### Search 24 February 2015

((NEXUS OR CCR OR National Emergency X-Radiography OR Canadian c-spine OR Canadian cervical spine OR Clinical pathway\* OR critical pathway\* OR treatment pathway\* OR clinical protocol\* OR treatment protocol\* OR algorithm\* OR guideline\* OR decision tree\* OR decision tool\* OR decision rule\* OR triage OR protocol\* OR neurolog\* assess\* OR neurolog\* exam\* OR neurolog\* sign\* OR physical\* exam\* OR physical\* assess\* OR physical\* sign\* OR clinical \* exam\* OR clinical\* assess\* OR clinical\* sign\* OR MRI\* OR CT\* OR Computed Tomography OR CAT scan\*OR X ray\* OR x-ray\* OR xray\* OR radiogra\* OR roentgenogra\* OR imaging) AND (cervical spine clear\* OR c-spine clear\* OR clearing the c-spine OR clearing the cervical spine OR cervical trauma\* OR cervical injur\* OR cervical fracture\* OR cervical sublux\* OR cervical disloc\* OR cervical avuls\* OR cervical instab\* OR Spinal cord injury without radiographic abnormality OR SCIWORA))

### ClinicalTrials.gov

#### Search 24 February 2015

((cervical spine OR c-spine) AND (fracture OR injury OR trauma OR avulsion OR dislocation OR instability) AND (NEXUS OR "National Emergency X-Radiography" OR "Canadian c-spine" OR clearance OR decision OR algorithm OR pathway OR triage))

### ICTRP

#### Search 24 February 2015



(NEXUS OR National Emergency X-Radiography OR Canadian c-spine OR Canadian Cervical Spine) AND (cervical fracture OR cervical injury OR cervical trauma OR cervical dislocation OR cervical instability OR cervical avulsion)

### CDSR, DARE, HTA

Searched 25 February 2015

clearance:ti,ab,kw or cervical spine

#### ARIF

Search 24 February 2015

Advanced search, all indexed fields: Clearance or cervical spine

#### **DTA Trials Register**

Searched 10 March 2015

We received the following report from the information specialist of the Renal group:

"There are no studies relating to your review in the DTAS Register. I used keywords from your review title plus other broader target condition words e.g. spinal injur\* spinal trauma, head injur\* etc. I found only 3 studies, all of which were in adults only, and which were using radiological modalities to screen for blunt trauma injuries, including cervical arteries. I also used the test names you mentioned, but did not retrieve anything."

#### Medion

Searched October 2013

ICPC code = Musculoskeletal OR Neurological

And

Abstract = clearance or "cervical spine"

## WHAT'S NEW

Date	Event	Description
22 March 2024	New search has been performed	We performed a search update on 13 December 2022 to identify new studies.
		Inclusion criteria have been expanded to include studies that evaluate the diagnostic accuracy of any clinical decision rule or clinical criteria for the evaluation of cervical spine injury in chil- dren; previously only studies that evaluated the diagnostic accu- racy of NEXUS and Canadian C-spine rule were included.
		The previous version of the review had reference standards of radiographic imaging or clinical follow-up if the index test score was negative or if children did not undergo radiographic imag- ing. Clinical follow-up was considered to be part of the initial trauma evaluation in the emergency department during the first 72 hours. In this updated review, we also included studies where the cervical spine was clinically cleared in the emergency depart- ment by the treating clinician. For children who did not undergo imaging, we preferred to include studies where children were fol- lowed up some time after discharge to ensure no cervical spine injuries were missed.
		We added to the patient selection domain an additional sig- nalling question to check if the data were collected prospective- ly as retrospective data are prone to selective and incomplete recording.



Date	Event	Description
22 March 2024	New citation required and conclusions have changed	Review update includes five studies: one study included in the previous review; one study excluded from the previous review and now included due to expanded eligibility criteria; and three new studies identified in the updated search. Two studies are awaiting classification, with authors contacted for further eligi- bility information, and we identified two ongoing studies.

### HISTORY

Protocol first published: Issue 5, 2015 Review first published: Issue 12, 2017

## CONTRIBUTIONS OF AUTHORS

- Designing the review update: ET, NE, NP, FB
- Co-ordinating the review: ET, NE
- Updating search strategies and undertaking search: Medical Librarian, RCH, Melbourne
- Screening updated search results: ET, VR, NE, JW
- Screening retrieved papers against inclusion criteria: ET, VR, JW, NE
- Appraising quality of papers: ET, NE
- Extracting data from papers: ET, NE
- Writing to authors of papers for additional information: NE
- Obtaining and screening data on unpublished studies: ET, NE
- Data management for the review: ET, NE
- Entering data into Review Manager 5: ET, NE
- Analysis of data: JW, ET, NE
- Interpretation of data: ET, JW, NE
- Providing a methodological perspective: ET, JW, NE
- Providing a clinical perspective: FB, NP
- Writing the review: ET, NE, NP, FB

## DECLARATIONS OF INTEREST

ET: none.

NE: none.

NP: is an author on one of the included studies but was not involved in the screening of papers and data extraction (Phillips 2021).

JW: none.

VR: none.

FB: is an author on one of the included studies but was not involved in the screening of papers and data extraction (Phillips 2021).

# SOURCES OF SUPPORT

## **Internal sources**

• Murdoch Children's Research Institute, Australia

Infrastructure supportRoyal Children's Hospital Foundation, Australia

Review author support (FB)



### **External sources**

• National Health and Medical Research Council, Australia

Centre of Research Excellence grant for Paediatric Emergency Medicine (GNT1058560)

## DIFFERENCES BETWEEN PROTOCOL AND REVIEW

We made the following changes from the protocol (Slaar 2015).

We planned to meta-analyse the sensitivity and specificity of the tools with a bivariate model. However, we identified only five studies that met the inclusion criteria, and the outcomes of the studies were too diverse for us to perform meta-analyses in this review. For the same reason, an analysis of heterogeneity could not be completed.

We did not anticipate in the protocol that we would encounter studies with mixed populations (adults and children) in which we could not extract the data for both groups ourselves. When this occurred during the review process, we attempted to contact the authors to obtain these data.

For this update, the inclusion criteria for studies was expanded to include all clinical decision rules rather than just NEXUS and the Canadian C-Spine Rule and weaker study designs (case-control studies) were excluded.

## INDEX TERMS

### **Medical Subject Headings (MeSH)**

Cervical Vertebrae [diagnostic imaging] [\*injuries]; Checklist; Cohort Studies; \*Decision Support Techniques; Magnetic Resonance Imaging; Radiography; Reference Standards; Spinal Injuries [\*diagnosis] [diagnostic imaging] [etiology]; Tomography, X-Ray Computed; Triage [\*methods]; Wounds, Nonpenetrating [\*complications] [diagnostic imaging]

### **MeSH check words**

Child; Humans