Reference values for Cobb angles when evaluating the spine in the sagittal plane: a systematic review with meta-analysis

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REVIEW ARTICLE

ABSTRACT

The present systematic review of observational studies with meta-analysis aim to identify the reference values of the spinal curvatures in the sagittal plane, as evaluated using the Cobb angle in X-rays, in healthy individuals. Electronic searches were undertaken in MEDLINE, Scopus, ScienceDirect and LILACS. Studies that evaluated the spinal curvature of healthy children, adolescents, adults, and elderly using Cobb method and presented reference values for those curvatures were incluced. Thirty-one studies were eligible for inclusion. The reference values found (confidence interval 95%) were: for children, thoracic (28.7°-37.9°), lumbar L1-L5 (34.5°-44.8°), and lumbar L1-S1 (41.7°-54.1°); for adolescents: thoracic (31.5°-39.2°), lumbar L1-L5 (39.8°-45.6°), and lumbar L1-S1 (51.9°-59.1°); for adults: thoracic (37.7°-40.3°), lumbar L1-L5 (38.1°-45.6°), and lumbar L1-S1 (54.2°-61.7°); and for the elderly: thoracic (37.7°-50.4°), and lumbar L1-S1 (56.6°-65.9°). For the cervical region, it was impossible to establish consistent reference values. The present study supports that precise reference intervals were identified for the sagittal curvatures of the thoracic and lumbar spine in healthy children, adolescents, adults and elderly, as evaluated by means of the Cobb Method.

Keywords: kyphosis, lordosis, radiograph, reference values.

INTRODUCTION

individuals without musculoskeletal In dysfunction and apparently healthy, the spine is usually structured by successive opposed convexities in the sagittal plane (Perriman et al., 2010). The magnitude of each curve varies from individual to individual, however, when maintaining the upright position, they should result from a balance of muscular forces and external loads (Singh, Bailey & Lee, 2010). An accentuated curvature may cause the intervertebral disks to protrude, while an excessively straight spine may lead to overload on "articular facets" (Singh, Bailey & Lee, 2010). Thus, the spine should be neither excessively curved nor straight, but within a range of normality. Based on values that quantify the deviation and its progression (for example, curvature), quantitative increasing spinal postural evaluation procedures are used to identify subtle alterations to the spinal curvature or accompany treatment (Oliveira et al., 2012).

The sagittal curvature of the spine can be measured using several invasive and/or noninvasive postural evaluation methods. The gold standard evaluation method for the sagittal plane is the latero-lateral X-ray (Zaina, Donzelli, Lusini & Negrini, 2012), in which the Cobb angles, represented by the crossing of tangents originating from the cranial and caudal vertebral bodies, are calculated (Briggs et al., 2007; Goh, Price, Leedman & Singer, 2000).

Regarding the range of normality, the literature is extremely divergent. For example, in the lumbar spine, the range of normality is found to vary from 26° to 58° Cobb-angle (Prospst-Proctor & Blac, 1983), or from 13° to 78° Cobb-angle (Bernhardt & Bridwell, 1989). Thus, the range of sagittal spinal curvature considered to be within the normal range when measured using the Cobb method is very wide, varying by up to 60° (Briggs et al., 2007; Chaise et al., 2011; Goh et al., 2000; Oliveira et al., 2012; Zaina et al., 2012). Hence, the absence of adequate

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parameters may compromise the diagnosis and, consequently, the feasibility of effective intervention (Zubovic, Davies, Berryman & Pynsent, 2008).

It should be noted that this broad spectrum of normal curvature values is due to the variability of the human being, while also being dependent on, for example, age range and the vertebral bodies considered when calculating the curvature. Therefore, it would be useful to objectively synthetize the normal values arising from studies that have adopted the Cobb measurement in healthy individuals. Moreover, in clinical practice, the definition of a narrower range of normality, mainly for children and adolescents, would increase the chances of early diagnosis and might improve preventive strategies and interventions (Monticone et al., 2014).

Therefore, based on a systematic review with meta-analysis, this study aims to identify the reference values of the spinal curvature in the sagittal plane, as evaluated using the Cobb angle in X-rays, in healthy individuals.

METHOD

Identification and selection of studies

A systematic review with meta-analysis of observational studies presenting reference values of the sagittal curvature of the spine, determined using the Cobb angle in X-rays was conducted. This systematic review followed the recommendations proposed by the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) (Moher, Liberati, Tetzlaff & Altman, 2009). The Project was registered in the PROSPERO of the Centre for Reviews and Dissemination at the University of York (http://www.crd.york.ac.uk/PROSPERO), under the number CRD42015025691.

A systematic search was carried out by two independent investigators and in duplicate for studies in the following databases: MEDLINE (Medical Literature Analysis and Retrieval System Online) accessed by PubMed; Scopus; ScienceDirect; and LILACS (Literatura Latino-Americana e do Caribe em Ciências da Saúde) accessed by BVS (Biblioteca Virtual em Saúde), in the month of June 2016. The search strategy adopted in the PubMed data base is presented in Table 1. No year of publication restriction was used in the systematic search. The studies should be written in English, Spanish or Portuguese.

The PICO model was adopted to establish the inclusion criteria, where the "P" (Patient, Population, or Problem) is defined as children, adolescents, adults and the elderly; the "I" (Intervention, Prognostic Factor, or Exposure) is the sagittal curvature assessment based on applying the Cobb method to X-ray images; the "C" (Comparison or Intervention) is not applicable to this review; and the "O" (Outcome) is the range of normal values regarding the sagittal curvature of the cervical, thoracic and lumbar spine.

Thus, the inclusion criteria were studies that: 1) evaluated the spinal curvature using the twoor four-line Cobb method; 2) evaluated the sagittal curvatures of the spine (lumbar, thoracic or cervical); and 3) presented reference values for those curvatures. The exclusion criteria were studies that: 1) solely revised the literature; and 2) evaluated non-healthy populations with specific pathologies.

Table 1

Search strategy conducted in the PubMed*

#1	"Lordosis" [Mesh] OR "Kyphosis" [Mesh] OR "Kyphoses "Cervical" OR "Thoracic" OR "Lumbar"	" OR "Spinal	Curvatures" OR	"Sagittal	Curvatures" OF	"Spine C	urvatures" OR
		" OD (D	1.1 D. (' VD "	OD ((D)	VD	D 1:1 " OD

"Radiography" [Mesh] OR "Radiology, Diagnostic X Ray" OR "Radiology, Diagnostic X-Ray" OR "Diagnostic X Ray Radiology" OR
"Diagnostic X-Ray Radiology" OR "X Ray Radiology, Diagnostic" OR "X-Ray Radiology, Diagnostic" OR "Roentgenography" OR "X Ray, Diagnostic" OR "X-Ray, Diagnostic" OR "X-Rays, Diagnostic" OR "Diagnostic X-Ray" OR "Diagnostic X Ray" OR "Diagnostic X-Ray" OR "X rays" OR "X-ray" OR "X-rays" OR "Radiologic" OR "Radiologic" OR "Radiologic" OR "Radiologic" OR "Radiologic" OR "Cobb"

"Reference Values" [Mesh] OR "Reference Value" OR "Values, Reference" OR "Value, Reference" OR "Range, Reference" OR "Reference Ranges"
#3 OR "Values, Normal" OR "Value, Normal" OR "Normal Value" OR "Normal Values" OR "Ranges, Normal" OR "Range, Normal" OR "Normal Value" OR "Normal Values" OR "Ranges, Normal" OR "Range, Normal" OR "Normal Ranges" OR "Normal Ranges" OR "Normal Range"

#4 #1 AND #2 AND #3

Note. *The search was conducted using title/abstract/study keywords.

The titles and abstracts of the studies identified using the search strategy were independently assessed in duplicate by two investigators (T.S.F and J.A.S). Those studies in which the abstracts contained sufficient information regarding the inclusion and exclusion criteria were selected for full evaluation of the article. In the second stage, the same investigators, independently and in duplicate, assessed the full texts and made the selection eligibility criteria. according to the Disagreements between investigators were resolved by consensus, and when disagreements persisted, a third investigator (C.T.C.) evaluated the article. In addition, the reference sections of the selected studies were checked in an attempt to find suitable studies not revealed by the electronic search.

Assessment of characteristics of studies

Two investigators (T.S.F. and J.A.S.), independently and in duplicate, extracted the data referring to the methodological characteristics and findings, and evaluated the risk of bias. Disagreements were resolved by consensus or by a third investigator (C.T.C.). Using a standard form, the following information was extracted: spine regions assessed, level used to calculate the Cobb angle, assessed population and reference values reported for spinal curvatures.

The studies that fulfilled the inclusion criteria were assessed regarding risk of bias using the Guidelines for Critically Appraising Studies, which consists of a check list of 8 criteria that can be answered "yes", "no" or "impossible to determine" (Loney, Chambers, Bennett, Roberts & Stratford, 1998). This checklist was developed and validated to critically appraise research studies that estimate the prevalence or incidence of a disease or health problem (Loney et al., 1998), and investigates three main areas: 1) the internal validity (design, sampling frame, sample size, outcome measures, measurement and response rate); 2) the interpretation of the results; and 3) the applicability of the findings. The risk of bias of the studies is presented according to the total number of criteria classified

as "yes" in each article. Each criterion was given a point and each study was scored out of a total of eight (Davoren, Demant, Shiely & Perry, 2016).

Data analysis

To better organize the extracted data, with a view to the meta-analysis, the studies were grouped according to the assessed population: children (under 12 years), adolescents (between 12 and 17 years), adults (between 18 and 59 years) and the elderly (60 plus). Also, in the lumbar region, the studies were grouped according to the vertebral levels used in the evaluation of the Cobb angle: between vertebra L1 and L5, and between vertebra L1 and S1. This division according to vertebra levels in lumbar spine came about because the Cobb angles calculated based on a sacral vertebra presented a considerable discrepancy in relation to those calculated based on lumbar vertebrae. The large inclination of the sacrum in relation to lumbar vertebrae is mainly responsible for this discrepancy (Marty et al., 2002). Thus, the groups were: 1) Cervical Children, 2) Cervical Adolescents, 3) Cervical Adults, 4) Cervical Elderly, 5) Thoracic Children, 6) Thoracic Adolescents, 7) Thoracic Adults, 8) Thoracic Elderly, 9) Lumbar Children L1-L5, 10) Lumbar Children L1-S1, 11) Lumbar Adolescents L1-L5, 12) Lumbar Adolescents L1-S1, 13) Lumbar Adults L1-L5, 14) Lumbar Adults L1-S1, 15) Lumbar Elderly L1-L5, and 16) Lumbar Elderly L1-S1.

The Comprehensive Meta-Analysis software (CMA, www.metaanalysis.com) was used in the meta-analysis. The studies were grouped according to the assessed population and, only in the case of lumbar curvature, the level of the vertebra used to evaluate the Cobb angle. The means and 95% confidence intervals (CI) were calculated for the angles of the cervical, thoracic and lumbar curvatures with random effect models for the different populations. The Z test was used to evaluate the statistical significance of the means and the confidence interval for the analyzed curvatures. The significance value of 0.05 was adopted.

The statistical heterogeneity for each metaanalysis was assessed using the Cochran Q test and inconsistency test I^2 . For the purpose of interpretation, an I^2 value close to 0% indicates no heterogeneity among the studies, close to 25% indicates low heterogeneity, close to 50% indicates moderate heterogeneity, and close to 75% indicates high heterogeneity among the studies (Higgins & Thompson, 2002; Higgins, Thompson, Deeks & Altman, 2003).

RESULTS

Flow of studies through the review

Of the 1785 potentially relevant studies recovered from the electronic data bases and found in the references, 31 met the inclusion criteria (Figure 1).



Figure 1. Flowchart of the included studies

Characteristics of studies

Table 2 summarizes the main features of the included studies: age, number of subjects and the protocols for the X-ray examination and Cobb angle calculation. In relation to age, most of the studies assessed young adults, while the least assessed population was the elderly. The good reproducibility of the two- or four-line Cobb method was confirmed in most of analyzed studies.

Table 3 shows the reference values (in Cobb degrees) of the sagittal curves. Few studies assessed the cervical curvature, with only one study being found involving children and

adolescents (Kasai, Ikata, Katoh, Miyake & Tsubo, 1996). Therefore, it was not considered in the meta-analysis. The lumbar curvature of the elderly population was only assessed using L1-S1 vertebral levels. Therefore, in the elderly, vertebral level L1-L5 was not used in the meta-analysis. In general, the Cobb angles calculated on basis of the sacral vertebra (L1-S1) are greater than those calculated on lumbar vertebrae alone (L1-L5). This issue is raised in the studies by Champain *et al.* (2006) and Damasceno *et al.* (2006) which analyze the same sample using different methods (Table 3).

Table 2

Features of the studies: age, number of subjects and the protocols for the X-ray examination and Cobb angle calculation

5	0 /	J	1 1 1	8
First author (year)	Age* (years old)	Number of subjects	X-rays	Cobb angle
Le Huec (2015)	38.0 (18-76)	106	Upright, looking forward and their fingertips on their clavicles	Digital, two observers, excellent reliability
Zhu (2013)	34.3±12.6	260	Upright, horizontal arms and resting on supports	Digital, excellent reliability
Endo (2012)	31.5 ± 7.4	50	Upright, hands in front of their trunk	Digital, two observers, excellent reliability
Lee (2012)	11.7 ± 4.4	181	Upright, arms elevated 90°	Digital, good reliability
Yukawa (2012)	20-70	1230	Upright, looking forward	Digital, experienced observer
Chanplakorn (2011)	33.3±6.8	100	Upright, hands hold together behind the neck	Manual, two observers, excellent reliability
Lee (2011)	28.19	86	Upright, horizontal arms and resting on supports	Manual (protractor and ruler), two observers, good reliability
Janssen (2009)	26.5	60	Upright, looking forward , arms elevated 45°	Automatically after 3D digital reconstruction of the vertebrae, excellent reliability
Gonçalves (2008)	15.8 ± 1.4	22	Upright, looking forward	Manual
Champain (2006)	43±10.5	60	Not explained	Digital, two observers, good reliability
Damasceno (2006)	29.0±8.24	350	Upright, arms resting on supports	Manual, two observers, good reliability
Cil (2005)	3-15	151	Upright, arms elevated 30°	Digital, same observer, good reliability
Vialle (2005)	35.4±12	300	Upright, shoulders flexed on 45° and resting on supports	Digital, three observers
Hammerberg (2003)	76.3 ± 4.1	50	Upright, horizontal arms and resting on supports	Low inter-observer error
Boseker (2000)	5-19	121	Upright, horizontal arms and resting on supports	Experienced observer
Korovessis (1999)	20-79	120	Upright, horizontal arms and resting on support	Two observers, excellent reliability
Chernukha (1998)	1-30	199	Supine horizontal	Three observers, good reliability
Jackson (1998)	39.4±9.45	50	Upright, arms below chest level and resting on support	Digital, two observers, good reliability
Korovessis (1998)	52.7±15	99	Upright, horizontal arms and resting on support	Two observers, strong agreement
Vedantam (1998)	13±8	88	Upright, arms elevated 60°	Two observers, low error between measurements
Hardacker (1997)	38.5±9.3	100	Upright, looking forward, flexion of shoulders and resting on support	Single observer
Troyanovich (1997)	27.1±8	50	Upright, hands on top of the head	Manual, good reliability
Kasai (1996)	1-18	360	Not explained	Not explained
Gelb (1995)	57±11	100	Upright, horizontal arms	One or two observers, low error between measurements
Lin (1992)	50	149	Recumbent position with hip in 45° flexion	Single observer
Wright (1991)	12	112	Upright	Single observer
Singer (1990)	15-93	286	Upright	Manual and digital
Bernhardt (1989)	12.8	102	Upright, horizontal arms and resting on support	Not explained
Voutsinas (1986)	5-20	670	Upright, flexion of shoulders and resting on support	Single observer, good reliability
Propst-Proctor (1983)	2-19	104	Upright, arms elevated 90°	Manual
Fon (1980)	30.63 (6-75)	316	Upright, horizontal arms	Manual, good reliability
Note *Age is described	as presented i	n the studies	e meaner means + standard deviat	ion, mean (minimum maximum), or

Note. *Age is described as presented in the studies: means; means ± standard deviation; mean (minimum-maximum); or minimum-maximum

Table 3

Data extracted from the studies with reference values (in Cobb degrees) for the sagittal curves

First author (year)	Cervical	Cervical	Cervical	Cervical	Thoracic	Thoracic	Thoracic	Thoracic	Lumbar children		Lumbar adolescents		Lumbar adults		Lumbar elderly
	children	adolescents	aduits	elderly	children	adolescents	adults	elderly	L1-L5	L1-S1	L1-L5	L1-S1	L1-L5	L1-S1	L1-S1
Le Huec (2015)			4.9 ± 12.8	4.9 ± 12.8											
Zhu (2013)							27.8 ± 11.4							48.2 ± 9.6	
Endo (2012)													33.3±11.3		
Lee (2012)					31.7±9	34.9 ± 9.4				48 ± 10.5		49.6 ± 9.9			
Yukawa (2012)			11.5 ± 11.3	18.7±11											
Chanplakorn (2011)														54.7 ± 9.9	
Lee (2011)							32 ± 9.2							49.6 ± 9.6	
Janssen (2009)							36 ± 8.9							58.5 ± 9.6	
Gonçalves (2008)											44.4 ± 6.9	57.5 ± 6.1			
Champain (2006)							42.5 ± 18.4						49 ± 18	57±22	
Damasceno (2006)													45.1 ± 10.8	60.9 ± 10.7	
Cil (2005)					46±10.8	53.3 ± 9.1				50.1 ± 10.8		54.6 ± 9.8			
Vialle (2005)							40.6±10						43±11.2		
Hammerberg (2003)								52.5 ± 12.2							57.4±13.7
Boseker (2000)					33±10.6	33 ± 10.6									
Korovessis (1999)							36.5 ± 9	49 ± 13.9						62.5 ± 13	62 ± 20.9
Chernukha (1998)										39 ± 8.6		51.1 ± 8.8		52.6 ± 11.6	
Jackson (1998)							47 ± 9.7							62.1 ± 10.8	
Korovessis (1998)							41.8 ± 13						45.7±12		
Vedantam (1998)						38±10						64±10			
Hardacker (1997)							49.4 ± 10.9							60.1 ± 12.1	
Troyanovich (1997)														65 ± 10.6	
Kasai (1996)	13.5 ± 4.7	14.3 ± 6.6													
Gelb (1995)							34±11	34±11						64±10	64±10
Lin (1992)													33.2±12.1		
Wright (1991)									35±10						
Singer (1990)						33.2 ± 6.8	31.6±11.4	45.3 ± 10.2							
Bernhardt (1989)					36±10	36±10	36±10		44±12		44±12		44±12		
Voutsinas (1986)					37.1±7.5	38 ± 8.1				54.2 ± 10.2		56.4 ± 8.7			
Propst-Proctor (1983)					27±10.6	27±10.6			40±11		40±11				
Fon (1980)					22.3 ± 7.8	25.5 ± 7.8	30.1±7.8	39.9 ± 7.4							

Risk of bias in included studies

In the risk of bias assessment (Table 4), the criteria that involve measurement issues (criterion 4) and confidence intervals (criterion 7) were found in all the included studies. In 28 studies (90.3%), the response rate was adequate (criterion 6); in 26 studies (83.9%) there are no unbiased assessors (criterion 5); and 25 studies

(80.6%) presented detailed subject descriptions (criterion 8). Only 13 studies (40.3%) presented an unbiased sampling frame (criterion 2) and 2 studies (6.4%) presented a random sample (criterion 1). None of the studies scored in the third scale criterion (sample size), because they did not provide a sample size calculation or justification.

Table 4

Risk of bias assessment using the guidelines for critically appraising studies

Studies	Criteria Guidelines for Critically Appraising Studies								
1 st author (year)	1	2	3	4	5	6	7	8	Total (Nr of ✓)
Le Huec (2015)	Х	\checkmark	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	6
Zhu (2013)	Х	\checkmark	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	6
Endo (2012)	Х	Х	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	5
Lee (2012)	Х	\checkmark	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	6
Yukawa (2012)	Х	Х	Х	\checkmark	?	\checkmark	\checkmark	\checkmark	4
Chanplakorn (2011)	Х	Х	Х	\checkmark	\checkmark	?	\checkmark	\checkmark	4
Lee (2011)	Х	Х	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	5
Janssen (2009)	Х	Х	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	5
Gonçalves (2008)	Х	Х	Х	\checkmark	?	\checkmark	\checkmark	\checkmark	4
Champain (2006)	Х	Х	Х	\checkmark	\checkmark	\checkmark	\checkmark	Х	4
Damasceno (2006)	Х	Х	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	5
Cil (2005)	Х	\checkmark	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	6
Vialle (2005)	Х	Х	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	5
Hammerberg (2003)	Х	Х	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	5
Boseker (2000)	Х	\checkmark	Х	\checkmark	?	\checkmark	\checkmark	\checkmark	5
Korovessis (1999)	Х	Х	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	5
Chernukha (1998)	Х	\checkmark	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	6
Jackson (1998)	Х	Х	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	5
Korovessis (1998)	Х	Х	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	5
Vedantam (1998)	Х	\checkmark	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	6
Hardacker (1997)	Х	Х	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	5
Troyanovich (1997)	Х	?	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	5
Kasai (1996)	\checkmark	Х	Х	\checkmark	\checkmark	?	\checkmark	Х	4
Gelb (1995)	Х	Х	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	5
Lin (1992)	Х	\checkmark	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	6
Wright (1991)	Х	Х	Х	\checkmark	\checkmark	\checkmark	\checkmark	Х	4
Singer (1990)	\checkmark	\checkmark	Х	\checkmark	\checkmark	\checkmark	\checkmark	Х	6
Bernhardt (1989)	Х	\checkmark	Х	\checkmark	?	?	\checkmark	?	3
Voutsinas (1986)	Х	\checkmark	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	6
Propst-Proctor (1983)	Х	\checkmark	Х	\checkmark	?	\checkmark	\checkmark	Х	4
Fon (1980)	Х	\checkmark	Х	\checkmark	\checkmark	\checkmark	\checkmark	\checkmark	6

Note. Criteria Checklist Critical Appraisal of Studies: 1) Are the study design and sampling method appropriate for the research question? 2) Is the sampling frame appropriate? 3) Is the sample size adequate? 4) Are the objective and standard criteria used for measurement of the health outcome suitable? 5) Is the health outcome measured in an unbiased fashion? 6) Is the response rate adequate? Are the refusers described? 7) Are the estimates of prevalence or incidence given with confidence intervals and in detail by subgroup, if appropriate? 8) Are the study subjects and the setting described in detail and similar to those of interest to you? ANSWERS: \checkmark = Yes; X = No; ? = Unable to determine.

Meta-analyses of Exposure

The meta-analysis of the child population presents the mean of the results found in the literature for the Cobb values of the thoracic and lumbar curves and the 95% CI (Table 5). None of the meta-analyses carried out for the sagittal curves presented heterogeneity. In the thoracic spine, the z value was 14.182 (p<0.001), while in the lumbar spine, the z values were 15.068 (p<0.001) and 15.237 (p<0.001), demonstrating that the means and confidence intervals of the reference values for both curvatures are statistically significant in children.

Table 5

Mean Meta-analysis for the reference values of the thoracic and lumbar spine curvatures in children

			St	atistics for e	ach study				
Curve	Study Name	Sample size	Mean	Standard Error	Variance	Lower limit	Upper limit	Z-Value	p-Value
	Lee (2012)	99	31.7	9	0.818	29.9	33.5	35.046	< 0.001
U	Cil (2005)	120	46	10.8	0.972	44.1	47.9	46.658	< 0.001
	Boseker (2000)	121	33	10.6	0.929	31.1	34.9	34.245	< 0.001
rac	Bernhardt (1989)	102	36	10	0.980	34.1	37.9	36.358	< 0.001
ioh	Voutsinas (1986)	670	37.1	7.5	0.084	36.5	37.7	128.041	< 0.001
H	Propst-Proctor (1983)	104	27	10.6	1.080	25.0	29.0	25.976	< 0.001
	Fon (1980)	49	22.3	7.8	1.242	20.1	24.5	20.013	< 0.001
			33.3	2.4	5.522	28.7	37.9	14.182	< 0.001
Hete	rogeneity: Tau ² =0.000; Q=	2.075, df=6	(p=0.55)	7); I ² =0.000	%				
ч	Wright (1991)	112	35	10	0.893	33.1	36.9	37.041	< 0.001
ıba -L5	Bernhardt (1989)	102	44	12	1.412	41.7	46.3	37.032	< 0.001
L un	Propst-Proctor (1983)	104	40	11	1.163	37.9	42.1	37.084	< 0.001
			39.6	2.6	6.917	34.5	44.8	$\begin{array}{c cccc} Z-Value & p-V\\ \hline 35.046 & <0.\\ 46.658 & <0.\\ 34.245 & <0.\\ 36.358 & <0.\\ 128.041 & <0.\\ 25.976 & <0.\\ 20.013 & <0.\\ 14.182 & <0.\\ \hline 37.041 & <0.\\ 37.032 & <0.\\ 37.084 & <0.\\ 15.068 & <0.\\ \hline 32.386 & <0.\\ 15.237 & <0.\\ \hline \end{array}$	< 0.001
Hete	rogeneity: Tau ² =0.000; Q=	0.661, df=2	(p=0.882	2); I ² =0.000	%				
	Lee (2012)	99	48	10.5	1.114	45.9	50.1	45.485	< 0.001
) 1	Cil (2005)	120	50.1	10.8	0.972	48.2	52.0	50.816	< 0.001
l-S	Chernukha (1998)	51	39	8.6	1.450	36.6	41.4	32.386	< 0.001
(El	Voutsinas (1986)	670	54.2	10.2	0.155	53.4	55.0	137.542	< 0.001
			47.9	3.1	9.9	41.7	54.1	15.237	< 0.001
Hete	rogeneity: $Tau^2 = 0.000$; Q=	2.155, df=3	(p=0.82)	7); $I^2 = 0.000$	%				

Table 6

Meta-analysis for the reference values of the thoracic and lumbar spine curvatures in adolescents

			St	tatistics for e	ach study				
Curve	Study Name	Sample size	Mean	Standard Error	Variance	Lower limit	Upper limit	Z-Value	p-Value
	Lee (2012)	82	34.9	9.4	1.078	32.9	36.9	33.621	< 0.001
	Cil (2005)	31	53.3	9.1	2.671	50.1	56.5	32.611	< 0.001
	Boseker (2000)	121	33	10.6	0.929	31.1	34.9	34.245	< 0.001
. <u>.</u>	Vedantam (1998)	88	38	10	1.136	35.9	40.1	35.647	< 0.001
ac	Singer (1990)	13	33.2	6.8	3.557	29.5	36.9	17.604	< 0.001
ioh	Bernhardt (1989)	102	36	10	0.980	34.1	37.9	36.358	< 0.001
H	Voutsinas (1986)	670	38	8.1	0.098	37.4	38.7	121.433	< 0.001
	Propst-Proctor (1983)	104	27	10.6	1.080	25.0	29.0	25.976	< 0.001
	Fon (1980)	49	25.5	7.8	1.217	23.3	27.7	23.117	< 0.001
			35.4	1.9	3.806	31.5	39.2	18.132	< 0.001
Hete	rogeneity: Tau ² =0.000; Q=3	3.678, df=8	(p=0.349	9); I ² =0.000%	6				
Lumbar (L1-L5)	Gonçalves (2008)	22	44.4	6.9	2.164	41.5	47.3	30.182	< 0.001
	Bernhardt (1989)	102	44	12	1.412	41.7	46.3	37.032	< 0.001
L ju	Propst-Proctor (1983)	104	40	11	1.163	37.9	42.1	37.084	< 0.001
			42.7	1.5	2.196	39.8	45.6	28.820	< 0.001
Hete	rogeneity: Tau ² =0.000; Q=-	4.786, df=2	(p=0.912	2); I ² =0.000%	6				
	Lee (2012)	82	49.6	9.9	1.195	47.5	51.7	45.368	< 0.001
-S1	Gonçalves (2008)	22	57.5	6.1	1.691	51.9	59.1	44.213	< 0.001
Ē	Cil (2005)	31	54.6	9.8	3.098	51.2	58.1	31.020	< 0.001
, Ч	Chernukha (1998)	108	51.1	8.8	0.717	49.4	52.8	60.346	< 0.001
Iba	Voutsinas (1986)	670	56.4	8.7	0.113	55.7	57.1	167.802	< 0.001
un,	Vedantam (1998)	88	64	10	1.136	61.9	66.1	60.037	< 0.001
Г			55.5	1.8	3.355	51.9	59.1	30.318	< 0.001
Hete	rogeneity: Tau ² =0.000; Q=2	2.212, df=5	(p=0.697	7); I ² =0.000%	6				

The meta-analysis of the adolescents presents the mean of the results found in the literature for the Cobb values of the thoracic and lumbar curves and the 95% CI (Table 6). None of the metaanalyses carried out for the sagittal curves presented heterogeneity. In the thoracic spine, the z value was 18.132 (p<0.001), while in the lumbar spine, the z values were 28.820 (p<0.001) and 30.318 (p<0.001), demonstrating that the means and confidence intervals of the reference values for both curvatures are statistically significant in adolescents. The meta-analysis of the adult population presents the mean results for the Cobb values of the cervical, thoracic and lumbar curves and the 95% CI (Table 7). None of the meta-analyses presented heterogeneity. In the cervical spine, the z value was 2.519 (p=0.012) and in the thoracic spine, the z value was 22.144 (p<0.001). While in the lumbar spine, the z values were 21.842 (p<0.001) and 30.230 (p<0.001), demonstrating that the means and confidence intervals of the reference values for the curvatures are statistically significant in adults.

Table 7

Meta-analysis for the reference values of the cervical, thoracic and lumbar spine curvatures in adults

			S	Statistics for	each study				
Curve	Study Name	Sample	Mean	Standard	Variance	Lower	Upper	7-Value	n-Value
	Brudy Hume	size	Mean	Error	variance	limit	limit	2 Value	p varue
	Le Huec (2015)	106	4.9	12.8	1.546	2.5	7.3	3.941	< 0.001
Cervical	Yukawa (2012)	1230	11.5	11.3	0.104	10.9	12.1	35.692	< 0.001
			8.3	3.3	10.878	1.9	14.8	2.519	0.012
Hetero	geneity: Tau ² =0.000; Q=	0.257, df=1	(p=0.61)	2); $I^2 = 0.000$	%				
	Zhu (2013)	260	27.8	11.4	0.500	26.4	29.2	39.321	< 0.001
	Lee (2011)	86	32	9.2	0.984	30.1	33.9	32.256	< 0.001
	Janssen (2009)	60	36	8.9	1.320	33.7	38.3	31.332	< 0.001
	Champain (2006)	60	42.5	18.4	5.643	37.8	47.2	17.891	< 0.001
	Vialle (2005)	300	40.6	10	0.333	39.5	41.7	70.321	< 0.001
	Korovessis (1999)	80	36.5	9	1.013	34.5	38.5	36.274	< 0.001
Thoracic	Jackson (1998)	50	47	9.7	1.882	44.3	49.7	34.262	< 0.001
moracie	Korovessis (1998)	99	41.8	13	1.707	39.2	44.4	31.993	< 0.001
	Vedantam (1998)	100	34	10	1.000	32.0	36.0	34.000	< 0.001
	Hardacker (1997)	100	49.4	10.9	1.188	47.3	51.6	45.321	< 0.001
	Gelb (1995)	100	34	11	1.210	31.8	36.2	30.909	< 0.001
	Singer (1990)	147	31.6	11.4	0.884	29.8	33.4	33.608	< 0.001
	Bernhardt (1989)	102	36	10	0.980	34.1	37.9	36.358	< 0.001
	Fon (1980)	316	28.9	8.3	0.317	29.0	31.2	53.472	< 0.001
			37.0	1.7	2.794	33.7	40.3	22.144	< 0.001
Hetero	geneity: Tau ² =0.000; Q=	4.726, df=1	3 (p=0.5	79); I ² =0.00	0%				
	Endo (2012)	50	33.3	11.2	2.554	30.2	36.4	20.838	< 0.001
	Champain (2006)	60	49	18	5.400	44.4	53.6	21.086	< 0.001
Lumbar	Damasceno (2006)	350	45.1	10.8	0.333	44.0	46.2	78.124	< 0.001
(I 1 I 5)	Vialle (2005)	300	43	11.2	0.418	41.7	44.3	66.498	< 0.001
(L1-L3)	Korovessis (1998)	99	45.7	12	1.455	43.3	48.1	37.892	< 0.001
	Lin (1992)	149	33.2	12.1	0.983	31.3	35.1	33.492	< 0.001
	Bernhardt (1989)	102	44	12	1.412	41.7	46.3	37.032	< 0.001
			41.8	1.9	3.666	38.1	limit2 + 0.000 7.3 3.941 <0.001	< 0.001	
Hetero	geneity: Tau ² =0.000; Q=	1.466, df=6	(p=0.96	2); I ² =0.000	%				
	Zhu (2013)	260	48.2	9.6	0.354	47.0	49.4	80.959	< 0.001
	Chanplakorn (2011)	100	54.7	9.9	0.980	52.8	56.6	55.253	< 0.001
	Lee (2011)	86	49.6	9.6	1.072	47.6	51.6	47.914	< 0.001
	Janssen (2009)	60	58.5	9.6	1.536	56.1	60.9	47.202	< 0.001
	Champain (2006)	60	57	22	8.067	51.4	62.6	20.069	< 0.001
Lumbar	Damasceno (2006)	350	60.9	10.7	0.327	59.8	62.0	106.480	< 0.001
(L1-S1)	Korovessis (1999)	80	62.5	13	2.113	59.7	65.3	43.001	< 0.001
	Chernukha (1998)	40	52.6	11.6	3.364	49.0	56.2	28.679	< 0.001
	Jackson (1998)	50	62.1	10.8	2.333	59.1	65.1	40.659	< 0.001
	Hardacker (1997)	100	60.1	12.1	1.464	57.7	62.5	49.669	< 0.001
	Troyanovich (1997)	50	65	10.6	2.247	62.1	67.9	43.360	< 0.001
	Gelb (1995)	100	64	10	1.000	62.0	66.0	64.000	< 0.001
			57.9	1.9	3.671	54.2	61.7	30.230	< 0.001
Hetero	geneity: Tau ² =0.000; Q=	6.550, df=1	1 (p=0.9	24); I ² =0.00	0%				

The meta-analysis of the elderly population presents the mean results for the Cobb values of the curves and the 95% CI (Table 8). None of the meta-analyses presented heterogeneity. In the thoracic spine, the z value was 13.597 (p<0.001), while in the lumbar spine, the z value was 25.639 (p<0.001), demonstrating that the means and

confidence intervals of the reference values for curvatures are statistically significant in the elderly. By contrast, in the cervical spine, the z value was 1.718 (p=0.086), which demonstrates that the mean and the confidence interval of the normality values are not statistically significant.

Table 8

Meta-analysis for the reference values of the cervical, thoracic and lumbar spine curvatures in the elderly

			St	atistics for e	ach study				
Curve	Study Name	Sample size	Mean	Standard Error	Variance	Lower limit	Upper limit	Z- Value	p-Value
	Le Huec (2015)	106	4.9	12.8	1.546	2.5	7.3	3.941	< 0.001
Cervical	Yukawa (2012)	1230	18.7	11.2	0.102	18.1	19.3	58.557	< 0.001
			11.9	6.9	47.607	-1.7	25.4	1.718	0.086
Heterogeneity: Tau ² =0.000; Q=0.257, df=1 (p=0.612); I ² =0.000%									
	Hammerberg (2003)	50	52.5	12.2	2.977	49.1	55.9	30.429	<0.001
	Korovessis (1999)	40	49	13.9	4.830	44.7	53.3	22.295	< 0.001
Thoracic	Gelb (1995)	100	34	11	1.210	31.8	36.2	30.909	< 0.001
Thoracte	Singer (1990)	126	45.3	10.2	0.826	43.5	47.1	49.852	< 0.001
	Fon (1980)	25	39.9	7.4	2.190	37.0	42.8	26.959	< 0.001
			44.0	3.2	10.488	37.7	50.4	13.597	< 0.001
Heterog	geneity: Tau ² =0.000; Q=2	.765 df=4 (p=0.554)	; I ² =0.000%					
Lumber	Hammerberg (2003)	50	57.4	13.7	3.754	53.6	61.2	29.626	< 0.001
(11 S1)	Korovessis (1999)	40	62.0	20.9	10.920	55.5	68.5	18.762	< 0.001
(LI-51)	Gelb (1995)	100	64	10	1.000	62.0	66.0	64.000	< 0.001
			61.2	2.4	5.705	56.6	65.9	25.639	< 0.001
Heterog	Heterogeneity: Tau ² =0.000; Q=3.450, df=2 (p=0.835); I ² =0.000%								

DISCUSSION

One significant contribution of the present study, besides the systematization of the results presented in the literature, is the presentation of the intervals of normality organized according to the age of the assessed individuals in relation to each region of the spine. The lumbar curvature was the most frequently investigated, followed by the thoracic curvature. Few studies (Kasai *et al.*, 1996; Le Huec, Demezon & Aunoble, 2015; Yukawa, Kato, Suda, Yamagata & Ueta, 2012) have attempted to evaluate the cervical spine, therefore the results are inconsistent.

The literature reports that, on average, the normal thoracic curvature is 40° (Harrison, Harrison, Troyanovich & Harmon, 2000), while that of the lumbar curvature ranges from 45° to 50° (Damasceno, Catarin, Campos & Defino, 2006), with some variation according to the population investigated. However, in general, the studies show highly variable results, with wide confidence intervals (Table 3), compromising the classification process. It can be seen that most of the studies have standard deviations greater than

10°, which results in a high 95% CI, with around 40° of variability. The result of this meta-analysis for the thoracic and lumbar curvatures is associated with the most restricted confidence interval, around 10° of variability, in the different vertebral levels and populations (Tables 5-8). In fact, when a wide range of normality is used in the assessment, there is a greater chance of an individual, who in fact requires treatment, being considered normal. This situation is particularly undesirable in children and adolescents who would benefit from early diagnosis and treatment. On the other hand, a narrow range of normal values, might lead to the indication of unnecessary treatment. It should be noted that an adequate diagnosis and/or follow-up depends not only on the information regarding the angles of the spine, but also on the knowledge of the spine's morphology, kinesiology and biomechanics. Thus, we understand that, the narrow range identified in the present metaanalysis could help clinicians to better classify the individuals within a normal range and adequately monitor treatment effects.

The results of the meta-analysis show an increase in the average thoracic and lumbar curvatures which occurs with aging, reaching the highest averages in the elderly (44° in the thoracic curvature and 61° in the lumbar curvature). These findings corroborate those in the literature which show a tendency towards postural flexion in the elderly, with consequent increase in the magnitude of the sagittal curvatures in the vertebral column, due to the increased degeneration of the intervertebral disks (Benedetti, Berti, Presti, Frizziero & Giannini, 2008).

Regarding the cervical curvature, the result of the meta-analysis was inconsistent. In the elderly, the results for the cervical spine were not statistically significant, and the normality interval encompassed a range from cervical kyphosis, indicated by the negative sign (-1.7°), to cervical lordosis, indicated by the positive sign (25.4°). In adults, despite the results being significant, the range was rather more limited (1.9°-14.8°), while the signal was always positive. Even so, there is a great variability in the magnitude of the curvature.

Physiologically, the cervical spine has a lordotic curvature (Harrison, Janik, Troyanovich & Holland, 1996) and the normal alignment of cervical lordosis is one of the most important factors for the movement and function of this region (Miyazaki et al., 2008). Moreover, the loss of normal lordosis may induce pathological changes and accelerate the degeneration of the cervical spine (Miyazaki et al., 2008), as well as cause headaches (Nagasawa, Sakakibara &Takahashi, 1993) and neckaches (Harrison et al., 2004; Miyazaki et al., 2008). Therefore, given the morphological nature of the cervical lordotic curvature, the normality values indicated for adults and the elderly by the meta-analysis appear to inadequately represent this region of the spine. Hence, it is suggested that more studies be conducted with the aim of identifying a reference interval of normality indicated for the cervical spine in different age groups.

When analyzing our findings, some limitations should be taken into account. One of which is inherent to the adopted search method, which is based exclusively on electronic data bases and in the English, Spanish and Portuguese languages, which may have failed to identify possibly eligible studies. It is important to point out that, although the included studies meet the vast majority of criteria used to evaluate risk bias, criterion 3 (sample size) was not met by any of studies, and criterion 1 (random sample) was only met by two studies. Considering the intrinsic variability of spinal curvatures, the failure to meet these criteria may explain why the studies individually present a wide range of normality. Thus, by gathering these results systematically and compiling them according to the adopted methodologies, we were able to significantly reduce the range of normality.

It should noted be that important methodological differences exist between the included studies, for example: most of the studies involving evaluation of the thoracic region report shoulder positioning close to 90° of flexion. However, there are studies that refer to shoulders positioned at 60° (Vedantam, Lenke, Keeney & Bridwell, 1998), and between 30° and 45° (Cil et al., 2005; Jackson, Peterson, McManus & Hales, 1998; Korovessis, Stamatakis & Baikousis, 1998; Vialle et al., 2005), while there are even studies that fail to mention the position of shoulders (Boseker, Moe, Winter & Koop, 2000; Champain et al., 2006; Gelb, Lenke, Bridwell, Blanke, McEnery, 1995; Korovessis et al., 1998; Singer, Jones & Breidahl, 1990; Voutsinas & MacEwen, 1986). It is believed that this positioning, as well as the support or otherwise of the arms on some surface, may alter the angle of thoracic kyphosis, leading to variation in the results.

When referring to the assessments of the lumbar region, important differences are also found, with most studies conducting X-ray examinations with the patient in the upright position. However, there are reports of examinations conducted with the patients in the supine (Chernukha, Daffner & Reigel, 1998) and dorsal decubitus (Lin, Jou &Yu, 1992) positions, as well as studies that do not report the patient's position (Champain *et al.*, 2006). Therefore, when the objective of the examination is to measure the spinal curvature, placing the subject in the upright position is recommended in order

to minimize the variability inherent in the clinical examination.

CONCLUSION

Based on the present systematic review with meta-analysis, it was possible to identify a narrow interval of reference values for the thoracic and lumbar curvatures of the spine for healthy children, adolescents, adults and the elderly, assessed using the Cobb method with X-rays. For the cervical spine, the reference values were inconsistent due to the large variability between studies, which warrants further research.

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Conflict of interests:

Nothing to declare.

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