Compartment syndrome and regional anaesthesia: Critical review

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Acute compartment syndrome (ACS) is a potential orthopaedic/traumatology emergency. Without prompt, precise diagnosis and immediate treatment with surgical decompressive fasciotomy it can lead to neurological dysfunction and disability. The role of regional anaesthesia (RA) in patients at risk for ACS/ and in those with developed ACS is controversial. The aim of this critical review was to answer the question, whether regional anaesthesia can delay the diagnosis. The authors use an evidence-based approach to discuss these high risk patients in considering RA as a method of choice for effective analgesia.

To the date of data collection, there was no single case report identified where RA alone led to delay in ACS diagnosis and surgical treatment. In four clinical cases, epidural analgesia can be associated with delayed ACS diagnosis. Frequent clinical evaluation and breakthrough pain despite a functional RA in combination with intracompartment pressure measurement remains the keystone of recommended management for patients at risk of ACS.

Key words: compartment syndrome, regional anaesthesia, peripheral nerve block/blockade

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INTRODUCTION

Compartment syndrome (CS) is a clinical condition where increase in pressure within a closed compartment leads to perfusion abnormalities, ischemia and neurological damage. CS is considered a real orthopaedic/traumatology emergeency¹. There are several pathophysiologic theories for describing this syndrome including the arteriovenous gradient theory and the ischemia-reperfusion syndrome². However, whether there is a extraluminal or intraluminal pressure increase, the main principle remains the same - the pressure increase leads to capillary blood flow restriction, decrease in tissue pO, and consequently to tissue hypoxia and ischaemia². The pressure elevation in the compartment can arise from the inflammatory response, oedema, haemorrhage or ischaemia reperfusion syndrome, arterial pathology and also from external compression from casts, tight bandages, dressings or burns³⁻⁵. CS has been diagnosed in a variety of anatomical compartments including leg, gluteal region, thigh, tibia, foot, arm, forearm, hand and abdomen⁴. Fractures of the long bones, predominantly tibia (diaphysis) are the cause of approximately 75% of ACS and the risk is even higher in the case of comminuted fractures⁶⁻⁷. Soft tissue injuries without fracture represent the cause of around 20% of ACS. The reported incidence of ACS is 3.1 per 100 000, and ten times higher in men⁸. The incidence in the paediatric population is even lower although due to physiologic higher intracompartment pressure (13-16mmHg vs. 0-10 mmHg) children are at greater risk of ACS (ref.⁹). Identified risk factors for ACS are: male gender, young age <35 years, open fracture, intramedullary nailing, anticoagulation therapy, high-energy and penetrating trauma, vascular injuries, tourniquet use, haemophilia and complications of intravenous and intraosseous infusions^{8,10-19}.

Prompt diagnosis of ACS is crucial and significantly lowers the morbidity. Clinical signs of ACS are known as 5/6P: pain out of proportion to injury and pain with acute stretch, pallor, pressure, paraesthesia, paralysis and pulselesness²⁰⁻²¹. Pulselessness is not frequent, it is considered as a late sign²² and is associated with bad prognosis²²⁻²⁴. Clinical findings for ACS are however unreliable and can lead to delay in treatment^{5,25} because ACS can also present without any pain (in up to 10% of patients with ACS) (ref.^{3,26}). The most reliable method for diagnosis seems to be frequent assessment of a patient's clinical condition combined with intracompartment pressure monitoring in high risk patients. Intracompartment pressure monitoring (ICP) is considered as a "gold standard" for ACS diagnosis and for decompressive fasciotomy indication^{5,14,16,24,27}. The normal intracompartment pressure is in between 0-8 mmHg or below 10-12 mmHg according to different data^{9,28-30}. Pain is common with the ICP pressure over 20 mmHg and pressure over 30 mmHg is considered to be indication for emergency fasciotomy^{8,10,27,31,32}, but the precise value varies in the literature between 30-45 mmHg³³⁻³⁴.

Other authors recommend the Δ pressure - perfusion pressure (Δ pressure = diastolic blood pressure - intracompartmental pressure) with a threshold of 30mmHg³⁵. Δpressure has been reported as more sensitive than the absolute measured pressure and can lead to minimizing the risk of unnecessary fasciotomy³⁶⁻³⁷. Data are consistent for the urgent indication for fasciotomy in ACS and it appears that performing the fasciotomy within 6 hours after diagnosis (after onset of ACS) is associated with good outcome - 88% good outcome, 3.2% amputation rate compared to fasciotomy after 12 hours - good outcome only in 15% of patients with 14% amputation rate³⁸⁻³⁹. The fasciotomy timing is not based on the robust evidence-based data (EBM) and neurological deficit can be diagnosed in cases with rapidly performed fasciotomy (in 2 h). Hence, this seems to indicate fasciotomy as quickly as possible after confirmation of the ACS (ref.⁴⁰).

Decompression with surgical fasciotomy is considered a gold standard for therapy CS. The timing of fasciotomy is important and the best functional results without neurologic consequences are achieved when fasciotomy is performed within six hours from onset of CS (ref.²⁷). Regional anaesthesia (RA) is considered as a possible risk factor that can delay the diagnosis of acute CS (ACS) (ref.⁴¹⁻⁴³). Data are however derived only from case reports and case series and currently there is no uniform opinion in the literature or in clinical practice on the role of RA in patients in ACS or at in risk of ACS formation. Therefore, we decided to conduct the review of literature to answer whether the regional anaesthesia can lead to delay in diagnosis and treatment of ACS.

RESULTS

In the initial results we identified 294 citations, 88 were related to the topic: compartment syndrome and regional anaesthesia in the reviewed time period. After exclusion of duplicates (n=56), articles not in English (n=3) and articles with no available abstract or full-text online (n=7) a total of 22 full-text articles were included in the final analysis: case reports (n=15), review of literature (n=5), clinical trial (n=1), systematic review (n=1). Search diagram is showed in Figure 1.

No randomized controlled study dedicated to the topic was found. The currently available published data are of low-EBM (evidence-based medicine) quality (case reports, case series, retrospective data) with the high risk of bias and that is the main limitation of this review.

In selected 15 case reports were 20 published clinical cases – Table 1. The case reports were divided into 3 sections according to type of anaesthesia: regional anaesthesia, intravenous regional anaesthesia and epidural or subarachnoid/spinal anaesthesia and analgesia. No single case (0/8) in regional anaesthesia subgroup was found to be related to delay in acute compartment syndrome (ACS) diagnosis. In 7 cases, breakthrough pain appeared despite functional regional analgesia and in 1 case there was swelling, blisters and passive extension excessive pain.

In the intravenous regional anaesthesia subgroup only

two case reports were included in the review. In both, pain immediately (0-30 minutes after surgery) led to ACS diagnosis.

10 case reports were included in the epidural/spinal anaesthesia and analgesia subgroup. In 6 cases anaesthesia/analgesia did not lead to delay in ACS, because clinical signs of compartment syndrome were present despite functional blockade or the ACS appeared after a prolonged interval after epidural analgesia discontinuation. In 2 published cases Tang et al.⁴⁴ and Somayaji et al.⁴⁵ the epidural analgesia could have led to delay in ACS diagnosis however, in the first case there were some clinical signs of ACS - swollen calf, diminished capillary return. In the second case, dense block were persistent 24 h after epidural induction. In both cases the delayed diagnosis led to permanent neurological deficit. In Kumar et al. 46 case 3 and case 4, the pain appeared 4 h after epidural analgesia discontinuation. Epidural analgesia led to the delay. However there were clinical signs of ACS: erythema, tense and tender area and swelling in both patients. After urgent fasciotomy both patients fully recovered with no neurological deficit.

The only one included clinical trial (Maletis et al.⁴⁷), compared the frequency of ACS in patients undergoing closed reduction of closed lower leg shaft fractures under general or intravenous regional anaesthesia. General anaesthesia had favourable effect with only 13% vs. 27% rate of ACS diagnosis. The reason for the higher incidence may be, according to the authors, explained by the thigh tourniquet use.

In the recently published systematic review by Driscoll et al.⁴⁸, the authors compared regional anaesthesia and the patient-controlled analgesia (PCA) in patients with compartment syndrome. Although they used different search methods and a different approach for considering the effect of anaesthesia and analgesia on the delay in ACS diagnosis, in 6/8 (75%) patients in the PCA group ACS was diagnosed with a delay compared to the RA group where delay in diagnosis was detected in 19/29 (65%) case reports (Table 1). When comparing the possible influence of RA on delay in time (considering the evolution in regional anaesthesia – lower dosing, lower concentrations, ultrasound guidance), the more recently published data (after 2009) claimed that in 75% cases, no association between RA and delay in ACS was detected⁴⁸.

DISCUSSION

Our results showed no relationship between functional regional anaesthesia and delay in ACS diagnosis. Intravenous regional anaesthesia can be associated with risk of ACS development, because the prolong and excessive cuff pressure can lead to ACS formation. However, currently, intravenous regional anaesthesia is used only occasionally. Regional continuous analgesia is considered superior to opioid-based analgesia and can lead to reduction of in-hospital stay and chronic pain incidence ⁴⁹. Regional anaesthesia (RA) in patients with ACS or in patients at risk is controversial, however we found no

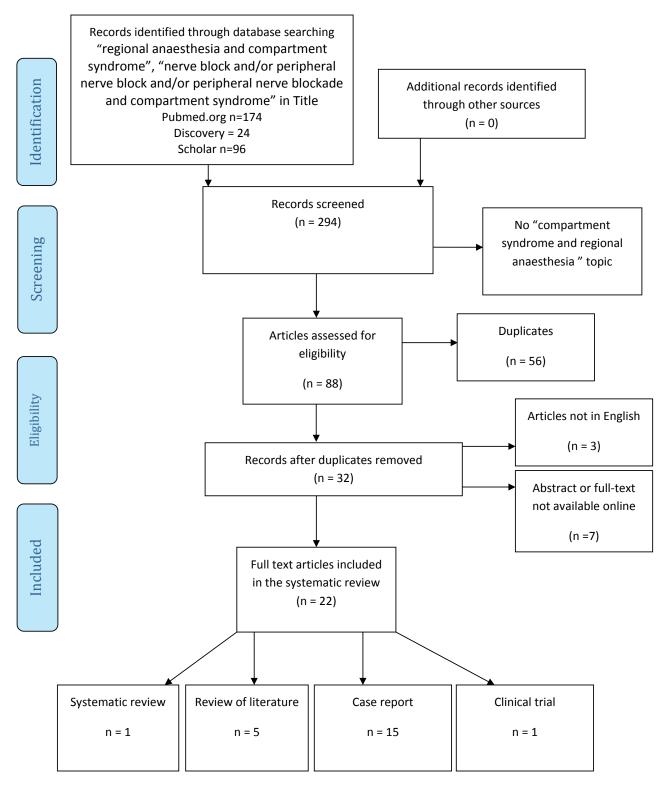


Fig. 1. Data search diagram.

published case report confirming that regional anaesthesia was the primary cause of the delay in ACS (ref.⁵⁰⁻⁵¹), Aguiree et al.³⁸ analysed 5 published case reports blaming RA for delay in ACS diagnosis, however in all cases the breakthrough pain was missed. From an ethical and pathophysiological point of view the patient's pain should be treated and there are several published case reports on opioid-based pain management has led to delay in ACS

diagnostics⁵⁰⁻⁵¹. For this reason, the analgesic method per se cannot be considered as the reason for delay in diagnostics^{50,52}. Cometa et al.⁵³ and Walker et al.⁵⁴ reported that despite functional continuous regional blockade, ischaemic pain was unaffected and patients can present with breakthrough pain.

It seems that compartment syndrome can be safely detected despite functional regional blockade⁵⁵ and pa-

Table 1. List of case reports.

Author	Clinical condition, type of surgery	Type of anaesthesia	Symptoms of Acute compartment syndrome (ACS)	Clinical manage- ment	Time of ACS diagnosis	Intracompartment Fasciotomy pressure values	Fasciotomy	Did regional anesthesia/analgesia mask the symptoms of ACS	Neurologic recovery	Authors comments	Ref.
Regional anaest	Regional anaesthesia / analgesia										
Uzel et al.	closed femoral fracture internal fixation	single-injection femoral block + general anesthesia	1st day after surgery – disproportionate pain	Intracompartment pressure monitor- ing, ACS diag- nosed, fasciotomy performed	1st postop- erative day	compartment pressure 54 mmHg	YES	ON	Full	3 h and 40 min after surgery breakthrough pain (despite functional regional blockade) - subcutaneous morphine	58
Cometa et al.	distal femur and proximal tibia osteotomy with external fixation	continuous femoral and sciatic nerve blocks + general anesthesia	2nd postoperative day, severe pain despite effective blocks and oral opioids	ACS diagnosed, fasciotomy per- foremd	2nd postop- erative day	over 30 mm Hg	YES	NO - breakthrough pain despite functional regional analgesia	Full	breakthrough pain despite functional regional analgesia	53
Kucera, et al. Case 1	right lateral ankle ligament reconstruction	single-injection femoral and sciatic nerve blocks	90 min after the cast dress- ing - severe pain despite persistent complete sensory and motor blockade	cast removal and reapplied	90 min after surgery	NOT measured	OZ	NO - breakthrough pain despite functional regional analgesia	Full	breakthrough pain despite functional regional analgesia	57
Aguirre et al.	complex distal humerus fracture	infraclavicular catheter + general anesthesia, patient-controlled regional analgesia postoperatively	severe forearm pain 14 hours after surgery despite functional regional analgesia	Intracompartment pressure monitor- ing, ACS diag- nosed, fasciotomy performed	14 h after surgery	40 mmHg	YES	NO - breakthrough pain despite functional regional analgesia	Full	breakthrough pain despite functional regional analgesia	38
Rauf et al.	open reduction and internal fixa- tion of the radius	supraclavicular bra- chial plexus block + general anesthesia	disproportionately intense pain despite an otherwise fully functioning sensory and motor block, swollen, tender, loss of radial pulse	revision surgery - haematoma evacuation	after surgery - time not specified	NOT measured	YES - revision surgery	NO – breaktrhrough pain despite functional regional blockade	Full	clinical signs of ACS despite functional regional blockade	59
L. Sermeus et al.	forearm osteochondroma resection	Infraclavicular block + general anesthesia	Breakthrough pain despite functional regional block. Motor function and sensitive deficit, swollen fingers	forearm cast removal - pain disappeared while motor function and sensation recovered.	In the night - time not specified	NOT measured	ON	NO - breakthrough pain despite functional block	Full	ACS diagnosed - loss of sensation, motor function deficit and break- through pain	4
Munk- Andersen et al.	open tibial shaft fracture	single-shot distal sciatic nerve preoperatively and postoperatively, lst postoperative day - distal sciatic nerve catheter was inserted	calf muscles were tense and sore, patient suddenly experienced severe pain in the lower leg	ACS diagnosed and fasciotomy performed	1st postop- erative day	NOT measured	YES	NO – break-through pain despite functional regional analgesia	Full	breakthrough pain despite functional regionaol analgesia	09

Table 1. Continued

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Neurologic Authors comments recovery	Clinical signs of ACS were present despite functional regional blockade		ACS diagnosed immediately after surgery, intravenous regional anaesthesia complication	ACS diagnosed 30 min after surgery, intravenous regional anaesthesia complication		clinical sign of ACS - diminished capillary return, authors describe full anesthesia below pelvis ith epidural analgesia on the 2nd day	dense block presistent 24 h after epidural induction need further clinical
Neurologic recovery	Permanent neuro- logical deficit		Permanent deficit	Full		Permanent neurologic deficit	Permanent deficit
Did regional anesthesia/analgesia mask the symptoms of ACS	NO – clinical signs of compartment – loss of sensation, no active movement, severe pain with passive movement		NO - pain immedi- ately after end of the operartion	NO - pain 30 minutes after end of the oper- artion		YES	YES
Fasciotomy	YES		Several fascioto- mies	YES		YES	YES
Intracompartment Fasciotomy pressure values	46 and 50 mm Hg in the anterior compartment and a reading of 22 mm Hg in the posterior compartment of the forearm were noted		NOT measured	NOT measured		The pressure of the anterior, peroneal, superficial, and deep posterior compartments was 65, 75, 75, and 80 mm Hg	NOT measured
Time of ACS diagnosis	24 h after surgery		immediately after surgery	30 min after surgery		2nd postoperative day	36 h postop- eratively
Clinical management	ACS of the forearm diagnosed and fasciotomy was performed		ACS diagnosed and fasciotomy was performed	ACS diagnosed and fasciotomy was performed		intracompartment pressure monitoring, ACS diagnosed, Fasciotomy performed	epidural infusion stop, ASC diagnosed, Fasciotomy
Symptoms of Acute compartment syndrome (ACS)	24 hours after surgery loss of sensation in the fingers; multiple swellings over the forearm and hand, oedematous, elbow with multiple haemorrhagic blisters, no active movements of the fingers and wrist, passive movements were painful and limited, passive extension of the fingers caused severe pain		increased forearm muscle tension, hand anaesthesia, pallor, and limited motor function developed	pain and swelling of the forearm 30 min after operation		swollen calf immediately after surgery, 2nd postoperative day – diminished capillary return	dense block, tachycardia and a temperature of 38.5 °C, severe pain after epidural analgesia stopping, disconfigat in both bushold.
Type of anaesthesia	axillary nerve block, the patient was discharged the same day, as he was pain free and comfortable		iv regional anesthesia with 360 mg lidocaine and sedation, tourniquet time was 107 min at a pressure of 260 mmHg	iv regional anesthesia, tourniquet time was 64 min at a pressure of 250 mmHg	ılgesia	epidural anaesthesia/ analgesia postopera- tively	epidural anethesia/ analgesia + general anesthesia
Clinical condition, type of surgery	internal fixation radius	Intravenous regional anesthesia	left-hand Dupuytren's contracture	Dupuytren fasciotomy	Epidural/spinal anesthesia and analgesia	total knee arthroplasty	hip arthroplasty
Author	Ganeshan et al.	Intravenous reg	Anantha- narayan et al. Case 1	Anantha- narayan et al. Case 2	Epidural/spinal	Tang et al.	Somayaji et al.

Table 1. Continued

Ref.	57	63	64	65	46	46
Authors comments	breakthrough pain despite functional regional analgesia	late ACS diagnosis not masked by epidural analgesia	ACS diagnosed in patient recieving parenteral opioids	breakthrough pain appeared despite functional epidural analgesia, the pain was treated with i.v. opioids	pain appeared 28 h after epidural infu- sion stopping	pain appeared 16 h after epidural infu- sion stopping
Neurologic recovery	amputa- tion of the 4th and 5th finger	NOT clear	Full recovery	Full recovery	Full recovery	Full recovery
Did regional anesthesia/analgesia mask the symptoms of ACS	NO - breakthrough pain despite functional regional analgesia	NO - pain appeared after epidural removal	NO – at the time of ACS diagnosis	NO - despite functional epidural analgesia the breakthrough pain appeared	NO - pain appeared 28 h after epidural infusion stopping	NO - pain appeared 16 h after epidural infusion stopping
t Fasciotomy	NO - vascular bypass surgery	revision	YES	YES – several hours after breakthrough	YES - 48 h after initial surgery (28 h after epidural infusion stopping)	YES - 44 h after the initial operation
Intracompartment Fasciotomy pressure values	NOT measures	NOT measured	NOT measured	YES (values not included in the text)	Not measured	Not measured
Time of ACS diagnosis	immediately after surgery	dull pain after epidural removal, per- sistent pain for 3 days after epidural removal, dysesthesia	2nd day – neurological deficit	7 h after surgery	48 h after initial sur- gery (28 h after epidural infusion stopping)	appeared 16 h after epidural infu- sion stopping
Clinical management	intravenous opioids insufficient - brachial to palmar arch bypass failure, amputation of the 4th and 5th finger	epidural removal	epidural infusion stopped, paren- teral analgesia	ACS diagnosed a fasciotomy performed	acute gluteal compartment diagnosed and fasciotomy was performed	acute gluteal compartment diagnosed and fasciotomy was performed
Symptoms of Acute compartment syndrome (ACS)	despite dense sensory and motor blockade, she still reported excruciating pain	swollen leg 4 days after surgery	hyposthesia and mobility limitations 1st postopera- tive day	7 h after surgery when she experienced ever increasing pain despite intravenous opioids	pain in the left buttock 28 h after discontinuation of the epidural infusion, tense, tender swelling no abnormal neurological findings.	severe right buttock pain 16 h after discontinuation of the epidural infusion, firm, tense, tender swelling with erythema
Type of anaesthesia	continuous cervical paravertebral block	epidural anesthesia / analgesia	subarachnoid block + epidural catheter insertion	caudal epidural injection + general anesthesia	epidural anaesthesia, the epidural infusion was discon- tinued after 20 h introduced at the L4/5 level	epidural anaesthesia at the L4/5 level, epidural infusion for 28 h
Clinical condition, type of surgery	necrotic lesions of 4th and 5th digits	tibial soft tissue defect – fibular transfer	total knee arthroplasty	6-year-old child, midshaft tibial fracture	total knee arthroplasty	total hip arthro- plasty
Author	Kucera, et al. Case 2	Streker et al.	Pinheiro et al.	O. Ross et al.	Kumar et al. Case 1	Kumar et al. Case 2

Table 1. Continued

Author	Clinical condition, type of surgery	Type of anaesthesia	Symptoms of Acute compartment syndrome (ACS)	Clinical manage- ment	Time of ACS diagnosis	Intracompartment pressure values	Fasciotomy	Clinical manage- Time of ACS Intracompartment Fasciotomy Did regional anesthe- Neurologic Authors comments ment diagnosis pressure values sia/analgesia mask the recovery symptoms of ACS	Neurologic recovery	1	Ref.
Kumar et al. Case 3	right hip arthroplasty	right hip arthro- epidural anaesthesia plasty at the L3/4, epidural infusion for 19 h	left buttock pain 4 h after discontinuation of the epidural infusion, erythematous, tense, and tender area over the left buttock and pain on passive flexion at the hip.	acute gluteal compartment diagnosed and fasciotomy was performed	4 h after epidural infu- sion stopping	not measured	YES - 28 h after the initial operation	YES - pain appeared Full 4 h after epidural infurecovery sion stopping	Full	other clinical signs of ACS were present - erythema, tense and tender area	46
Kumar et al. Case 4	total knee arthroplasty	epidural anaesthesia at the L1/2, epidural for 43 h	the L1/2, epidural anaesthesia at right foot drop - 4 h after the L1/2, epidural the epidural infusion stop- for 43 h	acute gluteal compartment	4 h after the not measured epidural infusion stopping		YES - timing is not clear	YES - pain appeared 4 h after epidural infu- sion stopping	Full	tourniquet was applied for 1 h and 50 mi, other clinical signs of ACS were present - swelling and tenderness over the right buttock	94

tients present with the classical signs of ACS (ref.²³). The French Society of Anaesthesia does not consider the risk of ACS as a contraindication for regional anesthesia⁵⁶ and because there is evidence that regional blockade can increase the blood flow through the partial sympathetic blockade without blocking the warning signs of ACS, the implementation of regional anaesthesia also in patients in the risk of ACS should be encouraged.

For minimizing the risk of delay ultrasound guidance for regional blockade and the catheter insertion, which can lead to lowering the dose of local anaesthetics are strongly recommended. Low concentrations of local anaesthetics can facilitate the motor function of the limb and allow breakthrough pain detection^{3,23,38,57}.

Several cases (Kumar et al. 46) reported, that epidural analgesia can lead to delay in ACS diagnosis, however the ACS is a complex syndrome and can present also without pain. In the majority of these cases there were other clinical signs of ACS presentation. These high-risk patients should be more frequently evaluated and intracompartment pressure measurement should be used. Frequent clinical re-evaluation of the patient's condition, increasing demands for analgesia and presence or reoccurrence of the breakthrough pain despite the functional RA should be incorporated in to management of the patient in the risk of ACS.

One interesting observation can be found in Driscoll et al. 48, that most of the articles pinpointing the delay in ACS diagnosis on RA were published in surgical/orthopaedic journals and most of the articles defending RA were published in anaesthesiology journals, which corresponds daily clinical practice.

Currently the EBM data considering the ideal analgesic method in patients with the risk of ACS is insufficient. Every single analgesic method can be associated with the delay in ACS diagnosis if pain is the sole diagnostic criteria. As opioids (PCA) can lead to even higher incidence of delay compared to regional anesthesia⁴⁸ and it is highly unethical not to treat the pain, the complex management of the whole patient at risk of ACS appears to be more important than the analgesic method alone.

Summary of evidence

We found no prospective RCT dedicated to this theme and the reviewed data can be considered as low-EBM quality (case reports, case series). Currently clinical practice is based on several reported cases and the theme is considered controversial.

Limitations

The main limitation is sparse data of low quality, search strategy limited by 3 databases (PubMed, Scholar. google, EBSCO – Discovery) and inclusion only the articles in the English language.

CONCLUSION

Results of the review didn't show, that RA can be associated with higher risk in delay ACS diagnosis compared

to other analgesic methods (intravenous opioid analgesia, epidural analgesia, PCA) and the cornerstone of diagnosis and treatment has to be the complex management including the intracompartment pressure monitoring in selected high-risk patient. Pain alone must not be considered as a single diagnostic sign of compartment syndrome.

However it must be noted that the evidence is spare and no clear guideline is currently available. There is an urgent need for discussion between orthopaedic/traumatology and anaesthesiology communities with the aim of establishing clear recommendation how to proceed in this clinical scenario.

Search strategy and selection criteria

This article followed the PRISMA Statement (www. prisma-statement.org) for meta-analysis and reviews.

We searched the following databases: Scholar Google (https://scholar.google.com), PubMed (http://www.ncbi. nlm.nih.gov/pubmed/), Discovery - EBSCO discovery service for keywords: regional, anaesthesia and compartment syndrome, nerve block or peripheral nerve block or peripheral nerve blockade and compartment syndrome located in the title or in the abstract.

We searched for data published between 1980 and 12/2016. All full-text articles in English related to the topic regional anaesthesia and compartment syndrome were included. Inclusion criteria were: articles found by selected search strategy (keywords, published between 1980-2016, selected databases), written in English language, full-text available and dedicated to the topic, compartment syndrome. Exclusion criteria were: abdominal compartment syndrome, articles not related to the topic CS and regional anaesthesia.

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