Supplementary Table 1: Specific data on PNE

Name - Scientific literature

Reference	Specific data
	Must answer the question: "What term refers to PNE?
Adenis et al (2020) [17}	Pain neuroscience education, pain neurophysiology education
Dolphens et al (2014) [62]	Pain neuroscience education, Therapeutic pain neuroscience education
Lane et al (2018) [19]	Pain neuroscience education
Malfliet et al (2017) [20]	Pain neuroscience education
Malfliet et al (2019) [22]	Pain neuroscience education
O'Keefe et al (2015) [69]	Pain education
Vier et al (2018) [70]	Pain education
Werner et al (2010) [57]	Intensive neurophysiology education
Wälti et al (2015) [71]	Neurophysiological education, education on the neurophysiology of pain
Agarwal et al (2020) [23]	Pain neuroscience education
Anandkumar et al (2018) [24]	Therapeutic pain neuroscience education, pain neuroscience education
Bodes Pardo et al (2018) [54]	Pain neurophysiology education
Clarke et al (2011) [5]	Pain neurophysiology education
Gallan Martin et al (2019) [53]	Pain neuroscience education
King et al (2018) [48]	Pain neurophysiology education
Louw et al (2012) [58]	Neuroscience education
Malfliet et al (2019) [21]	Pain neuroscience education
Mansouri et al (2018) [73]	Pain neurophysiology education
Moseley (2004) [46]	Pain physiology education
Moseley (2005) [44]	Pain physiology education
Moseley et al (2004) [47]	Neurophysiology education, pain neurophysiology education
Moseley (2003) [45]	Pain physiology education
Nijs et al (2017) [27]	Pain neuroscience education
Nijs et al (2014) [26]	Pain neuroscience education, therapeutic pain neuroscience education

Nijs et al (2011) [31]	Pain physiology education, 'pain (neuro)physiology education' or 'pain biology education'
Orhan et al (2019) [28]	Pain neuroscience education
Petersen et al (2019) [29]	Pain neuroscience education
Pires et al (2015) [53]	Pain neurophysiology education
Puentedura et Louw (2016) [30]	Pain neuroscience education
Rizzo et al (2018) [67]	Pain biology education, pain education
Rufa et al (2018) [31]	Pain neuroscience education
Ryan et al (2010) [68]	Pain biology education
Saracoglu et al (2020) [32]	Pain neuroscience education
Saracoglu et al (2020) [33]	Neuroscience pain education
Tegner et al (2018) [65]	Neurophysiological pain-education
Tellez-Garcia et al (2015) [49]	Neuroscience pain education, neuroscience education, neurophysiology education
Toomey et al (2020) [34]	Pain neuroscience education
Ünal et al (2020) [35]	Pain neuroscience education
Van Wilgem et Keizer, (2012) [59]	The sensitization model
Wood et Hendrick (2019) [60]	PNE has been named in different ways: therapeutic neuroscience education, or "explain pain". Pain Neuroscience Education is used for the purpose of the article
Moseley et Butler (2015) [2]	PNE has been named in different ways: therapeutic neuroscience education, pain biology education, pain neuroscience education. Explain pain is used for the purpose of the article
	Parpose of the million
Louw et al (2016) [36]	PNE has been named in different ways: Explain Pain, pain neuroscience education, pain biology education, therapeutic neuroscience education. Pain Neuroscience Education is used for the purpose of the article
Moseley (2003) [42]	Pain physiology education
Louw et al (2017) [37]	Pain neuroscience education, de-education
Diener et al (2016) [12]	Therapeutic neuroscience education
Wijma et al (2016) [55]	Pain neuroscience education
Louw et al (2017) [60]	Pain neuroscience education
Moseley (2002) [41]	Neurophysiology education
Louw et al (2017) [51]	Neuroplasticity explanation
Lotze et Moseley (2015) [40]	Explaining Pain
Blickenstaff et Pearson, (2016) [38]	Pain neuroscience education
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Barbari et al (2020) [14]	Pain science education
Ibrahim et al (2019) [56]	Patient education
Ibrahim et al (2018) [61]	Patient education
Nijs et al (2015) [39]	Pain neuroscience education
Galan-Martin et al (2020) [25]	Pain neuroscience education

Name - Clinician educational resources

Specific data	
Must answer the question: "What term refers to PNE?	
Pain neuroscience education	
Explain pain	
-	Must answer the question: "What term refers to PNE? Pain neuroscience education

Definition - Scientific literature

References	Specific data
	Must answer the question: "What is PNE?"
Adenis et al (2020) [17]	PNE is an educational strategy aimed at explaining to the patient their pain experience, prognosis, and treatment options from a neurobiological perspective.
Agarwal et al (2020) [23]	PNE is a new cognitive therapy for patients with chronic pain teaches the patient about the biology and physiology of their pain experience.
Galan Martin et al (2019) [18]	PNE is an educational health strategy that aims to provide up-to-date information from neuroscience regarding chronic pain.
King et al (2018) [48]	PNE is an educational and cognitive-behavioral intervention that involves explaining pain biology to the patient.
Lane et al (2018) [19]	PNE is an educational intervention used by physical therapists to help patients understand the biology, physiology, and psychosocial factors that influence their pain experience, and to reconcile them with the maladaptive thoughts and beliefs associated with chronic pain.
Louw et al (2012) [58]	PNE is a form of CBT which aim of increasing knowledge and understanding of pain, and thus decrease fear associated with musculoskeletal disorders.
Moseley (2004) [46]	PNE is an individualized educational session that provides information on nociception and pain.
Nijs et al (2014) [26]	PNE is about explaining central sensitization to the patient using evidence from modern neuroscience.
Nijs et al (2011) [31]	PNE is education on central sensitization and its role in chronic pain
Orhan et al (2019) [28]	PNE is an educational model that explains pain from a biological and psychological perspective to the patient.
Pires et al (2015) [53]	PNE is a cognitive-behavioral intervention that uses the explanation of the neurophysiology of pain to alter pain-related illness beliefs, disease perception, and coping strategies, in order to promote a return to normal activities, and reduce levels of pain and disability related to movement.
Puentedura et Louw (2016) [30]	PNE is teaching people about the neurobiology and neurophysiology of pain.
Ryan et al (2010) [68]	PNE is a new cognitive-behavioral intervention that attempts to reduce pain and disability by educating the patient about pain biology.

Saracoglu et al (2020) [32]	PNE is one of the many cognitive therapy methods often implemented alongside manual therapy and exercise.
Saracoglu et al (2020) [33]	PNE is a new cognitive education method to explain pain to patients with chronic low back pain.
Tegner et al (2018) [6]	PNE is a cognitive-behavioral intervention in the form of a verbal intervention that provides education in the neurophysiology of pain to change maladaptive
	disease beliefs, alter maladaptive pain cognitions, and re-conceptualize pain.
Tellez-Garcia et al (2015) [49]	PNE consists of educational sessions describing the neurobiology and neurophysiology of pain, as well as the treatment of pain in the nervous system.
Vier et al (2018) [70]	PNE is a biopsychosocial approach that demystify fear-avoidance beliefs about pain and movement, and explains how pain is not necessarily linked to tissue
	damage or degeneration, and how biological, psychological, and cognitive factors influence pain perception.
Wälti et al (2015) [71]	PNE is a cognitive approach.
Werner et al (2010) [57]	PNE is a cognitive education program that is theoretically based on contemporary knowledge of pain mechanisms.
Wood et Hendrick, (2018) [60]	PNE differs from traditional pain education by aiming to desensitize the neural system by focusing on neurophysiology, neurobiology, pain representation and
	pain meaning, instead of using an anatomical and traditional biomedical model.
Moseley et Butler, 2015) [2]	PNE is a range of educational interventions aimed at helping the patient to change their understanding of pain; its meaning, purpose, and biological processes
	that underpin it. It refers to the theoretical framework used for treatment and the approach itself, rather than a series of specific techniques or procedures. PNE
	focuses on helping the patient to gain a functional understanding of pain by explaining the key biological concepts behind it. PNE is different from other
	educational strategies due to its emphasis on conceptual change, and is not: 1) advice on how to move in pain, manage their pain, manage their activity, set goal
	or solve problems (CBT); 2) an explanation of Gate Control Theory; 3) a statement that the cause of the pain is linked to an irreducible sensitization of the
	central nervous system; 4) a rejection of biomedical models in favour of only focusing on psychosocial factors; or 5) a discrete intervention to be used in
	addition to treatment strategies based on a model of structural pathology.
Louw et al (2016) [36]	PNE is a strategy for explaining to people the biology and physiology of their pain experience, with the aim of reducing their pain and disability
Diener et al (2016) [12]	PNE is an educational biopsychosocial approach aimed at helping the patient understand their painful experience from a neurological and physiological
	perspective in order to produce a therapeutic outcome.
Wijma et al (2016) [55]	PNE is a patient-centered and individualized explanation of central sensitization, including biopsychosocial interactions.
Louw et al (2017) [37]	PNE is an educational strategy that focuses on explaining the neurobiological and physiological processes involved in pain, with the aim of altering beliefs and
	cognitions about their pain experience.
Lotze et Moseley (2015) [40]	PNE is a range of educational interventions that aim to change the person's understanding of what pain really is. PNE differs from other pain education strategie
	derived from CBT, which are limited to the application of scientific principles (e.g. pain does not equal injury, moving despite the pain, pain is inevitable but
	suffering is optional), while PNE covers the science itself. PNE is a management approach rather than a set of techniques, and is integrated with CBT.
Blickenstaff et Pearson (2016) [38]	PNE is a biopsychosocial, cognitive, and educational approach whose goal is to modify the patient's beliefs and thoughts about pain through the lens of
	physiology.
Louw et al (2016) [64]	PNE aims to teach the patient about pain from a biological and physiological point of view, increasingly adopting a biopsychosocial approach.
Barbari et al (2020) [14]	1) Educational strategies are the processes used by clinicians to encourage patients to make informed decisions and use reasoning related to their condition; 2)
	They differ from communication strategies, whose goal is to maximize the interaction and exchange of ideas or concepts between the clinician and the patient to
	enhance the therapeutic bond and adherence to treatment; 3) PNE differs from educational strategies that focus on a biomedical model, such as ergonomics,
	biomechanics, back schools, postural exercises, or traditional TM; 4) Rather, it is based on a biopsychosocial vision that is in line with recent advances in
	neuroscience and CLBP research.
Ibrahim et al (2019) [56]	PNE is a psychosocial or cognitive education that emphasizes self-management strategies.

Definition - Clinician educational resources

Reference	Specific data Must answer the question: "What is PNE?"
Louw et al (2018) [13]	 PNE is the explanation to the patient of his painful experience with the latest knowledge on pain, while de-emphasizing him from tissue problems; 2) Pain reconceptualization by PNE refers to cognitive restructuring, while the physical treatment refers to the behavioural part; 3) One point of view is that PNE is more than a treatment technique, but an approach that permeates all aspects of care.
Moseley et Butler (2017) [18]	1) In a broad sense: PNE can be defined as a way of thinking about pain; 2) In a more targeted sense: PNE is not a single technique but a range of conceptual change strategies based on pain science.

Objective(s) - Scientific literature

Reference	Specific data
	Must answer the question: "What is the goal of PNE?"
Adenis et al (2020) [17]	The main objective is to help the patient reconsider their pain, from a sign of tissue damage / deterioration, to a sign of the sensitivity of their protective systems, influenced by a combination of biological, psychological and social factors. The other goal is to increase their adherence to active rehabilitation strategies based on a biopsychosocial framework and to enhance their preceived benefits.
Agarwal et al (2020) [23]	The objective is to teach the patient the biology and physiology of their painful experience.
Anandkumar et al (2018) [24]	The objective is to re-conceptualize the pain, address concerns about their back, and provide reassurance.
Bodes Pardo et al (2018) [54]	The objective is to alter patients' beliefs, as a pre-requisite for behavioral change.
Galan Martin et al (2019) [18]	The objective is to change cognitions by altering erroneous beliefs and decreasing catastrophizing, kinesiophobia and avoidance.
Clarke et al (2011) [5]	The objective is to improve patients' understanding of their pain.
King et al (2018) [48]	The aim is to reduce maladaptive beliefs and behaviours in order to reduce pain and disability.
Lane et al (2018) [19]	The objectives are to: 1) help patients better understand the biology and psychosocial factors associated with their pain experience; 2) reconcile them with their inappropriate thoughts and beliefs; 3) increase intrinsic motivation through self-efficacy and therapeutic alliance.
Louw et al (2012) [58]	The objective is to increase knowledge and understanding of pain, thus reducing fear related to musculoskeletal disorders.
Malfliet et al (2019) [22]	The objective is to reduce the perceived threat of pain by increasing knowledge about pain and re-conceptualizing it.
Malfliet et al (2017) [20]	The objectives are to: 1) Reconceptualize pain; 2) Optimize the therapeutic alliance (cornerstone of the neuroscience-based approach).
Moseley (2004) [46]	The objective is to help the patient understand their pain by understanding the physiological mechanisms underlying it, rather than just having a cognitive- behavioral response to it.
Moseley (2005) [44]	The objective is to modify patients' understanding of their painful condition (reconceptualizing the problem).
Nijs et al (2017) [27]	The objectives are to: 1) Transfer knowledge of pain neuroscience to the patient, enabling them to better understand their own pain and thus better manage it; 2) Enhance beliefs about pain, decreasing pain threat, including painful flare-ups from exercise and daily physical activity; 3) Prepare for cognitive-targeted exercise therapy.

Nijs et al (2014) [26]	The objectives are to: 1) Change beliefs by re-conceptualizing pain; 2) Convince patients that pain is not a result of tissue damage; 3) Make sense of pain
	without imaging abnormalities; 4) Transfer this knowledge to patients to prepare them for exercise therapy that is time-contingent, rather than symptom-
	contingent.
Nijs et al (2011) [31]	The objective is to modify understanding of their painful condition and to re-conceptualize their pain.
Orhan et al (2019) [28]	The objectives are to: 1) enhance coping strategies in the face of pain by changing misguided pain beliefs; 2) shift their understanding of pain from a threatening
	signal to an alarm signal that protects the body, emphasizing the disproportionate nature between tissue damage and pain.
Petersen et al (2019) [29]	The objective is to address high levels of fear-avoidance beliefs.
Pires et al (2015) [53]	The objective is to modify illness-related beliefs about pain, perception of disease, and coping strategies, in order to introduce a return to normal movements and
	activities, and to reduce levels of pain and disability.
Puentedura et Louw (2016) [30]	The objectives are to: 1) Reconceptualize pain by teaching more about the neurobiological and neurophysiological processes involved in the painful experience
	rather than focusing only on tissue pathology; 2) Shift the focus away from tissue as the main source of the problem and emphasize a thorough understanding of
	the pain experience as a path to eventual recovery.
Rizzo et al (2018) [67]	The objective is to help the patient understand the biological processes involved in the experience of pain.
Rufa et al (2018) [31]	The objective is to alter the beliefs of patients with chronic pain. PNE aims to influence elements such as kinesiophobia and catastrophism in order to reduce
	pain, increase tolerance to movement and reduce disability.
Ryan et al (2010) [68]	The objectives are to: 1) Reduce pain and disability by explaining the biology of pain; 2) De-medicalize the painful condition; 3) Shift attitudes towards a
	biopsychosocial self-management approach; 4) Emphasize that pain is not equal to injury; 5) Decrease fear associated with activity through physical activity
	sessions trained in a graduated manner.
Saracoglu et al (2020) [32]	The objectives are to: 1) Change misperceptions and incorrect beliefs about pain; 2) Educate the patient that pain can occur even without an injury, and that pain
	can become overprotective.
Saracoglu et al (2020) [33]	The objective is to alter patients' attitudes and beliefs regarding pain.
Tegner et al (2018) [6]	The objective is to change maladaptive disease beliefs, alter maladaptive pain cognitions, and re-conceptualize pain
Tellez-Garcia et al (2015) [49]	The objective is to help patients increase their understanding of their pain and thereby decrease fear of movement or re-injury, by explaining to them the
	neurophysiological mechanisms underlying chronic pain.
Toomey et al (2020) [34]	The objectives are to: 1) Increase the patient's understanding of the neurobiology of pain, and specifically the weak correlation with tissue damage; 2) Provide
	the therapist with the opportunity to move the patient's focus away from a persistent tissue issue; 3) Explore the psychological factors and social influences
	affected by pain.
Ünal et al (2020) [35]	The objective is to enable the patient to understand the neurobiology and neurophysiology of their pain experience.
Van WIlgem et Keizer (2012) [59]	The objectives are to: 1) Reconceptualize the somatic beliefs of an anatomical defect linked to their pain; 2) Provide an opportunity to provide a physical
	explanation for chronic pain through changes in the nervous system.
Wälti et al (2015) [71]	The objectives are to: 1) Reduce patients' perception of pain and disability, reconsider protective behaviors and self-restraint resulting from fear of movement; 2)
	Help them regain a confident and positive outlook on their abilities and recognize the beneficial effects of activity.
Wood et Hendrick (2019) [7]	The objective is to re-conceptualize pain as a marker of body tissue protection rather than a marker of injury.
Moseley et Butler (2015) [2]	The objectives are to : 1) Help the patient change their understanding of pain: what it represents, its function, and the biological processes that it is based on; 2)
	Change a person's conception of pain from a marker of tissue damage (or disease) to a perceived need to protect body tissues; 3) The goal is for patients to

	understand their pain and to incorporate this new understanding into their beliefs, attitudes, behaviors, treatments, and lifestyle; 4) Change the understanding of
	pain from a structural biomedical paradigm to a truly biopsychosocial paradigm; 5) Acquire a functional pain literacy.
Louw et al (2017) [21]	The objectives are to: 1) Help people reduce their pain and disability; 2) Help people modify their behavior that contributes to pain and disability (through PNE
	and associated cognitive-behavioral therapies); 3) Help people understand (reconceptualize) that their pain is not a direct indication of the state of their tissues,
	without implying that the pain is "all in their head".
Moseley (2003) [42]	The objectives are to: 1) Reduce the perceived threat of pain; 2) Increase the patient's knowledge of human physiology, and re-conceptualize their pain.
Diener et al (2016) [12]	The objectives are to: 1) Help the patient gain an understanding of their pain experience from a neurological and physiological perspective; 2) Produce a
	therapeutic effect.
Wijma et al (2016) [55]	The objective is to help progress through the stages of behavior change (Prochaska)
Louw et al (2017) [37]	The objectives are to: 1) Change beliefs and cognitions about their painful experience; 2) Convince the patient that the presence of central sensitization implies
	that the brain produces more pain signals independently of what is happening in the tissues.
Moseley (2002) [41]	The objective is to address cognitive and behavioral factors by re-conceptualizing the problem and providing information.
Lotze et Moseley (2015) [40]	The objectives are: 1) to provide a biological justification for the treatment; 2) To shift the understanding of pain from a sign of tissue damage or pathology, to a
	sign of the perceived need to protect body tissues.
Blieckenstaff et Pearson, 2016) [32]	The objectives are to: 1) Change the patient's beliefs and thoughts about pain; 2) Provide experiences aimed at reducing perceived pain-related threat; 3)
	Generate positive expectations for improved pain and function; 4) Create a reasonable context to explain the pain and invite the person to explore movement.
Louw et al, (2017) [64]	The objectives are to: 1) Help patients gain an understanding of their pain from a biological and physiological perspective; 2) Facilitate a change in behavior and
	encourage movement despite the pain.
Ibrahim et al (2019) [61]	The objectives are to: 1) Provide non-threatening information to enable a better understanding of pain; 2) Change unhelpful beliefs about low back pain; 3)
	Encourage active coping strategies and secure pacing; 4) Integrate self-management strategies; 5) Promote positive attitudes and healthy behaviors.
Nijs et al (2015) [39]	The objectives are reconceptualization and deep learning.

Objective(s) - Clinician educational resources

References	Specific data
	Must answer the question: "What is the aim of PNE?"
Louw et al (2018) [13]	The objectives are to : 1) Help people understand and re-conceptualize their pain from a biological and physiological perspective based on current knowledge; 2)
	Processing maladaptive cognitions to modify the pain experience; 3) Helping people understand that pain is a normal human experience.
Moseley et Butler (2017) [18]	The objectives are to: 1) Re-conceptualize his pain deeply; 2) Identify and increase SIMs (Cognitive and Behavioral Factors of Safety). Target concepts targeted
	by the conceptual change: i) Understanding why and how it hurts is part of the treatment; ii) Pain receptors do not exist; iii) Pain depends on the perceived
	danger/safety balance; iv) Pain depends on the context; v) Pain involves multiple brain activity; vi) When the danger persists, the protection systems are
	sensitized; vii) Pain is one protection system among others; viii) We are bioplastics; ix) Learning about pain can help move forward; x) Active strategies
	promote recovery.
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Content - Scientific literature

References	Specific data
	Must answer the question: "What does PNE cover?"
Adenis et al (2020) [17]	Educational content covers: 1) Deconstructing erroneous patho-anatomical and patho-mechanical beliefs initially; 2) Second stage education in pain biology; 3)
	The persistence of pain is less related to healed tissues, but more related to the sensitization of protective systems influenced by a combination of
	biopsychosocial factors; 4) Treatment options include movement, physical activity, stress and emotional management, with an emphasis on dosage and
	progressivity (rather than biomechanics and the need for tissue protection).
Agarwal et al (2020) [23]	PNE utilizes metaphors, examples, and visuals to educate patients about the neurobiology and neurophysiology of their painful experience.
Anandkumar et al (2018) [24]	Educational content covers: 1) Explanation of the origin of pain and its treatment through the nervous system; 2) Difference between acute and chronic pain; 3)
	Factors that favor the persistence of pain; 4) Focus on the multidimensional link between exercise addiction and persistent low back pain; 4) Establishing
	concerns about back fragility. Examples: i) lack of association between pain and MRI abnormalities; ii) spontaneous regression of herniated discs (with images)
	iii) explanation of the effect of mistaken beliefs and catastrophism on pain with the neuromatrix model; iv) video of the 3 tissue healing phases; v) pictures of
	visual illusions to demonstrate the importance of the brain's inaccurate perception; vi) normal time for tissue healing; vii) spine model to show that the spinal
	cord is well protected; viii) permanent co-activation of the trunk muscles and repetitive exercises can increase the load on already sensitized spinal structures an
	contribute to the maintenance of pain; ix) relationship between poor body perception, thoughts and beliefs, and persistence of pain; etc.
	contribute to the mannenance of pain, ix) relationship between poor body perception, thoughts and benefits, and persistence of pain, etc.
Bodes Pardo et al (2018) [54]	Educational content is based on the Explain Pain book and Pain in Motion content.
Clarke et al (2011) [5]	PNE provides a detailed explanation of the underlying neurophysiology of chronic pain using pictures and metaphors, with a special emphasis on the brain and
	its role in thoughts and attitudes.
Galan Martin et al (2019) [18]	Educational content covers: 1) Epidemiology of chronic pain; 2) Pain as an alarm system; 3) Pain is not equal to damage; 4) Psychosocial and cultural aspects of
	pain (session 1); 5) Differences between acute and chronic pain; 6) Danger assessment system; 7) Amplification and inhibition systems; 8) Pain as a brain
	response (session 2); 9) Consequences of chronic pain and central sensitization; 10) Movement, motor control disorder, and kinesiophobia; 11) Fear-avoidance
	behaviors; 12) Pain catastrophizing; 13) Structural and functional disturbances that generate persistent pain (session 3); 14) Knowledge as a tool in the
	evaluative process of sensory stimuli; 15) The belief system and its epigenetic effects; 16) Reversibility of structural and functional changes; 17) Neuroplasticit
	mediated by cognitive and somatosensory stimuli, and physical exercise; 18) Benefits of group exercise (session 4); 19) Review of the contents covered in the
	first four sessions, and of the most relevant aspects of the PE group sessions.
Dolphens et al (2014) [63]	Educational content is focused on the physiology of the nervous system in general and pain in particular. Content and pictures are based on the 'Explain Pain'
	book."
King et al (2018) [48]	Educational content is based on the book 'Explain Pain'".
Louw et al (2012) [58]	Educational content includes pictorial and metaphorical explanations of: 1) the peripheral nervous system; 2) the central nervous system and neuromatrix; 3)
	other protective systems; 4) treatment options. Summary 1: An analogy of an ankle sprain is used to explain lumbago (injury, inflammation, sensitization of
	peripheral nerves, desensitization, scarring). Acute pain is contrasted with persistent pain where nerve sensitization persists despite tissue scarring. This
	phenomenon is explained by the presence of certain factors such as fear of pain, stress related to family, work, anxiety, and different explanations. The good
	news is that the more we understand this, the more the nerves become desensitized.
Malfliet et al (2017) [20]	PNE states that 100% of pain is generated by the brain, and that chronic pain is more likely caused by hypersensitivity of the central nervous system rather tha
	by local tissue damage.
Mansouri et al (2018) [73]	Educational content covers: the origin of pain (tissues and nociceptive pathways), the role of the brain (cerebral decoding based on the assessment of perceived
Mansouri et al (2018) [73]	Educational content covers: the origin of pain (tissues and nociceptive pathways), the role of the brain (cerebral decoding based on the assessment of perceived danger) and the factors that influence the perception of pain. Different classifications of pain are based on duration (acute or chronic) and mechanism
Mansouri et al (2018) [73]	Educational content covers: the origin of pain (tissues and nociceptive pathways), the role of the brain (cerebral decoding based on the assessment of perceived danger) and the factors that influence the perception of pain. Different classifications of pain are based on duration (acute or chronic) and mechanism (nociceptive or neuropathic). Other components include sensory-discriminative, affective-emotional, cognitive and behavioral aspects. The ascending gate

Moseley (2004) [46]	Educational content covers neurophysiology of nociception and pain, more specifically: 1) Neuron (receptor, axon, terminal); 2) Synapse (neurotransmitter, ior
	channels, postsynaptic membrane, action potential); 3) Descending pathways of facilitation and inhibition; 4) Peripheral sensitization; 5) Central sensitization
	(increase in synaptic efficiency and number of postsynaptic receptors).
Moseley (2005) [44]	PNE provides information on the physiology of the nervous system and the mechanisms of pain in particular, using pictures, examples and metaphors. This
	information can be understood by people who have not received an education. The comprehensive content is described in the "Explain Pain" book.
Moseley et al (2004) [47]	Educational content includes information specific to the nervous system, such as the functional importance of each component, neurotransmitters, postsynaptic
	receptors, and the concept of a "volume knob" of post-synaptic excitation and inhibition. It is content-based on the current knowledge in neurophysiology of
	pain based on the work of Wall and Melzack. However, there is no specific information on the lumbar spine or mention of emotional and behavioral patterns
	associated with chronic pain (e.g. catastrophism, kinesiophobia). There are three different sections: 1) Presentation of the nervous system in general with a focu
	on nociceptive pathways and pain; 2) Presentation of synapses with a focus on the 2nd neuron (medullary); 3) The adaptability of the nervous system (includes
	afferent and efferent pathways, physiological peripheral and central sensitization, receptor synthesis, axonal germination, neuronal response to activity and
	control of movement).
Moseley (2003) [45]	Educational content covers the physiology of pain and injury
Nijs et al (2014) [26]	The content of the sessions is based on the book Explain Pain and covers several themes: 1) the characteristics of acute versus chronic pain; 2) the function of
	acute pain; 3) how pain arises in the nervous system (nociceptors, ion channels, neurons, action potential, nociception, peripheral sensitization, synapse,
	intersynaptic space, inhibitory and facilitatory chemicals, spinal cord, descending inhibitory and facilitatory pathways, role of the brain, pain memory, pain
	perception), 4) how pain becomes chronic (plasticity of the nervous system, modulation, central sensitization, pain neuromatrix theory).
Nijs et al (2011) [31]	The educational content is based on the book Explain Pain, covering the physiology of the nervous system and that of pain in particular. The various themes
	include: 1) the characteristics of acute versus chronic pain; 2) the function of acute pain; 3) how pain arises in the nervous system (nociceptors, ion channels,
	nurses action potential postantian participated constitution suppose intersupportionances inhibitory and facilitatory chamicale animal acted descending
	neurons, action potential, nociception, peripheral sensitization, synapse, inter synaptic space, inhibitory and facilitatory chemicals, spinal cord, descending
	inhibitory and facilitatory pathways, role of the brain, pain memory, and pain perception); 4) how pain becomes chronic (plasticity of the nervous system,
	inhibitory and facilitatory pathways, role of the brain, pain memory, and pain perception); 4) how pain becomes chronic (plasticity of the nervous system,
O'Keeffe et al (2014) [69]	inhibitory and facilitatory pathways, role of the brain, pain memory, and pain perception); 4) how pain becomes chronic (plasticity of the nervous system, modulation, central sensitization, and pain neuromatrix theory); and 5) potential maintenance factors for central sensitization, such as emotions, stress,
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	nerves: signal transmission (including synapses and neurotransmitters), the ability to modify peripheral nerves; 3) central nervous system: the role of the brain in pain production, the influence of beliefs, emotions, and attention on pain production; 4) chronic pain: the disconnect between pain and tissue damage, its
	unsuitable nature; 5) treatment strategies: the benefits of knowledge, stress relief, sleep hygiene, exercise, and gradual activity re-entry.
Ryan et al (2010) [68]	Educational content includes information on the biology of pain
Saracoglu et al (2020) [32]	Educational content covers: 1) Pain physiology; 2) Nociplastic pain; 3) The virtual body in the brain; 4) Changes in bodily perception related to pain; 5) The psychosocial dimensions of pain. PNE uses neurophysiological information to make the patient understand that pain can occur even in the absence of injury and that pain can become overprotective. Session 1: Peripheral neuropathic pain, peripheral nerve sensitization, allodynia, central sensitization, hyperalgesia; Session 2: Neuroplasticity, diffuse pain, central sensitization, hyperalgesia, allodynia; Session 3: Biology of stress, immune response, emotional overload, fear, catastrophizing and pain; Session 4: How to deal with pain? The role of exercise and manual therapy.
Saracoglu et al (2020) [33]	Educational content includes (with metaphors, images, graphics and anecdotes) teaching about the physiology of pain, central sensitization, body representation in the brain, changes in body representation due to pain, psychosocial dimensions of pain, fear of pain, and how these are affected by treatment. During the PNE sessions, it was explained that the nervous system can be overprotective and that in addition to the sensitivity of the central nervous system, nociceptive transmission can be affected by an individual's thoughts, beliefs, and environment. Session 1: Definition of the following terms: peripheral neuropathic pain, peripheral nerve sensitization, central sensitization, allodynia, hyperalgesia and neuroplasticity; Session 2: The role of central sensitization, hyperalgesia, allodynia on their symptoms. Why their pain expands, the role of neuroplasticity in chronic pain; Session 3: Biology of stress, immune response, emotional overload, fear, catastrophizing and their role in pain; Session 4: How to deal with pain? The potential effect of manual therapy and pain education in neuroscience.
Tegner et al (2018) [65]	PNE is explaining to the patient the key biological concepts underlying pain
Tellez-Garcia et al (2015) [49]	Educational content covers the neurophysiology of pain, without reference to the spine, discussing the distinction between acute and chronic pain. With additional information on the impact of beliefs and attitudes on pain.
Toomey et al (2020) [34]	Educational content includes: 1) Reassuring the patient that their pain is real even though the tissues may not be at risk; 2) Explaining the neurobiological and psychological mechanisms of manual therapy and the low biological plausibility of certain concepts such as "correcting subluxations"; 3) Understanding the biological processes that underlie pain; 4) Understanding that pain can become overprotective; 5) Understanding how psychosocial factors can positively and negatively modulate pain; 6) Using a diary to reflect the factors of good and bad days; 7) Identifying specific movements that the patient is apprehensive about, and breaking them down into small, manageable chunks; 8) Using breathing and mindfulness techniques. Resources: a) YouTube video (Tame the Beast, Why Things Hurt, Understanding Pain in Less than 5 Minutes); b) Books (Painful Yarns, Explain Pain); c) Analogies (Pain as an Alarm System; The Role of Expectations in Treatment; How Manual Therapy Can Provide Relief in Realigning Bones).
Unal et al (2020) [35]	Educational content covers pain mechanisms; central pain treatment; how the nervous system is sensitized in chronic pain; factors that cause chronic pain; side effects related to fear avoidance. But also: neurophysiology of pain, nociception and nociceptive pathways; neurons, synapses, action potential, spinal inhibition and facilitation; sensitization device; awareness center; plasticity of the nervous system. All explained with simple images, stories, leaflets, metaphors, drawings.
Van Wilgem et Keizer (2012) [59]	Anti-theft alarm metaphor to explain central awareness: "It's like having an anti-theft alarm system in the house that is always turned on." Metaphor of the thermostat to explain continuous chronic pain (e.g. fibromyalgia, headache): "It's like the thermostat is set too low, which causes the house to constantly overheat."
Vier et al, (2018) [70]	Educational content covers: 1) the importance of pain in our lives; 2) how pain is a useful warning sign that something may not be right; 3) what factors contribute to the chronicization of pain; 4) the significance of psychosocial factors such as stress, anxiety, and kinesiophobia in the chronicization of pain; 5) the role of insufficient sleep and physical activity in back pain; 6) the benefits of gradual exposure and daily movements; 7) the neurophysiology of pain; 8) the nocebo effect; 9) the lack of correlation between posture, imaging, and pain. In the other sessions: reinforcing the concepts mentioned above.
Wälti et al (2015) [71]	Educational content is based on a Randomized Controlled Trial (RCT) by Moseley et al. It provides an overview of the biopsychosocial model of chronic low back pain, including cortical dysfunction of pain and bodily perception. The focus is on how this model can explain the participants' characteristics of low back pain experiences.

Werner et al (2010) [57]	The content consists of an understanding of pain that differs somewhat from the traditional injury model. The content is based primarily on the neurophysiology of pain, reflecting sensitization and neuronal response to inactivity and control of movement. Based on this, the program has three basic elements: 1) reducing the perceived threat; 2) targeting the patient's understanding of pain; 3) exposure to threatening inputs.
Wood et Hendrick (2019) [60]	Educational content focuses on neurophysiology, neurobiology, representation and meaning of pain, instead of relying on the traditional anatomical and biomedical model (where is the lesion and how to fix it).
Moseley et Butler (2015) [2]	PNE aims to explain key biological concepts related to pain. These concepts are: 1) the variable relationship between nociception and pain; 2) the concept's powerful influence on pain; 3) upregulation of the nociceptive system (sensitization) when pain persists; 4) the coexistence of several protection systems (including pain); 5) the potential influence of other protective systems on pain; 6) adaptability, and therefore the ability to re-train our biology (including but not limited to the concept of neuroplasticity); 7) and the knowledge that this adaptation is likely to be slow. PNE may contain: 1) explanation of the biological processes underlying the pain; 2) PNE relies on multimedia design and principles to present the biology of pain; 3) the teaching that pain can become overprotective; 4) teaching that nociceptive messages can be up-regulated in the spinal cord; 5) teaching that nociceptive messages can be down-regulated by the brain; 6) the teaching that the nociceptive system can become hypersensitive leading to more nociceptive messages, but it is still the brain that decides whether or not to produce pain; 7) it is reassuring that the pain is still real even if the tissues may not be in danger.
Louw et al (2016) [36]	Educational content covers: 1) De-education: sharing normative data on imaging exams (e.g. Brinkji chart 2015) and helping them understand that abnormalities may not be associated with pain (and reduce catastrophizing and kinesiophobia); 2) Pictorial and simplified description of the nervous system; 3) Comparison of the nervous system to a highway network; 4) Comparison of the nervous system to an alarm system; 5) Analogy of the foot stepping on a nail to explain tissue injury, peripheral sensitization and desensitization, scarring; 6) Explanation of chronic pain: nerve tenderness persists while tissues have healed (graphs); 7) Proposal of factors (yellow flags) to explain why for some people, the sensitivity persists (images); 8) Impact of nervous system sensitization on function and morale (graph); 9) Proposal of a treatment plan to desensitize the nerves (including PNE and other conservative therapies).
Moseley (2003) [42]	PNE differs from other conventional education strategies (which focus on the anatomy and physiology of the spine), and includes detailed information on the neurophysiology of pain: 1) the neuron (specific receptors, nerve endings, axon) ; 2) the action potential (all or nothing law, postsynaptic membrane potential, propagation, dromic and antidromic flow); 3) the synapse (neurotransmitters, inhibitory and facilitating inputs, chemically controlled ion channels, ion channel synthesis and uptake); 4) primary nociceptors (danger receptors); 5) 2nd order nociceptors (sum of inhibitory and facilitating inputs, inhibitory interneurons, project to several parts of the brain); 6) cerebral production depends on the total perception of danger, all information relevant to this decision-making is taken into account: thoughts, memories, beliefs, explanatory models, consequences); 7) descending inhibitory and facilitatory pathways (dependent on the total perception of danger); 8) state-dependent functioning of the primary nociceptor: potentiation and summation, ectopic discharge, dorsal root ganglion, neurogenic inflammation, allodynia and hyperalgesia; 9) state-dependent functioning of 2nd order nociceptors: potentiation, blocking of active ion channels, increase in receptor synthesis, germination, activation by endocrine mediators
Diener et al (2016) [12]	Educational content covers: 1) Dispelling misconceptions about tissue health; 2) Explaining pain from neurobiology and physiology.
Louw et al (2017) [60]	PNE includes: 1) An alarm system metaphor for understanding central sensitization. The nervous system is compared to an alarm system to contrast sensitivity before and after the onset of pain. In some people, the alarm system does not settle down, leaving them with an oversensitive alarm system; 2) Manual therapy is presented as a transitional technique to recover range of motion and facilitate the activation of endogenous opioids.
Moseley (2002) [41]	Educational content is focused on the neurophysiology of pain without reference to the lumbar spine.
Louw et al (2017) [51]	Educational content includes: 1) A simplified, literal explanation of Penfield's homunculus as a virtual map of the body; 2) When we move less in a region, the map of that region blurs; 3) When we move a blurred region, we experience more pain; 4) It is possible to redraw the maps of the brain; 5) Manual therapy can help with this process.
Barbari et al (2020) [14]	Educational content includes: 1) Explanation of concepts and presentation of the neurophysiology of pain; 2) Explaining the nervous system, synapses and neuroplasticity based on the "Textbook of Pain" by Melzack and Wall, with diagrams and examples; 3) Focusing on topics such as the origin of acute pain in the nervous system, the transition from acute to chronic pain, central sensitization, the role of the brain in the perception of pain, the role of psychosocial factors related to pain, pain-related cognitive and behavioral responses, pain flare-ups and pacing through the use of metaphors and images; 4) Biology of pain using diagrams and freehand drawings; 5) Neurophysiology of acute and chronic pain, as well as the role of beliefs in relation to pain based on the book Explain Pain; 6) Content focused on vicious circles of pain, psychosocial factors involved in pain, awareness and the difference between pain and injury; 7) Neurophysiology of pain, cortical dysfunction, and body perception.

Nijs et al (2015) [39]	Pain neuroscience content rather than psychology content. Introduction to time contingent exercise.
	back pain red flags.
	cause of pain; 6) the basis of pain physiology; 7) staying active and resuming normal activities; 8) coping and pacing strategies; 9) modifying lifestyle; 10) Low
brahim et al (2019) [51]	Educational convent covers: 1) the meaning of low back pain; 2) facts about low back pain; 3) beliefs about low back pain; 4) the strength of the spine; 5) the
	Coping and pacing strategies; b) Self-management; c) Postural hygiene; d) Increasing activity levels; e) Modifying lifestyle; f) Warning signs and what to do.
	ligament, and joint strain, and stopping movement can cause these structures to become weak. To keep the back healthy, it needs to be active; ix) Other topics: a)
	presence of red flags; vii) Pain can be felt without any physical changes in the body; viii) Return to normal activities: Most pain in the spine is due to muscle,
	of the structures of the spine and difficulty of injuring it; 6) Cause of pain: poor radio-clinical correlation, poor prognostic predictor, use of imaging in the
	back pain (pain equals injury, physical activity, rest, etc.). Fear-avoidance beliefs and behaviors can promote the persistence of pain; 5) Basic anatomy: solidity
	associated costs; 3) Low back pain facts: common, not serious, recurrence is common but not serious; 4) Beliefs about low back pain: some beliefs about low
	back pain; 2) Meaning of low back pain: definitions of low back pain (nonspecific vs specific, acute vs chronic, epidemiology, explosion of disability and
	lingering pain, then starting to move on. Importance of setting realistic goals or action plans. Participants were allowed to tell their story/experience about low
Ibrahim et al (2018) [56]	Educational content covers: 1) Interactive session: Meaning of low back pain: Justification of treatment and expected objectives. Accepting that you have

Content - Clinician educational resources

Specific data
Must answer the question: "What does PNE cover?"
Educational content addresses the neurophysiological mechanisms of pain, and signs/symptoms, factors, and treatment options in a neurophysiological manner.
1) Neurophysiological mechanisms of pain (ion channels, nociception, peripheral sensitization, central sensitization, spinal inhibition & facilitation, pain is a
production of the brain according to the perceived threat, the neuromatrix of pain, the homunculus biology of stress , sympathetic, endocrine and immune
responses, inflammatory responses, impaired brain function, cortical blurring, positive neuroplasticity): 2) signs and symptoms (hyperalgesia, allodynia, diffuse
pain, sensitivity to cold and stress, loss of function, swelling problems, physical fatigue and sensitive muscles, intestinal problems, neuropathic pain); 3)
associated factors (stress, anxiety and fear, emotion, catastrophizing, sleep problem, past treatments, etc.); 4) treatment options (movement, exercise, pacing and
gradual exposure, membrane stabilizer). According to old Mosley papers, no mention of 1) anatomical and patho-anatomical aspects; 2) emotional and 3)
behavioral. However in this book, the authors propose that 1) anatomical and patho-anatomical aspects be addressed in PNE with the aim of "de-educating
patients" on their erroneous patho-anatomical beliefs; 2) They also propose that PNE address emotional aspects with the aim of helping patients understand why
they are suffering (how emotions can trigger, increase or biologically maintain pain); 3) finally they propose that the behavioral aspects be approached such as
"moving despite the pain"
Educational content includes short and long stories, metaphors, diagrams, analogies, literal explanations to explain biological concepts in order to aim for target
concepts: 1) broad on pain, neuroscience and anatomy (ex: solidity of the spine, pain is a protector, pain is produced by the brain, pain neuromatrix, pain
threshold vs tolerance threshold, Penfield homunculus, biologization of beliefs and thoughts, etc.); 2) tissue changes and nociception (no pain receptor, rarely
related pain and injury, tissue scarring, release of acidity related to prolonged positions, peripheral sensitization, etc.); 3) peripheral neuropathic pain (night
pains, nerves don't get stuck, ganglia, juice shots aren't injuries, importance of keeping nerves well oiled, etc.); 4) central sensitization (increase in nociceptive
volume, sensitivity to light, mirror pain, diffuse pain, etc.); 5) homeostatic systems (stress and homeostatic systems, stress and swelling, virus and pain, etc.; 6)
treatment (distraction is analgesic, hugs are analgesic, knowledge is analgesic, music is analgesic, movement lubricates joints, movement is medicine, pain can
improve, recovery is done fluctuations, sensitive but not hurt, flares up: avoid them but don't be afraid of them, neuro slips, movement is medicine, gradual
exposure, advice on belts & neck braces, manipulations, etc.; 7) diagnostic labels and radiological reports (a vertebra does not move, osteoarthritis reflects aging
and not pain, osteophytes are stabilizers, fibromyalgia reflects an overprotective alarm system, etc.). PNE stands out in terms of the content of expectation
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Theoretical framework and effects - Scientific literature

References	Specific data
	Must answer the question: "On which theoretical foundations is PNE based?" "How efficacy of PNE is explained?"
Adenis et al (2020) [17]	PNE increases the patient's adherence to an active rehabilitation program based on a biopsychosocial framework in order to desensitize their protective systems
Agarwal et al (2020) [23]	PNE explains its effectiveness by targeting catastrophism and thus reactivating endogenous inhibitory mechanisms.
Bodes Pardo et al (2018) [54]	PNE desensitizes the nervous system, especially when combined with therapeutic exercise. PNE targets the cognitive-behavioral components of pain, reduces top-down facilitation mechanisms, and encourages the activation of inhibitory mechanisms. A better understanding leads to more appropriate coping strategies, and enhances the effects of therapeutic exercises.
Galan Martin et al (2019) [18]	PNE could change cognitions, catastrophizing, kinesiophobia and avoidance by altering incorrect beliefs. This would consequently enhance adherence to physical activity, which encourages neurogenesis, results in neuroplastic changes in the brainstem and activates descending pathways that inhibit pain. These changes would together enhance quality of life, reduce pain and disability.
Clarke et al (2011) [5]	The underlying theory is that psychosocial factors can be strong predictors of persistent pain and disability (PNE target psychosocial factors)
Dolphens et al (2014) [63]	PNE is part of a neuroscience-based pain treatment that targets central factors (deficit in the descending inhibitory system, central sensitization), psychosocial factors (erroneous beliefs, catastrophizing) and peripheral factors (motor control dysfunction) within a larger biopsychosocial context. PNE specifically targets the enhancement of descending inhibition.
King et al (2018) [48]	In theory, the mechanism of action of PNE is through helping patients better understand their pain, correcting inappropriate beliefs, and reconceptualizing pain. Reconceptualizing pain has the potential to reduce perceived threat. Reducing this perceived threat could potentially: 1) reduce pain-related fear, pain-related distress and disability; 2) break the cycle of misguided problem solving; 3) and potentially reduce pain levels.
Lane et al (2018) [19]	The main mechanism would be pain reconceptualization, including concepts related to fear, knowledge and beliefs about pain. The pain reconceptualization would be a strategy through which therapists could promote self-efficacy, build the therapeutic alliance, and increase intrinsic motivation, thus promoting behavioral changes.
Louw et al (2012) [58]	The proposed mechanism of action is the pain reconceptualization in a way that increases patient confidence and activity levels. This is justified by the fact that studies have shown that maladaptive beliefs are associated with impairments in motor control.
Malfliet et al (2017) [20]	Chronic low back pain is a mixed condition combining peripheral mechanisms (e.g. impaired motor control) and central mechanisms (e.g. central sensitization, psychosocial factors). Therefore, treatment should target peripheral and central mechanisms within a larger biopsychosocial framework. This can be done through the use of PNE followed by an exercise program targeting cognitions
Moseley (2004) [46]	The underlying theory is that reconceptualizing the problem can improve maladaptive beliefs and attitudes. Educational intervention could reduce the levels of catastrophizing, leading to a decrease in vigilance to peripheral nociceptive impulses and a modification of the threshold of sensitivity or tolerance to pain, and thus of the performance on the Laseig test. This decrease in catastrophism could be explained by a modification of the meaning attributed to the nociceptive impulse, which would consequently modify the painful response.
Moseley (2005) [44]	One possible explanation is that the reconceptualization of the problem leads to a change in the belief that pain does not necessarily equate to injury, which in turn leads to increased confidence and increased levels of physical activity. It is highly likely that the change in fMRI during task performance is related to a decrease in perceived threat.
Moseley et al (2004) [47]	The explanatory hypothesis would be the reconceptualization of pain from a reliable indicator of vertebral pathology to the evaluation and adaptation of neural processes. Seen in this way, patients would be more likely to adhere to strategies of exposure to movements and activities that they are afraid of, and would more easily be able to challenge their catastrophic thoughts. PNE would become more clinically relevant when combined with other therapeutic strategies, thus increasing their effects. Another explanatory hypothesis would be that the reconceptualization of pain would reduce catastrophizing, which would have an influence on pain thresholds through a decrease in vigilance towards somatic signals. The authors explain that an education based on a structural model, implying that the underlying structures are vulnerable, would likely increase alertness levels and vigilance towards pain.

Moseley (2003) [45]	One of the hypotheses explaining the effect of PNE would be via the modification of beliefs and attitudes. Another hypothesis would be that modifying the
	meaning of pain (reconceptualization) would directly modify the intensity of pain, which is biologically plausible. Moreover, patients would be more likely
	resume their activities due to feeling less disabled.
Nijs et al (2017) [27]	PNE prepares for cognitive exercise targeted at systemically desensitizing the nervous system.
Nijs et al (2014) [26]	Central sensitization implies that the brain generates more warning signals, including pain, even when there is no tissue damage. Time-contingent treatment
	which is favored by PNE, could deactivate downward nociceptive facilitation. This viewpoint is supported by discoveries of decreased central nervous systemetry of the statemetry of the stateme
	hyperactivity and increased prefrontal cortical volume in response to time-dependent treatment in patients with chronic pain.
Nijs et al (2011) [31]	When only cognitive and behavioral responses are encouraged, without reconceptualizing pain, these responses can become counter-intuitive because pain
	always seen as a sign of injury for them (e.g. education in fear-avoidance model and gradual exposure without reconceptualizing pain). The hypothesis is
	therefore that by reconceptualizing pain, more appropriate cognitive and behavioral responses will follow as pain will be seen as less dangerous.
Wälti et al (2015) [71]	Various research groups have demonstrated the importance of focusing on the abnormal cortical processing of the central nervous system in patients suffering
	from chronic low back pain, including cognitive, sensory, and motor disorders. A cognitive approach (PNE) could be beneficial as part of a multimodal
	treatment.
Werner et al (2010) [57]	PNE is part of the evolution of the vision of low back pain from an injury model to a biopsychosocial model. This view incorporates peripheral and centra
	contributors, and acknowledges that the perception of their problem can influence their thoughts, emotions, and behaviors, which in turn can influence pain,
	may determine how to cope with current pain.
Wood et Hendrick (2019) [60]	The purpose of PNE is to desensitize the nervous system
Moseley et Butler (2015) [2]	PNE draws its key principles from psycho-education (particularly conceptual change strategies), health psychology and neuro-immune science related to pa
	PNE is based on theories of conceptual change: the questioning of existing knowledge rather than simply learning new information about potentially confron
	concepts. Considering a Bayesian perspective, pain can be viewed as a brain output that reflects the most advantageous response based on our brain's
	probabilistic evaluation. The hypothesis is that the brain is often overprotective due to the perception of an excessive threat to the body, which can be modula
	by any credible evidence of danger from nociceptive, somato-sensory, visual, cognitive, auditory, and social modalities. The hypothesis is that PNE alters the
	perception of threat by altering sensory modalities, such that the brain's response to the situational assessment shifts from "the situation requires protection
	(pain) to "does not require protection" (therefore no pain). Numerous clinical studies support that the modification of the perception of the threat linked to
	stimulus modifies the symptomatic response (Placebo, Nocebo). Numerous clinical studies support that the modification of the perception of the threat linker
	a stimulus, modifies the symptomatic response (placebo, nocebo). These effects have been investigated using fMRI, and several regions of the cerebral cort
	(anterior insular cortex and its connections with the periaqueductal gray matter, for example) seem to be involved in this process. Other studies seem to sugg
	that PNE would stimulate endogenous inhibition compared to control groups.
Moseley (2003) [42]	PNE works by reducing the perceived threat of non-nociceptive signals by enhancing a person's understanding of human physiology. Therapies that focus
	nociceptive cues should be implemented in an environment that does not reinforce threatening non-nociceptive cues. Current evidence shows that education
	limited effectiveness. The hypothesis is that the content of educational programs is counterintuitive for patients who have a structural biomedical understand
	of their pain (pain=injury). PNE addresses this limitation by helping the patient to reconceptualize their pain from a physiological perspective, assuming the
	appropriate cognitive and behavioral responses will follow. This theory is founded on the principles of deep learning (as opposed to surface learning) which
	suggests that information that is retained and understood can be applied to current problems. This deep learning is promoted when the participant is motivat
	and the information is made personally relevant, two factors that are favored by PNE. Thus, the goal of PNE is to reduce the perceived threat and therefore
	reduce the activity of the pain neuromatrix, as well as reduce its synaptic effectiveness, before introducing a therapy that: 1) targets the components of the
	neuromatrix pain without fully activating it; 2) increases physical and functional tolerance to threatening sensory and non-sensory stimuli.
Diener et al (2016) [12]	One of the possible explanations for the effectiveness of PNE would be related to the positive manipulation of beliefs and expectations (contextual effects) a
	their biological impact on the central nervous system.
Wijma et al (2016) [55]	PNE targets cognitions and perceptions that directly contribute to central sensitization, which in turn influence the patient's emotional and behavioural factor
	The patient's behaviours are influenced by their perceptions (as explained by the Common Sense Model). The link between these behaviours (avoidance and

Louw et al (2017) [60]	The nervous system and its plasticity are considered to be a therapeutic target for PNE and other treatments. PNE has been developed to target central sensitization. This hypothesis is supported by clinical trials showing an improvement in local and distant pressure thresholds after PNE. In this context, PNE offers a therapeutic space (decreased nervous sensitivity) for the introduction of a progressive mobilization approach. Another effect of PNE could be related to its ability to foster a therapeutic alliance through shared decision-making strategies.
Moseley (2002) [41]	The legitimacy of PNE was built in opposition to traditional educational approaches (e.g. back-to-school type) that do not target cognitive-behavioral aspects that may represent potential obstacles to recovery and lack of clinical efficacy. PNE is designed to target these cognitive-behavioral aspects through the reconceptualization of the problem.
Louw et al (2017) [51]	The hypothesis explaining the improvement of the SLR would be that education would positively influence muscle tone and blood circulation.
Lotze et Moseley (2015) [40]	PNE stands in opposition to the structural model of pain, which assumes that pain is a reflection of the state of the tissues and only offers treatments that aim to reduce nociception, the results of which have so far been disappointing. PNE is based on the theory that pain is a perceptual inference that elicits protection, and that pain will be modulated by all credible tissue and contextual (endogenous, exogenous, cognitive, emotional, sensory) cues that urge to protection, rather than the condition of the tissues alone, hence its variable relationship. It emphasizes the protective action of pain, and the fact that nociception is neither necessary nor sufficient to produce pain. According to this theory, changing the meaning of pain is likely to change the pain itself. PNE is also based on the theory that chronic pain is linked to maladaptive plasticity of the nervous system, disturbed inhibitory mechanisms and a less precise representation of the body. It is these mechanisms that are targeted by PNE and other associated therapies.
Blickenstaff et Pearson (2016) [38]	Introducing inconsistent cognitive experiences with the "pain does not equal injury" belief has the potential to decrease perceived pain-related threat. Reducing this threat can potentially: 1) help patients consider the influence they have on recovery; 2) change the way they approach movement, exercise and physical activity; 3) thus people are able to repeat movements with greater ease and frequency, creating more positive adaptations. However, for cognitive and behavioral changes to be stronger and more lasting, the generation of mental (PNE) and physical (movement or exercise) experiences is recommended. PNE reduces movement threat, and subsequent repeated movement confirms reduced threat.
Barbari et al (2020) [14]	 It has been documented that patients with chronic low back pain do not always adhere strictly to exercises, which can affect the outcomes; 2) PNE differs from educational strategies based on a biomedical model, such as those focused on ergonomics, biomechanics like back schools, postural exercises or traditional TM; 3) instead, it is based on a biopsychosocial perspective that is more in line with advances in neuroscience and CLBP research.
Ibrahim et al (2019) [61]	Chronic low back pain is often associated with psychosocial factors such as fear avoidance beliefs, catastrophizing, and negative emotions, which can be addressed by providing educational training.
Nijs et al (2015) [39]	The combination of pain neuroscience education and exercise therapy combined with in vivo exposure principles can desensitize the central nervous system by altering pain memory.

Theoretical framework and effects - Clinician educational resources

References	Specific data
	Must answer the question: "On which theoretical foundations is PNE based?"; "How efficacy of PNE is explained?"
Louw et al (2018) [13]	1)What you need to know about the neuroscience of pain ; 2) Traditional and old pain models ; i) Cartesian model of pain ; ii) Take your foot out
	of the fire; avoid the fire; iii) Put the fire out; iv) Cut the wire (or remove the painful part); 3) Changing beliefs about pain; 4) Input mechanism:
	Tissues; 5) Input mechanism: Environment; 6) Input mechanisms: Peripheral neurogenic; i) Ions channels; ii) Double crush and axoplasmic flow;
	iii) Blood flow; 7) Processing mechanism: Spinal cord, dorsal horn and second-order neurons; 8) Processing mechanism: Brain, the pain
	neuromatrix and functional changes in the brain; 9) Processing mechanism: The pain neuromatrix, yellow flags and "personalization" of the pain
	experience; 10) Output mechanisms: The stress response, endocrine system and immune system; i) Stress response; ii) Clinical manifestation of
	the output systems; 11) Plasticity and merging of systems.
Moseley et Butler (2017) [18]	1) You and your sneaky theories i) theory : a system of ideas intended to explain something ; ii) The biopsychosocial model of pain ; iii) Theory One : The
	Grand Poobah Pain Theory (GPPT); iv) Theory Two: Neurotags, collaboration and competition; v) Theory Three: The cortical body matrix theory; 2)

Supercharge your pain biology ; i) The relationship between damage, nociception and pain ; ii) Detecting tissue based events ; iii) A new look at primary nociception ; iv) What makes primary nociceptors fire? ; v) Two way traffic – efferent function of nociceptors; vi) Time for neuroimmune coupon – hang on to your haats! ; vii) The immune set point, TLR4 and why you should care; viii) The dorsal horn: time for a rethink; ix) Central sensitization; x) Smudging neurotags; xi) The grand finale – a pain mechanisms sheet.

Practical aspects - Scientific literature

References	Specific data
	Must answer the question: "PNE is delivered individually or in a group?"; "How long is a PNE session?"; "How many PNE sessions are delivered?"; "Ow
	what period of time is PNE delivered?"
Adenis et al (2020) [17]	Evaluation and synthesis stage (30 minutes) will be conducted individually, following the 4 hours of educational intervention spread over 4 days, consisting of
	sessions of 30 minutes per day conducted in group.
Agarwal et al (2020) [23]	There were 7 appointments over a 9-month period. Each session included PNE among other things (exam, consultation, home exercise). Each session lasts
	between 25 and 45 minutes.
Anandkumar et al (2017) [24]	For patient A, the first 2 sessions consisted of individual PNE of 50 minutes each. For patient B, the first 2 sessions consisted of individual PNE of
	approximately 30 minutes each.
Bodes Pardo et al (2018) [54]	Two educational group sessions (4 to 6 participants) of 30 to 50 minutes, with the second session taking place one month after the first.
Clarke et al (2011) [5]	2.5 - 4 hours of individual or group instruction. PNE can be provided in one or more sessions.
Dolphens et al (2014) [63]	The first three sessions are educational. These are individual educational sessions lasting approximately thirty minutes, with the exception of session one (a o
	hour group session with six participants per group), and session two (online).
Gallan Martin et al (2019) [53]	Six group PNE sessions (10 hours) were held. The first four sessions lasted 1.5 hours twice a week. A final session of two hours was held to reinforce the magnetic set of the session of two hours are held.
	concepts of the program, plus an educational book. A review session after three months emphasized doubts and problem solving.
King et al (2018) [48]	All participants received PNE as part of their usual NHS care. PNE session was delivered for a group of 10-12 people
Louw et al (2012) [58]	PNE was delivered in a private room for 45 minutes, within the first 75 minutes of the initial history and physical examination session. This initial session w
	followed by 7 physiotherapy sessions which included PNE, as well as other treatment strategies.
Malfliet et al (2017) [20]	Combination of a one group session (6 people per group), one group online session and one individual session (profitability argument) supplemented by an
	information brochure to be read at home.
Mansouri et al (2018) [73]	Pamphlet to be read within 3 weeks.
Moseley (2004) [46]	One individual PNE session (face-to-face) of three hours.
Moseley (2005) [44]	One individual session of two and a half hours.
Moseley et al (2004) [47]	1 individual 3-hour PNE session with a 20-minute break
Moseley (2003) [45]	4 individual sessions of 1 hour spread over 2 weeks or 1 group session of 4 hours (7 to 10 patients)
Nijs et al (2014) [26]	Two to three individual sessions spread across two weeks.

O'Keeffe et al (2015) [69]	Six PNE group sessions over six to eight weeks in groups of up to ten participants. Group sessions last an hour and fifteen minutes (PNE + exercises) with the PNE part lasting thirty minutes per session.
Orhan et al (2019) [28]	Two individual PNE sessions conducted one week apart. The first session lasted between 45 to 60 minutes. The second session lasted 45 minutes.
Pires et al (2015) [53	It took place in an outpatient clinic. PNE consisted of 2 group sessions of 90 minutes each.
Rizzo et al (2018) [67]	PNE was delivered in groups ranging from 1 to 7, twice a week, for 4 weeks
Rufa et al (2018) [31]	There were two semi-standardized individual face-to-face PNE sessions with a two-week interval. Session 1 lasted for sixty minutes, and Session 2 for thirty minutes.
Ryan et al (2010) [68]	Intervention lasting 2 hours and 30 minutes.
Saracoglu et al (2020) [32]	Four PNE sessions were held once a week, individually (face to face) for approximately 40-50 minutes.
Saracoglu et al (2020) [33]	In addition to manual therapy sessions, patients received PNE once a week for 4 weeks individually (face to face) lasting 45 to 50 minutes
Tegner et al (2018) [65]	PNE take different formats ranging from intensive one-on-one sessions, to small group sessions, and large seminars of up to 3 hours. The greatest effect is seen
6()[00]	when the sessions were individual and intensive (not surprising given the complexity and individual nature of the pain).
	when the sessions were manyoual and intensive (not surprising given the complexity and individual nature of the pain).
Tellez-Garcia et al (2015) [49]	30-minute individual PNE sessions, once a week for 2 weeks after dry needling application
Toomey et al (2020) [34]	12 sessions targeting PNE, manual therapy and exercises.
Unhal et al (2020) [70]	PNE is delivered in 40 minute sessions twice a week for 8 weeks
Vier et al (2018) [70]	There are sessions twice a week for 6 weeks individually
Wälti et al (2015) [71]	There are between 2 and 4 education sessions.
Werner et al (2010) [57]	The intervention is delivered in the form of 4 sessions of 30 minutes, with one session per week. Each session include specific educational content and one-on-
	one discussion between the patient and healthcare professional.
Wood et Hendrick (2019) [60]	PNE is delivered in many formats: from one-on-one sessions, to webinars, to group sessions.
Moseley et Butler (2015) [2]	PNE takes several formats: intensive individual sessions, sessions in small groups, or large seminars up to three hours long.
Louw et al (2016) [36]	PNE can be given in 10-15 minutes during the first 50-minute individual physiotherapy session (including anamnesis, physical examination and therapy), and
	then 10 minutes per session during the following 30-minute sessions.
Moseley (2003) [42]	Education can take time. It can be done individually or in a small group.
Moseley et al (2002) [20]	One hour of individual face-to-face educational session, once a week, for four weeks, plus a small workbook.
Louw et al (2017) [51]	Duration of 5 minutes.
Lotze et Moseley (2015) [40]	Several formats: 1) Face-to-face; 2) Small groups; 3) Large groups. Duration up to 3 hours. Approach adapted to preferences and financial considerations.
Blickenstaff et Pearson (2016) [38]	Some data suggest that individual PNE gives better results than PNE delivered in small groups
Barbari et al (2020) [14]	Individually or in a group, 1 to 8 sessions, 30 minutes to 3 hour session, generally once a week for multiple sessions.
Ibrahim et al (2018) [56]	Group sessions of 3 to 5 participants. Sessions begin with 15-20 minutes of interactive discussion followed by a 1-hour lecture.

Group sessions of 3 to 5 participants. Sessions begin with 15-20 minutes of interactive discussion followed by a 1-hour lecture.

 Galan-Martin et al (2020) [25]
 Patients assigned to the experimental group will perform a PNE program consisting of six sessions (10 hours) and eighteen sessions of therapeutic physical

 exercise to be performed in six weeks (18 hours), with a frequency of three sessions per week.

Practical aspects - Clinician educational resources

References	Specific data
	Must answer the question: "PNE is delivered individually or in a group?"; "How long is a PNE session?"; "How many PNE sessions are delivered?"; "Over
	what period of time is PNE delivered?"
Louw et al (2018) [13]	The timing and frequency of PNE sessions vary depending on the study. They can last anywhere from 30 minutes to 4 hours. Frequency can range from 1 single
	session to several sessions over the course of treatment. The most common way of delivering PNE is face-to-face verbal format (individual), but some studies
	have also looked at PNE in groups, using only educational booklets, or remotely (telecare, e-learning, e-mail). The choice of individual, group, or remote format
	could be based on the risk of chronicity.
Moseley et Butler (2017) [18]	1) Example of PNE Group: 8 sessions of 2 hours spread over 6 weeks for a group of 12 in a conference room; 2) Example of Individual PNE: 8 sessions of 30
	minutes (including gradual exposure and functional restoration) in the outpatient clinic of a hospital.

Learning modalities - Scientific literature

Reference	Specific data
	Must answer the question: "What learning strategies, methods and supports were used?"
Adenis et al (2020) [17]	Educational content is presented with the use of simple explanations, pictures and metaphors.
Agarwal et al (2020) [23]	Educational content was presented with thuse of examples, metaphors, and images to facilitate learning about the biology of pain was explored. Several format
	such as verbal discussion, YouTube video, educational cards (e.g. Why do I hurt?), and additional educational materials (figures and concepts from
	Supercharged EP) were used. Educational material was provided to the patient at each session for them to reflect upon, and to allow for further discussion durin
	each visit. Visual aids (e.g. spine models, drawings, flowcharts, and metaphors) were used during PNE sessions.
Anandkumar et al (2018) [24]	Educational content is mainly based on the book Explain Pain, including an individualized curriculum based on the main concerns of patients, is used during
	PNE sessions. Visual aids such as drawings, images, metaphors, flowcharts, PowerPoint presentations, YouTube videos, and animations are also used.
Bodes Pardo et al (2018) [54]	Educational content is based on the books Explain Pain and Pain In Motion. A small booklet is given out at the end of the sessions to reinforce key concepts.
Clarke et al (2011) [5]	Various models are presented to the patient using simple pictures and metaphors to explain the complex neurophysiology of pain.
Dolphens et al (2014) [63]	Educational content is presented verbally (therapist's explanations) and visually (summaries, pictures and computer diagrams). Following each session, patient
	are given a handout summarizing the educational content provided, and are asked to read it thoroughly. Content and pictures are based on the Explain Pain
	educational book.
Gallan Martin et al (2019) [52]	Educational books are given to participants. Reinforcement sessions with answers to questions and problem-solving are proposed.
Louw et al (2012) [58]	Educational content is accompanied by pictures, anecdotes and metaphors
Malfliet et al (2019) [22]	Educational content must be delivered with a strong interaction between the patient and the therapist (focusing on effective communication rather than one-sic
	lecturing).

Malfliet et al (2017) [20]	Examples and metaphors are used based on participants' experiences. Participants were invited to read the information brochure at home between the group and individual sessions. The online session builds on the content of the group session to facilitate a deeper understanding and assessment of knowledge and representations. The information gathered was used to tailor the 3rd session (individual) and Pain Neurophysiology Questionnaire as part of the program to further explore misconceptions and misunderstandings.
Mansouri et al (2018) [73]	The content of the booklet is adapted to the patient's understanding: comprehensive, simple, enjoyable, without imposing time constraints or complexity.
Moseley (2004) [46]	Educational content is presented with hand-drawn drawings and images accompanied by interactive commentary. There were no problem-solving exercises, coping strategies, or role-playing exercises, but examples were used to illustrate the concepts.
Moseley (2005) [44]	Educational content is presented with the use of pictures, examples and metaphors
Moseley et al (2004) [47]	During the educational sessions, people were free to sit, stand, or walk. There were no problem-solving exercises, coping skills training, or role-playing. Diagrams and examples were used to convey the concepts. At the end of the session, each participant received a workbook consisting of 10 sections. Participant were advised to read one section, then answer three questions about the material in it.
Nijs et al (2017) [27]	An information brochure, explanatory manual (e.g. Explain Pain) and website (e.g. Retrain Pain) designed specifically to explain pain to people with persistent pain are used during the PNE sessions.
Nijs et al (2014) [26]	Illustrations, examples and metaphors should be used frequently when delivering educational content to patients. Messages should be tailored to the level of intellectual ability and literacy of the patient.
Nijs et al (2011) [31]	Using a booklet with written recommendations is recommended. Illustrations, examples, and metaphors are often used when delivering educational content to patients. The educational content is presented verbally (explanations by the therapist) and visually (summary, images, and diagrams presented on computer or paper). During sessions, patients are encouraged to ask questions and their input should be utilized to personalize the information.
O'Keeffe et al (2014) [69]	Participants have the opportunity to ask questions. Visual aids are used (PowerPoint, Flipchart) and a copy of the slides is given to the participants. In addition, all participants are invited to read resources on the internet (pain-ed) and will receive brochures on sleep, relaxation, imagery and physical activity
Orhan et al (2019) [28]	The content is supplemented with visual aids such as images, drawings, graphics and metaphors. Patients receive a home education leaflet and are asked to reac it several times and write down any questions they have before the second session. The patients return to discuss their questions and the researchers ensure that everything was understood. Use of PowerPoint "Pain in Motion" support slides was discussed with the patient.
Pires et al (2015) [53]	Educational content is presented with the use of pictures and metaphors.
Rizzo et al (2018) [67]	Participants are encouraged to ask questions. At the end of each session, the information is summarized and a workbook is provided containing the same conter as during the lessons. Each participant is invited to review the material after each lesson.
Rufa et al (2018) [31]	Participants receive a booklet (Why Do I Hurt) to reinforce the content seen in the first session and are invited to read it before the next session.
Ryan et al (2010) [68]	Educational content is delivered using verbal communication, prepared diagrams and freehand drawings. In addition, all participants received a "Back Book"
Saracoglu et al (2020) [32]	Educational content is presented with the use of metaphors, anecdotes, and pictures utilizing PowerPoint.
Saracoglu et al (2020) [33]	To facilitate the transmission of concepts, metaphors, anecdotes, graphics, pictures, disseminated using a PowerPoint, were used.
Tegner et al (2018) [6]	The systematic review includes only verbal PNE, but educational content can be delivered through videos, books and pamphlets.
Tellez-garcia et al (2015) [49]	A PowerPoint presentation based on the Explain Pain educational book is used to facilitate understanding of the concepts. During the sessions, patients are encouraged to ask questions and their input is used to individualize the information they received. Written information on the pain physiology concepts discussed during the sessions is provided as homework between sessions.
Toomey et al (2020) [34]	Resources used are: YouTube video (ex: Taming the beast, understanding pain in less than 5 minutes, TEDx why things hurt, etc.), educational books (ex:

Unal et al (2020) [35]	Educational content is presented with the used of simple pictures, examples, booklets, metaphors. A workbook (with questions / answers) is also used.
Van Wilgem et Keizer (2012) [59]	The concept of sensitization is best explained with a metaphor (e.g. alarm system). If possible, the metaphor should correspond to the patient (e.g.: in connectio with his work, his hobbies).
Wälti et al (2015) [71]	Participants received a copy of the Explain Pain educational book. During the first 2 weeks, participants have to read ten pages of the book each day, and answe a questionnaire about the content.
Wood et Hendrick (2019) [60]	Individual explanations with the use of booklets and books to supplement the explanations.
Moseley et Butler (2015) [2]	PNE draws on conceptual change theory, instructional design and multimedia principles to explain the biology of pain. There are booklets and educational book available for patients.
Louw et al (2016) [36]	PNE works best using pictures, examples, and metaphors to promote deep learning.
Moseley (2003) [42]	 For patients to understand the educational material, it is necessary to use graphics (including whiteboards, hand drawings, and personalized manuals) and metaphors. Deep reconceptualization is facilitated when: a) the learner is motivated; b) the information presented is relevant to them.
Louw et al (2017) [60]	PNE is best delivered with metaphors, examples, and images. For example, the alarm system metaphor can be used to explain central sensitization.
Moseley et al (2002) [20]	A small book is used.
Louw et al (2017) [51]	Verbal explanations and pictures are used.
Barbari et al (2020) [14]	 Use of leaflets/booklets to reinforce the understanding of concepts; 2) Use of diagrams, examples, metaphors to facilitate learning; 3) Use of media such as books, booklets, PowerPoint presentations.
Ibrahim et al (2019) [61]	Verbal education with visual aids such as prepared slides and diagrams, as well as simple cultural metaphors to reinforce certain information.
Ibrahim et al (2018) [56]	An interactive session/discussion is followed by a lecture. Other: 1) visual aids (diagrams, slides); 2) simple and cultural metaphors to reinforce certain information; 3) simple and clear information in simple language to accommodate the participants' low levels of literacy and cultural understanding.
Nijs et al (2015) [39]	Educational content is provided using a Socratic style education and associative learning (violation of expectations). An educational booklet is also used.
Galan-Martin (2020) [58]	 Active listening; 2) Metaphors and visuals used; 3) Ensuring that the information is well understood; 4) Empathy; 5) Sending positive messages; 6) Resources, infographics, and video; 7) Stimulating nature of the group; 8) Individualized educational content.

Learning modalities - Clinician educational resources

References	Specific data
	Must answer the question: "What learning strategies, methods and supports were used?"
Louw et al (2018) [13]	Educational content is not presented in a lecture format, but rather designed as conversations where the person is encouraged to ask questions so that the
	educational material can be personalized. The use of stories, metaphors and examples helps to make the understanding of complex information such as the
	neurobiology of pain easier. The authors also believe they have the power to alleviate fears and promote behavior change. The use of visual resources can help
	promote understanding, retention and use of PNE (as humans are visually oriented). There are many educational resources available to provide educational
	content: patient books, educational cards, posters, workbooks, etc. There is also the possibility of moving away from commercial educational tools and creating
	your own tools (e.g. drawings) to further individualize education. The PNE approach promotes the active participation of the patient by proposing cognitive
	homework to reinforce the concepts that have been discussed.
Moseley et Butler (2017) [18]	1) Metaphor provides an opening to understand the patient's world; 2) Listening to the patient's metaphors, and offering back metaphors combined with
	transformative literal explanations (DIM->SIM) can be a respectful way to challenge unhelpful conceptions; 3) Quotations can be helpful, some learners love it;

4) Treating chronic pain can be a long journey, some metaphors related to perseverance and pacing can be helpful in keeping the person on board. 5) Importance of taking into account the strength, coherence and acceptance of a concept; 6) Importance of taking into account the motivation for change; 7) favoring the position of cognitive guide because it favors the reconceptualization of pain, rather than in a masterful way or by conditioning; 8) the message must be persuasive: understandable, plausible, coherent and attractive; 9) books alone are not enough; 10) PNE uses multimedia principles with a combination of words (printed or spoken) and pictures (illustrations, photos, animations, video); 11) use SIMs and non-DIMs pictures; 12) adapt according to the context

Associated treatment - Scientific literature

References	Specific data
	Must answer the question: "PNE is delivered alone or with other treatments? Which one?"
Adenis et al (2020) [17]	PNE is associated with multimodal and multidisciplinary treatment (balneotherapy, physiotherapy, ergonomics of the spine, stretching, muscle strengthening,
	relaxation, podiatry and speaking time).
Agarwal et al (2020) [23]	PNE is associated with opioid tapering
Anandkumar et al (2018) [24]	Mindfulness, breathing, quota-based reduction in exercises and modification of exercises into social activities, pleasurable activities, and hobbies.
Bodes Pardo et al (2018) [54]	The therapeutic exercise program was a multimodal exercise program that included motor control exercises for the lumbar spine, stretches and aerobic exercises
Clarke et al (2011) [5]	PNE alone or combined with treatment (physiotherapy, pain management program, motor control exercises).
Dolphens et al (2014) [63]	PNE is associated with exercise therapy with particular emphasis on motor control, time-contingent and cognition-focused. A home exercise program is
	established for each patient (with modalities described verbally and visually), who is strongly encouraged to continue during the 1-year follow-up.
Gallan Martin et al (2019) [53]	PNE is associated with group physical exercise, playful, dual task and promoting socialization. Tasks and challenges are carried out at home.
King et al (2018) [48]	PNE alone
Lane et al (2018) [19]	PNE is associated with current therapeutic approaches such as manual therapy and exercises.
Louw et al (2012) [58]	PNE is followed by a graded exercise/exposure approach such as Cognitive Behavioral Therapy (CBT). Therapies include cardiovascular exercise, neural
	mobilization exercises, spinal mobilization and stretching exercises, strengthening exercises, balneotherapy, hot physiotherapy during which the patient must
	focus on breathing, relaxation, summarizing the time, reflecting on goals and progress, and a home exercise program.
Malfliet et al (2019) [22]	PNE can be applied with other physical therapy interventions such as activity therapy, exercise therapy or manual therapy
Malfliet et al (2017) [20]	PNE is associated with motor control focused on cognitive processes and functional dynamic exercises.
Mansouri et al (2018) [73]	PNE is associated with a rehabilitation program
Moseley (2004) [46]	PNE alone
Moseley (2005) [44]	PNE alone
Moseley et al (2004) [47]	PNE alone
Moseley (2003) [45]	PNE is delivered with structured therapeutic exercises on motor control and home exercises. The motor control program is limited to a few variations to take
	into account the specificities of the PNE educational model: 1) no requirement for the position of activation of the trunk muscles, each patient has to choose the
	position where he feel most at home. ugly ; 2) progression implies an intermediate stage of motor imagery; 3) the exhibition focuses on movements and
	activities involving fear; 4) the final stages of progression involve training in conditions of cognitive or psychosocial stress.

Louw et al (2016) [36]	To desensitize the nervous system, PNE is combined with aerobic exercise, relaxation, meditation, manual therapy (joint mobilization, soft tissue treatment neurodynamics, stretching, etc.), diaphragmatic relaxation, goal setting, sleep hygiene, and so forth.
	of peripheral and central contributors to pain. PNE can promote strategies promoting neuroplasticity (or the stimulation of endogenous analgesic substances such as pharmacological means, hypnosis, meditation, or exercise.
Moseley et Butler (2015) [2]	PNE promotes biopsychosocial care, including, but not limited to, multimodal reactivation based on Cognitive Behavioral Therapies. PNE integrates treatme
Wood et al (2019) [60]	PNE can be given alone or in combination with other forms of physiotherapy treatment such as exercise, manual therapy, dry needling, or acupuncture.
Werner et al (2010) [57]	PNE is delivered in addition to their normal treatment.
Wälti et al (2015) [71]	PNE is associated with sensory retraining and motor retraining.
Vier et al (2018) [70]	PNE is associated with a lumbar manipulation (grade V).
Van Wilgem et Keizer (2012) [59]	PNE is associated with a multidisciplinary program.
Toomey et al (2020) [34] Unal et al (2020) [35]	PNE is associated with a multimodal approach based on the biopsychosocial model based on manual therapy and exercises. PNE alone
Tellez-Garcia et al (2015) [49]	PNE is associated with 3 dry-needling therapy sessions on active TrP.
Tegner et al (2018) [6]	PNE can be given alone or in combination with other treatment modalities (e.g. physiotherapy, CBT, manual therapy)
Saracoglu et al (2020) [33]	PNE is combined with manual therapy with a patient-centered model adapting techniques according to the patient's response.
	and pelvic warm-ups and stretching, followed by strengthening exercises. The goal of the program is to improve the strength and flexibility of the erector spi muscles, abdominals, hamstrings and quadriceps.
Saracoglu et al (2020) [32]	PNE is associated with individualized manual therapy based on the patient's response to treatment, as well as a group exercise program that begins with lumi
Ryan et al (2010) [68]	PNE is associated with training circuits based on graduated aerobic exercises and some core / stabilization exercises.
Rufa et al (2018) [31]	PNE alone
Rizzo et al (2018) [67]	PNE is associated with hypnosis
Puentedura et Louw, (2016) [30]	Hands-on therapies may initially seem more appropriate for use in conjunction with PNE, however this article outlines how hand-on therapy (manual therap could potentially be associated with PNE and supervised exercise / movement.
Pires et al (2015) [53]	PNE is associated with a group balneotherapy
Petersen et al (2019) [29]	PNE is integrated with targeted therapeutic exercises
Orhan et al (2019) [28]	PNE alone
O'Keeffe et al (2015) [69]	PNE is associated with progressive exercise circuits including aerobic, flexibility and strengthening exercises + a relaxation component at the end of each classical strengthening exercises are component at the end of each classical strengthening exercises are classical strengthening exercises are classical strengthening exercises.
Nijs et al (2011) [31]	Often followed by various components of a biopsychosocial oriented rehabilitation program, such as stress management, gradual activity and exercise therap
	exercises. The progression is gradual
Nijs et al (2014) [26]	Cognition-centered exercise divided into 2 stages: 1) motor control exercise, 2) functional exercises. Motor control exercises can be preceded by motor imag
Nijs et al (2017) [27]	PNE prepares for the cognitions-centered exercise. It may be associated with the normalization of ill-adapted provocative posture and movement patterns + CBT-I (change in negative thoughts about sleep, sleep restriction therapy, and learning relaxation techniques)

Moseley (2003) [42]	Graded motor and functional tasks, of insufficient intensity to activate the pain neuromatrix, sufficient to selectively activate components of this neuromatrix, and whose progression is effected by increasing the threat (sensory or non-sensory), making it possible to desensitize the neuromatrix
Louw et al (2017) [60]	PNE combined with manual therapy reinforces the message of manual therapy (contingent non-symptom approach)
Moseley (2002) [41]	Manual therapy (spinal mobilization / manipulation, massage, muscle and neurodynamic techniques) + motor control exercise + home exercises
Louw et al (2017) [51]	Grade II manual therapy technique
Lotze et Moseley (2015) [40]	Graded Motor Imagery
Blickenstaff et Pearson (2016) [38]	PNE associated with movement and exercise (e.g. gradual exposure technique), in order to increase the effectiveness of these approaches and reinforce the educational message of PNE
Barbari et al (2020) [14]	 PNE alone; 2) PNE associated with: i) therapeutic exercises; ii) balneotherapy; iii) dry needling; iv) sensory retraining; v) motor retraining (from motor imagery to functional activity; vi) Cognitive Functional Therapy (normalization of postures and behaviors, integration of these standardized patterns into activities of daily living, physical activity program based on patient preferences and impairments).
Ibrahim et al (2019) [61]	PNE is followed by motor control exercise, stretching and aerobic exercises.
Ibrahim et al (2018) [56]	PNE is followed by motor control exercise, stretching and aerobic exercises.
Nijs et al (2015) [39]	PNE is associated with exercise therapy based on in vivo exposure principles (e.g. motor control training, aerobic training or muscle strengthening).
Galan-Martin et al (2020) [25]	PNE is associated with therapeutic physical exercise

Associated treatment - Clinician educational resources

References	Specific data
	Must answer the question: "PNE is delivered alone or with other treatments? Which one?"
Louw et al (2018) [13]	PNE must be integrated with a physical treatment (PNE Plus) as it yields better results than PNE alone. The treatment (PNE Plus) can involve different strategies that can be tailored according to the patient's needs and the clinician's training. A set of pharmalogical and non-pharmalogical strategies, hands-off and hands-on, cognitive, behavioural therapy, etc. can be used (offering a total of 36 treatment options). However, according to evidence-based approaches and pain sciences, the key elements of a PNE+ program are: 1) PNE; 2) aerobic exercise; 3) sleep hygiene; 4) goal setting; 5) relaxation; and 6) education.
Moseley et Butler (2017) [18]	To achieve clinically significant results, PNE is combined with other treatments supported by an evidence-based biopsychosocial framework. PNE interconnects, combines and applies components of other evidence-based approaches such as motor imagery, gradual exercise, Pilates, yoga, taichi, hypnosis and psychological techniques, pharmacology and surgery, among others.

Clinical posture - Scientific literature

References	Specific data
	Must answer the question: "In what mindset or attitude is PNE delivered?"
Anandkumar et al (2018) [24]	Clinicians pay attention to the choice of words and explanatory models (avoiding anatomical and patho-anatomical models) due to their limited effectiveness and the potential for negative impacts (nocebo effect).
Galan Martin et al (2019) [18]	Aware of inhibitory system dysfunction, the exercises will not be symptom contingent and function-centered.

Louw et al (2012) [58]	A manual approach was purposefully excluded because it would contradict PNE: de-emphasizing tissues and focusing on sensitizing the central nervous system
Malfliet et al (2019) [22]	1) PNE must be delivered with a strong interaction between the patient and the therapist (communication rather than one-sided lecture); 2) if the therapist combines PNE with passive strategies, the communication must be adapted to a biopsychosocial (rather than biomedical) framework. Example: prefer the term
	desensitize (bps) rather than restore (biomedical). Choose less threatening and more neutral words (e.g. symptoms rather than pain)
Malfliet et al (2017) [20]	A strong therapeutic alliance is necessary to achieve clinical outcomes for pain.
Nijs et al (2017) [27]	The physical therapist must take into account the thoughts of the person, and extend the reconceptualization of pain to exercise and daily activities
Nijs et al (2014) [26]	More than an educational framework is needed: the term "neuroscience communication of pain" is applicable here. Such communication can pave the way for
	behavior change (including exercise therapy)
Van Wilgem et Keizer (2012) [59]	PNE is delivered according to a biopsychosocial approach to integrate physical, psychological, behavioral and environmental factors to understand why sensitization persists, and to promote their identification
Moseley et Butler (2015) [2]	One of the fundamental principles of PNE is that pain is a truly biopsychosocial phenomenon, where nociception and pain are 2 distinct entities, distinguishing themselves from structural biomedical models (nociception = pain).
Louw et al (2016) [36] Moseley (2003) [42] Diener et al (2016) [12]	A successful and adapted PNE must match the complexity of the person's painful experience. This requires the development of a therapeutic alliance with the person so that they have confidence, and to take into account a certain timing. Education must be an active, adapted and quantified process. Trust and the therapeutic alliance are critical components of PNE, and develop throughout the interview and the clinical examination: the patient must feel heard and examined conscientiously, without using fear-inducing words (nocebo), without focusing the examination on irrelevant details (e.g. tilting of the pelvis) and t be explained in a clear and reassuring manner the results of the examination (placebo). For PNE to work, images, language and explanations that refer to an anxiety-provoking patho-anatomical model should be eliminated, and a language of PNE should be encouraged. PNE-integrated care has many nuances that should not be overlooked, which are rooted in cognitive, motivational, educational domains and various principles of psychology, rather than just PNE. PNE promotes patient empowerment so that they become an actor in their care. PNE not only educates people about pain, but also deconstructs powerful biomedic models that have been shown to have limited efficacy but are likely to undermine the trust of patients in the medical field. PNE also deals with various psychological issues related to pain. Furthermore, it combines the psychological, biological and physiological dimensions of pain treatment to offer a biopsychosocial approach to treating people in pain. This requires spending time with the patient, engaging in meaningful conversations and providing physic support, all of which have been highly valued by patients receiving pain treatment.
Wijma et al (2016) [55]	movement and activity; 4) The need to explain the results of the examination to reduce the threat and empower the patient to make decisions. A biopsychosocial assessment allows an interactive and person-centered PNE
Louw et al (2017) [60]	A combination of PNE and manual therapy reinforces the message of PNE (pain can change independently of tissues) only if there is a shift in communication 1) a focus on function rather than symptom resolution is employed; 2) the strategy is presented as a transitional therapy.
Lotze et Moseley (2015) [40]	1) The importance of message consistency; 2) unifying principle is to provide evidence of safety (reassurance, education, therapeutic alliance): interaction, be caring, take the time, focus on the patient's needs, show that you are informed, explain the pain, explain approaching rehabilitation with simple words provide evidence of safety; 3) cognitive principles: i) rehabilitation should be load- or time-dependent (not pain-dependent); ii) when the pain increases or after the treatment, it is not a symptom of damage, but the sign of an overprotective alarm system; iii) balance between empathy and the need to hold the line to desensitize the nervous system; iv) remind the patient that he is responsible for his body and his rehabilitation; v) lead him to pay attention to the healthy aspective.

	of his body; vi) incorporate exercises at home; vii) teaching the patient to identify sources of safety; viii) learn and implement coping strategies to relieve pain; ix) give the patient the resources to control the situation and achieve his achievable goals; x) and help her understand the principles of slow and steady progression based on a modern understanding of her pain.
Blieckenstaff et Pearson (2016) [32]	 Need for exercise prescription and instructions to be consistent with PNE messages (e.g. not to associate PNE with a contingent symptom approach); For reconceptualization to occur, exploration and experimentation must occur at cognitive (PNE) and physical (gradual exposure) levels: PNE decreases threat to movement, and exposure confirms or refutes this new belief necessary for lasting change.

Clinical posture - Clinician educational resources

References	Specific data
	Must answer the question: "In what mindset or clinical attitude, PNE is delivered?"
Louw et al (2018) [13]	1) Importance of mastering the sciences of pain; 2) the spirit is to plant PNE seeds and water them with compassion; 3) focus on function rather than pain; 4) set limits; 5) all pain is real; 6) movement is the greatest pain reliever in the world; 7) each patient has a brain, and each brain is attached to a person; 8) the therapeutic alliance is a first fundamental step in PNE and essential and fundamental for the success of PNE; 9) the therapeutic alliance requires specific communication (need to be present, receptive, authentic and engaged) and a safe, welcoming, healing context 10) Need to soften words and use less provocative language (sensitive rather than pain for example)
Moseley et Butler (2017) [18]	 PNE requires a biopsychosocial mental framework; 2) broadly speaking, PNE can be a way of reasoning about pain based on concepts from the pain sciences; pay attention to your language: avoid structural metaphors and DIMs, favor SIMs; 4) the appropriate use of humor aids (SIMs); 5) the educator as a cognitive guide, rather than a teacher or learning through conditioning; 6) no ageism (age does not equal decline).

Patient educational resources

References	Content
0	Must answer the question: "what does PNE contain?"
Pain In Motion [63]	The booklet employs simplified literal information, analogies, metaphors, and diagrams to facilitate the transmission of biological concepts related to chronic pain, the distinction between pain and injury, and the contribution of biopsychosocial factors to the maintenance of the problem. Specifically, it covers: 1) the description of nociceptive pathways (nociceptors, ion channels, neurons, action potential, nociception, synapses, inhibitory and facilitating chemicals, spinal cord, descending pro and anti-nociceptive pathways) and pain (the role of the brain); 2) the process of acute pain becoming chronic (central nervous system sensitization at the spinal level); 3) the manifestations of central nervous system sensitization (pain can move, appear in different locations, and be disproportionate after tissue healing); 4) factors that may account for central nervous system to desensitize; and 6) recommendations for desensitizing the nervous system (understanding pain to overcome fear, identifying triggers, engaging in distraction activities, pacing, exercise and gradual activity, cognitive therapy, relaxation techniques).
Retrain Pain Foundation [65]	This educational tool utilizes concise PowerPoint slides, incorporating images, minimal text, metaphors, analogies, simplified literal explanations, and general concepts to facilitate the understanding of biological concepts (pain not necessarily indicating injury, pain context-dependent, perceived threat affects pain, chronic pain correlates with nervous system sensitization, biopsychosocial factors may sustain pain, chronic pain can improve). Specifically, the tool covers: 1) nociceptive pathways and pain; 2) peripheral sensitization; 3) central sensitization; 4) descending pro and anti-nociceptive pathways; 5) the influence of downward modulation on pain and the nervous system; 6) pain and modulation being dependent on the brain's perceived threat; 7) factors that may contribute to nervous system sensitization (beliefs, negative thoughts and emotions, hyper-focus, sleep, opioids); and 8) recommendations to facilitate nervous system desensitization (pain understanding, mindfulness meditation, sleep hygiene, appropriate medication).

Louw and al (2015) [10]	Introduction: Role of the book: 1) how pain works and 2) provide strategies to reduce pain. Section 1 helps to understand the difference between pain and disability; Section 2 provides an understanding of imaging and physical testing. Section 3 seeks to deconstruct beliefs about back pain: 1) deconstruction of patho-anatomical beliefs (osteoarthritis equals pain, pain equals injury), pathomechanical (hyper lordosis, hyper kyphosis, postures, MI inequalities, lack of strength, and lack of flexibility) and treatment (rest, medication, losing weight, etc.) + promoting strong spinal tissues. Section 4: helps to understand 1) why the pain persists (alarm system remains hypersensitive beyond the tissue healing time linked to the presence of stressors)?; 2) What does sensitization of the nervous system look like clinically?; 3) why certain pains are unpredictable (presence of more ion channels in the wall of the nerves because hypersensitive, these ion channels respond to various mechanical, thermal or chemical stimuli, therefore the pain can be caused by stress, cold or immune changes); 4) other (disability, pain not equal to injury) 3) why some pain is diffuse (explanation of central sensitization and immune changes); Section 5 explains: 1) what are the treatment options? 2) How does the treatment work? 3) How to apply the treatment? Treatment Options: i) Understanding Pain, ii) Problem Solving, iii) Pain Coping, iv) Specific Medications, v) Relaxation Techniques, vi) Gradual Exposure, vii) Aerobic Physical Activity, viii) Goal Setting , ix) drink water, x) pacing, xi) sleep hygiene, xii) lifestyle: limit sedentary lifestyle, weight loss, smoking cessation.
Lehman [66]	Why are you telling me all this? 1) Makes it possible to change the way of approaching the problem; 2) establish its own rehabilitation program; 3) better manage pain and 4) adopt the right behaviors. Chapter I: explanation of the mechanisms of pain (importance of understanding pain, what pain is, nociception, ascending nociceptive pathways, role of the spinal cord and brain, role of pain, pain memory, sensitization, desensitization, cortical reorganization) to better understand key messages, contributing factors and treatment; Chapter 2: the key messages make it possible to reconceptualize pain, to change beliefs and to adopt the right behaviors vis-à-vis pain (pain is an alarm that protects, pain weakly correlated with lesions, pain equals sensitivity rather than lesion, factors may influence sensitivity, protection may be over-amplified and persist after tissue has healed, pain may be bizarre, we are strong and adaptable, we respond positively to physical stress, dysfunction not equal to pain, no need to be put back in place, no movement is prohibited forever); Chapter 3: helps to understand that pain is not just a reflection of the state of the tissues, and how it can be influenced by a multitude of anatomical factors (osteoarthritis), physical (physical constraint, strength and flexibility, asymmetry) , emotional (catastrophism, kinesiophobia), behavioral (avoidance, persistence) or related to lifestyle (stress, sleep), social and comorbidities + demystification of the importance of certain factors (physical constraint, osteoarthritis, strength and flexibility, physical asymmetry). Understand that we don't need to be put in place by someone: concepts of self-efficacy, adapted behaviors and our body's ability to adapt; Chapter 4: 1) identify relevant factors; 2) appropriate recovery program in place. Treatment options: i) sports and physical activity, ii) resuming hobbies and activities that matter; iii) stress management; iv) sleep hygiene; v) gradual exposure; vi) diet, vii doing things that make us happ
Butler and Moseley (2013) [8]	Section 1: evokes general concepts with anecdotes, scientific studies, metaphors and analogies: 1) pain is normal; 2) nociception is neither necessary nor sufficient to produce pain; 3) pain is produced by the brain based on the perceived threat to the body; 4) pain depends on the context; 5) the amount of pain does not reflect the amount of degeneration or injury. Section 2 describes the neurophysiology of nociceptive pathways (nociceptors, ion channels, neurons, action potential, nociception, synapses, inhibitory and facilitative chemicals, spinal cord, descending pro and anti-nociceptive pathways), pain (neuromatrix, role of the brain) and other protective systems (sympathetic, endocrine, immune and motor), with simplified literal explanations, diagrams, graphs, metaphors and analogies to reinforce the general concepts mentioned above. Section 3: discusses the physiology and particularities (solidity, healing, inflammation, acid release) of different body tissues (skin, muscle, disc, bone, joint, dorsal root ganglion, nerves) with a focus on neuropathic pain. Section 4: describes the alteration of the central nervous system at the level of the spinal cord and the brain (sensitization of the pain neuromatrix) and of the other homeostatic systems (sympathetic, endocrine, immune and motor) inducing an over activity of the protective systems bodily + how thoughts and beliefs can help maintain this state. Section 5: discusses modern models of pain management (neuromatrix model, onion model, fear-based models, evolutionary model, clinical decision-based model), the need for the patient to be an actor of its management, and the exploration of cognitive-behavioral concepts (fears, coping, behavior) related to pain. Section 6: discusses care based on neuroscience, namely: 1) understanding your pain; 2) gradual exposure and pacing; 3) accessing the virtual body to desensitize the neuromatrix and protective systems.
Moseley and Butler (2015) [9]	Part 1: Reminder of concepts covered in Explain Pain: 1) pain is not a good indicator of tissue condition; 2) imaging is not a good indicator of pain; 3) pain is produced by the brain 100% of the time; 4) pain depends on context, all with literal explanations, anecdotes, reflective questions, scientific studies to reinforce these concepts. Part 2: presentation of the theory that pain occurs when the brain concludes that there is more evidence of harm than evidence of safety for body tissues. 1) SIMs/DIMs are neuro-immune circuits; 2) SIMs represent all situations (thoughts, behavior, words, context) that make a person feel safer; 3) DIMs represent all situations that make a person feel in danger; 4) explanation of the protectometer tool and the biological influence of DIMs/SIMs on it; 5) Protectometer user manual; 6) other concepts: i) other protection systems (sympathetic, endocrine, immune) can load the protectometer; ii) protection systems can become overactive when danger signals persist; iii) harmless events can now cause the pain, without really being the problem (with literal explanations & metaphors). Part 3: preparation: 1) check for red flags; 2) negative & positive bioplasticity; 3) how to promote positive bioplasticity (remove DIMs, add SIMs). Part 4: Focused on management based on the concepts covered: 1) planning for recovery; 2) promote active coping strategies; 3) limit maladaptive behaviors

	and promote gradual exposure and pacing with reasoning based on the biology of pain (tissues, pain, avoid flare-ups); 4) understand that knowledge is analgesic because it is the brain that produces pain; 5) favoring a SIM rather than DIM language; 6) understand that the protector is the key (with drawings, literal
	explanations, metaphors, graphics).
Louw and al (2013) [64]	This book uses texts, diagrams, metaphors, anecdotes, graphics, analogies, scientific studies, simplified biological explanations, and general concepts to
	facilitate the transmission of biological concepts. Section 1: discusses tissue injury, inflammation, neuroimmune sensitization, tissue scarring, desensitization +
	contrast with people whose pain threshold does not return to baseline despite tissue scarring. Link between nerve sensitization and decreased function. Proposal
	of psychosocial factors that may explain the maintenance of awareness. Explanation of ion channels to give meaning to unpredictable pain (stress, illness, cold,
	atmospheric pressure); Section 2: explanation of secondary hyperalgesia via the immune system (history of noisy neighbors); Section 3: explanation of
	secondary hyperalgesia via hypervigilance and pronociceptive facilitation (history of the PDG of the body); Section 4: explanation of the pain neuromatrix +
	Hebb's law (ex: board of directors, aerial maps); Section 5: pain is produced by the brain according to the perceived threat to protection (metaphor of the sprain
	and the bus); Section 6: physiological explanation of stress and long-term impact on protection systems + proposal of possible stressors (lion metaphor); Section
	7: proposed treatment to desensitize the nervous system: 1) knowledge; 2) progressive aerobic exercise; 3) specific medication (antidepressant type) to target
	descending inhibition; 4) sleep hygiene; 5) pacing and goal setting.
Moseley (2007) [68]	This book mainly uses anecdotes and metaphors to facilitate the transmission of biological concepts. Chapter 1: pain is an essential protection system for
3 () []	survival; Chapter 2: pain is a protector; Chapter 3: pain does not measure the condition of tissues; Chapter 4: pain is a complex conscious experience based on
	the interaction of many signals; Chapter 5: pain is the conscious reflection of the perceived threat to our tissues; Chapter 6: the virtual body; Chapter 7: Pain
	depends on the answer to the question "how dangerous is it"?; Chapter 8: Nociception is not sufficient to produce pain; Chapter 9: Nociception is not sufficient
	NOR necessary to produce pain; Chapter 10: the brain analyzes information based on perceived vulnerability and threat; Chapter 11: Neural networks that
	produce pain become more sensitive when pain persists.

Supplementary Table 2: General data of included studies

Author, year of publication, Study population Objective(s) Design Important result(s) Intervention and Outcome measure(s) title and country of the study comparator Adenis et al (2020), PNE + multidisciplinary Chronic low back pain Evaluation of efficacy of PNE Protocol study of a Roland Morris Disability N / A Evaluation of the effectiveness compared with conventional Randomized Control Trial Questionnaire (primary management versus of pain neurophysiology Conventional Education + education in a outcome) education compared to a multidisciplinary multidisciplinary program on conventional education management function at D90. combined with multidisciplinary rehabilitation, in chronic low back pain patients: a protocol for a randomized controlled trial, France [17] Dolphens et al (2014), PNE + cognition-targeted Chronic spinal pain To assess the effectiveness of Multi-center, triple-blind, The main outcome measures N / AEfficacy of a modern motor control training a modern neuroscience two-arm (1:1) randomized are pain (including symptoms neuroscience approach versus versus usual care approach, compared to usual clinical trial with 1-year and indices of central sensitization) and self-reported usual care evidence-based follow-up. physiotherapy care evidence-based physiotherapy on pain, physiotherapy, for reducing disability. disability and brain pain and improving characteristics in chronic functioning in patients with CSP spinal pain patients: protocol of a randomized clinical trial, Belgium [62] Lane et al (2018), The PNE training versus Chronic neck or back pain To determine the effectiveness Cluster randomized trial The primary outcome will be N/A effectiveness of training continue with usual care of providing physical the Patient Reported Outcomes therapists with PNE training Measurement Information physical therapists in pain neuroscience education on on patient-centered outcomes System (PROMIS) Physical patient reported outcomes for Function computer-adapted for patients with chronic neck patients with chronic spinal or back pain undergoing care test (PF-CAT) pain: a study protocol for a by a physical therapist cluster randomized controlled trial, USA [19] Malfliet et al (2017), Applying PNE + cognition-targeted Nonspecific chronic spinal To provides the treatment Protocol of a randomized N/A N/A controlled clinical trial contemporary neuroscience in motor control training pain protocol used in a large exercise interventions for versus usual care randomized controlled trial chronic spinal pain: treatment that aimed to assess the physiotherapy protocol, Belgium [20] effectiveness of a modern neuroscience approach compared to usual care evidence-based physiotherapy Malfliet et al (2019), The CBT-I combined with the Chronic spinal pain To examine whether CBT-I Protocol of a randomized Primary outcome measure: N / Aadded value of cognitive modern neuroscience combined with the modern controlled clinical trial self-reported pain severity

Protocol study (10)

behavioral therapy for	approach (PNE + cognition-		neuroscience approach (i.e.		(Brief Pain Inventory)	
insomnia to current best	targeted motor control)		pain neuroscience education			
evidence physical therapy for	versus modern		plus cognition-targeted			
chronic spinal pain: protocol	neuroscience approach		exercise therapy is superior to			
of a randomized controlled	alone.		the modern neuroscience			
clinical trial, Belgium [21]			approach alone to reduce pain			
			(primary outcome measure).			
O'Keeffe et al (2015),	Individualiz5.ed cognitive	Nonspecific chronic low	To examine the clinical	Protocol of a randomized	The two primary outcomes of	N / A
Individualised cognitive	functional therapy (CFT)	back pain	effectiveness of CFT, based on	controlled clinical trial	interest will be functional	
functional therapy compared	versus combined exercise		whether participants in the		disability (ODI) and pain	
with a combined exercise and	and pain education class		CFT arm report significant		intensity (NRS)	
pain education class for			improvements in the short,			
patients with non-specific			medium and long term on			
chronic low back pain: study			measures of functional			
protocol for a multicentre			disability and pain intensity,			
randomised controlled trial,			relative to those allocated to			
Ireland [69}			combined exercise and pain			
			education classes.			
Vier et al (2018), Effects of	Spinal manipulation + PNE	Chronic nonspecific low	To verify the pain control,	Protocol of an assessor and	The measures will be applied	N / A
spinal manipulation and pain	versus sham treatment +	back pain	functional and	subject blinded, 2-arm,	at baseline, six weeks, and	
education on pain in patients	PNE	-	neurophysiological effects of	randomized sham-	three months after	
with chronic low back pain: a			spinal manipulation, and pain	controlled trial	randomization. The primary	
protocol of randomized sham-			education in individuals with		outcome will be a pain	
controlled trial, Brazil [70]			chronic nonspecific LBP		intensity at six weeks	
			Ĩ		postrandomization	
Werner et al (2010). The	PNE versus normal	Nonspecific sub-	To evaluate whether a specific	Protocol of a cluster	The primary outcome is	N / A
COPE LBP trial: Cognitive	treatment	acute/chronic low back pain	cognitive based education	randomized controlled trial	function (disability) assessed	N/A
Patient Education for Low	ucament	of more than four weeks but	program for patients with LBP	randoniized controlled that	by the Roland Morris	
					Disability Questionnaire	
Back Pain - a cluster		less than 1 year's duration	in primary care is more			
randomized controlled trial in			effective than normal care in		(RMQ)	
primary care, Norway [57]			terms of increased function			
Ibrahim et al (2019), Effects	motor control exercise +	Chronic low back pain	To investigate the effects of	Protocol of an assessor-	Participants will be assessed	N / A
of motor control exercise and	PNE versus motor control		motor control exercise (MCE)	blind, 3-arm parallel	pre-intervention, immediately	
patient education program in	exercise versus PNE		and patient education (PE) for	randomized clinical trial	post-intervention and at 3-	
the management of chronic			the management of CLBP		month post-intervention.	
low back pain among			among community-dwelling		Primary outcomes will be pain	
community-dwelling adults in			adults in rural Nigeria		intensity and functional	
rural Nigeria: a study protocol					disability	
for a randomized clinical trial,						
Nigeria [61]						
Galan-Martin et al (2019).	PNE + therapeutic exercise	Chronic spinal pain	To compare the effectiveness	Randomized multicentre	Outcomes : quality of life. The	N / A
Pain neuroscience education	versus		of a PNE and PE combination	clinical trial protocol	outcome variables will be	
and physical exercise for	usual physiotherapy		therapy program versus usual		measured at the beginning of	
patients with chronic spinal			physiotherapeutic treatment		the intervention, after the	
pain in primary healthcare: a			used in PC physiotherapy units		intervention (week 11), at six	
			for CSP.		months, and a year.	
					montino, and a jour.	

Spain [18]

Randomized Control Trial (14)

Author, year of publication,	Intervention and	Study population	Objective(s)	Design	Outcome measure(s)	Important result(s)
title and country of the study	comparator					
Bodes Pardo et al (2018), Pain Neurophysiology Education	Group of therapeutic exercise + PNE <i>versus</i>	Chronic low back pain > 6 months	To assess the effect of a pain neurophysiology education	Single-blind randomized	The primary outcome was pain intensity rated on the	At 3-month follow-up, a large change in pain intensity
and Therapeutic Exercise for	therapeutic exercise		(PNE) program plus		numerical pain rating scale	(numerical pain rating scale:
Patients With Chronic Low			therapeutic exercise (TE) for		which was completed	2.2; 2.93 to 1.28; P<0,001;
Back Pain: A Single-Blind			patients with chronic low back		immediately after treatment	dZ1.37) was observed for the
Randomized Controlled Trial,			pain (CLBP)		and at 1- and 3-month follow-	PNE plus TE group, and a
Spain [54]					up	moderate effect size was
						observed for the secondary outcome measures
						outcome measures
Galan-Martin et al (2020),	PNE + therapeutic exercise	Chronic spinal pain	To compare the effectiveness	Randomized multicentre	The main measure of outcome	The experimental treatment
Pain Neuroscience Education	versus		of a PNE and PE combination	clinical trial	was the difference between	improved quality of life (d =
and Physical Therapeutic	usual physiotherapy		therapy program versus usual		groups in the change in health-	1.8 in physical component
Exercise for Patients with			physiotherapeutic treatment		related quality of life (HRQL)	summary), catastrophism (d =
Chronic Spinal Pain in			used in PC physiotherapy units		at different times (initial	1.7), kinesiophobia (d = 1.8),
Spanish Physiotherapy			for CSP.		assessment, post-intervention,	central sensitization (d = 1.4),
Primary Care: A Pragmatic					and 6 months). The Spanish	disability (d = 1.4), pain
Randomized Controlled Trial,					version of the SF-36 v2 health	intensity $(d = 3.3)$, and
Spain [25]					survey was used	pressure pain thresholds (d =
						2). Differences between the
						groups (p < 0.001) were
						clinically relevant in favour of
						the EG. Improvements post-
						intervention (week 11) were
						maintained at six months
Moseley et al (2004), A	individual PNE versus back	Chronic low back pain	To evaluate the effect of a	Blinded randomized	Cognitions were evaluated	There was a significant
randomized controlled trial of	anatomy education	patients	formal intensive	controlled trial	using the Survey of Pain	treatment effect on the
intensive neurophysiology			neurophysiology education		Attitudes (revised)	SOPA(R), PCS, SLR, and
education in chronic low back			program in chronic LBP		(SOPA(R)), ; Pain	forward bending. There was a
pain, Australia [47]			patients		Catastrophizing Scale (PCS);	statistically significant effect
					Behavioral measures included	on RMDQ; however, the size
					the Roland Morris Disability	of this effect was small and
					Questionnaire (RMDQ), and 3	probably not clinically
					physical performance tasks;	meaningful
					(1) straight leg raise (SLR), (2)	
					forward bending range, and (3)	
					an abdominal "drawing-in"	
					task, which provides a	
					measure of voluntary	
					activation of the deep	

					abdominal muscles	
Moseley (2003), Joining Forces – Combining Cognition-Targeted Motor Control Training with Group or Individual Pain Physiology Education: A Successful Treatment For Chronic Low Back Pain, Australia [45]	Individualize8d PNE <i>versus</i> group PNE	Chronic unremittent low back pain	To answer the two questions posed above with the hypotheses: (i) intervention based on a cognition-specific motor control training approach combined with pain physiology education is effective in reducing pain and disability associated with chronic LBP, and (ii) group physiology education is cheaper than, and equally as effective as individualized physiology education	Randomized comparative trial	 8-item Roland Morris Disability Questionnaire (RMDQ) 14 and (0-10) Numerical Rating Scale (NRS) for pain. The first assessment, second assessment (36 ± 4 days later), and third assessment (25 ± 2 days later) were performed. A follow- 88up telephone assessment was conducted 348 ± 13 days after the third assessment. 	Both groups reduced pain (numerical rating scale) and disability (Roland Morris Disability Questionnaire). IE showed bigger decreases, which were maintained at 12 months (P<0.05 for all). The combined motor control and education approach is effective. Although group education imparts a lesser effect, it may be more cost- efficient
Orhan et al (2019), Culture- sensitive and standard pain neuroscience education improves pain, disability, and pain cognitions in first- generation Turkish migrants with chronic low back pain: a pilot randomized controlled trial, Belgium [28]	culture-sensitive PNE <i>versus</i> standard PNE	First generation Turkish migrants with chronic low back pain	To compare the effectiveness of culture-sensitive and standard pain neuroscience education (PNE) on pain knowledge, pain intensity, disability, and pain cognitions in first-generation Turkish migrants with chronic low back pain (CLBP)	Pilot randomized controlled trial	Primary (pain knowledge, pain intensity, and disability) and secondary outcomes (pain beliefs, catastrophization, and fear of movement) were evaluated at baseline, immediately after the second session of PNE (week 1), and after 4 weeks	There was a significant main effect of time in pain knowledge ($p < .001$), pain intensity ($p = .03$), disability (p = .002), organic and psychological pain beliefs ($p =$.002, $p = .01$), catastrophization ($p = .002$), and fear of movement ($p =$.02). However, no significant difference was found between groups in terms of all outcome measures ($p > .05$)
Pires et al (2014), Aquatic exercise and pain neurophysiology education versus aquatic exercise alone for patients with chronic low back pain: a randomized controlled trial, Portugal [53]	aquatic exercise and PNE <i>versus</i> aquatic exercise alone	Chronic low back pain	To compare the effectiveness of a combination of aquatic exercise and pain neurophysiology education with aquatic exercise alone in chronic low back pain patients	Single-blind randomized controlled trial	The primary outcomes were pain intensity (Visual Analogue Scale) and functional disability (Quebec Back Pain Disability Scale) at the baseline, 6 weeks after the beginning of the aquatic exercise program and at the 3 months follow-up	Fifty-five participants completed the study. Analysis using mixed-model ANOVA revealed a significant treatment condition interaction on pain intensity at the 3 months follow-up, favoring the education group (mean SD change: -25.4 ± 26.7 vs $-6.6\pm$ 30.7, P < 0.005)
Saracoglu et al (2020), The effectiveness& of pain neuroscience education combined with manual therapy and home exercise for chronic low back pain : a single-blind randomized controlled trial, Turkey [32]	PNE + manual therapy + home exercise <i>versus</i> manual therapy + home exercise	Chronic low back pain	To investigate the short- and mid-term effects of pain neuroscience education (PNE) combined with manual therapy (MT) and a home exercise program (HEP) on pain intensity, back performance, disability, and kinesiophobia	Prospective, randomized, controlled, single-blind study	The participants' pain intensity, disability, low back performance, and kinesiophobia were assessed. All assessments were executed before intervention, at 4 weeks, and at 12 weeks post- intervention by the same	Analysis of pain level (<i>p</i> < .05), back performance (<i>p</i> < .05), disability (<i>p</i> < .05) and kinesiophobia (<i>p</i> < .05) revealed significant time, group, and time-by-group interaction effects. The participants in Group 1

			in mations - with stars 1		him dash share to all the	and determined and the second
			in patients with chronic low		blinded physiotherapist	exhibited greater improvement
			back pain (CLBP)			in terms of pain intensity and
						kinesiophobia compared to the
						participants in Group 2 and the
						control group. Level of
						disability was significantly
						decreased in both Group 1 and
						Group 2 compared to the
						control group
Saracoglu et al (2020). The	PNE + manual therapy	Chronic low back pain	To investigate the short-term	Randomized controlled	Outcome measures were the	Both intervention and control
short-term effects of	versus traditional patient		effects of PNE in patients with	study	Short Form-36 (SF-36) quality	group had significant effects of
neuroscience pain education	education + manual therapy		CLBP treated with manual		of life (QoL) questionnaire and	pain and quality of life.
on quality of life in patients			therapy		the Numeric Pain Rating Scale	Between-group comparisons
with chronic low back pain: A					for low back pain measured	revealed a significant
single-blinded randomized					before and after treatment	difference in pain severity (p =
controlled trial, Turkey [33]						0.01) and SF-36 physical
2 Diens						function subscale ($p = 0.04$) in
						favor of the NPE group
			T 14 1 1 4			
Téllez-Garcia et al (2015),	PNE + trigger point dry	Chronic non specific low	To determine the short-term	Preliminary clinical trial	Pain intensity (Numerical Pain	Patients treated with TrP-DN +
Neuroscience education in	needling versus trigger	back pain	effects of trigger point dry		Rating Scale, 0-10), disability	EDU experienced a
addition to trigger point dry	point dry needling alone		needling (TrP-DN) alone or		(Roland-Morris Disability	significantly greater reduction
needling for the management			combined with neuroscience		Questionnaire-RMQ-,	of kinesiophobia (P = 0.008)
of patients with mechanical			education on pain, disability,		Oswestry Low Back Pain	and greater increases in PPT
chronic low back pain: A			kinesiophobia and widespread		Disability Index-ODI),	over the transverse process of
preliminary clinical trial,			pressure sensitivity in patients		kinesiophobia (Tampa Scale of	L3 ($P = 0.049$) than those
Spain [49]			with mechanical low back pain		Kinesiophobia-TSK), and	patients treated only with TrP-
			(LBP)		pressure pain thresholds (PPT)	DN. Both groups experienced
					over the C5-C6 zygapophyseal	similar decreases in pain, ODI
					joint, transverse process of L3	and RMQ, and similar
					vertebra, second metacarpal,	increases in PPT over the
					and tibialis anterior muscle	C5/C6 joint, second
					were collected at baseline and	metacarpal, and tibialis anterior
					1-week after the intervention	after the intervention (all, P >
						0.05)
Unal et al (2020),	Myofacial induction	Chronic low back pain	to comparatively investigate	Prospective, randomized-	Primary outcome measure :	Within both groups, all
Investigating the effects of	therapy versus PNE	×	the effects of myofascial	controlled and single-blind	Roland Morris disability	outcome scores showed a
myofascial induction therapy			induction therapy (MIT)	study	questionnaire, McGill pain	significant improvement
techniques on pain, function			against pain neuroscience	,	questionnaire	(p < 0.05). After 8-week, SF-
and quality of life in patients			education (PNE) on pain and		1	36 physical function, physical
with chronic low back pain,			function in patients with			role and mental health scores
Turkey [35]			chronic low back pain (CLBP)			significantly improved in MIT
1 arroy [55]			entoine low back pain (CLDF)			
						group compared with PNE
						group, finger floor test score
						significantly decreased in MIT
						group compared with PNE
						group, and FABQ score
						significantly decreased in PNE

						group compared with MIT group ($p < 0.05$)
Wälti et al (2015), Short-term effect on pain and function of neurophysiological education and sensorimotor retraining compared to usual physiotherapy in patients with chronic or recurrent non- specific low back pain, a pilot randomized controlled trial, Switherland [71]	Multimodal treatment (PNE + sensory training + motor control training) <i>versus</i> usual physiotherapy treatment	Patients with nonspecific chronic low back pain, considerable disability (five or more points on the Roland and Morris Disability Questionnaire (RMDQ) and medium or high risk of poor outcome on the Keele Start Back Tool (KSBT)	To investigate the feasibility of MMT, prior to a larger RCT, with focus on patients' adherence and the evaluation of short-term effects on pain and disability of MMT when compared to usual physiotherapy	Randomized controlled trial (RCT)	The primary outcome was pain (NRS 0-10) and the secondary outcome was disability (RMDQ	Pain reduction (NRS; [95% CI]) was 2.14 [1.0 to 3.5] in the MMT and 0.69 [-2.0 to 2.5.] in the UPT. The between-group difference was 1.45 [0.0 to 4.0] ($p = 0.03$), representing a moderate effect size of 0.66 [- 0.1 to 1.5]. Reduction in disability on the RMDQ was 6.71 [4.2 to 9.3] in MMT and 4.69 [1.9 to 7.4] in UPT, with a non-significant between-group difference of 2.02 [-1.5 to 5.6] ($p = 0.25$)
Moseley (2002), Combined physiotherapy and education is efficacious for chronic low back pain, Australia [41]	PNE + manual therapy + exercise versus management as directed by their general practitioners	Chronic low back pain	To determine the efficacy of a combined physiotherapy treatment that comprised all of these strategies	Randomized controlled trial	The following items were used as outcome measures: the 18- item Roland Morris Disability Questionnaire (RMDQ; Roland and Morris 1983) and the 0-10 Numerical Rating Scale (NRS) for pain	Outcome data from 49 subjects (86%) showed a significant treatment effect. The physiotherapy program reduced pain and disability by a mean of 1.5/10 points on a numerical rating scale (95% CI 0.7 to 2.3) and 3.9 points on the 18-point Roland Morris Disability Questionnaire (95% CI 2 to 5.8) , respectively. The number needed to treat in order to gain a clinically meaningful change was 3 (95% CI 3 to 8) for pain, and 2 (95% CI 2 to 5) for disability. A treatment effect was maintained at one- year follow-up
Louw et al (2016), The effect of manual therapy and neuroplasticity education on chronic low back pain: a randomized clinical trial, USA [51]	Neuroplasticity explanation (part of PNE) + manual therapy technique <i>versus</i> mechanical explanation + manual therapy technique	Chronic low back pain	To determine if a neuroplasticity educational explanation for a manual therapy technique will produce a different outcome compared to a traditional mechanical explanation	Randomized clinical trial	Following consent, demographic data were obtained as well as pain ratings for low back pain (LBP) and leg pain (Numeric Pain Rating Scale), disability (Oswestry Disability Index), fear-avoidance (Fear- Avoidance-Beliefs Questionnaire), forward flexion (fingertips-to-floor), and straight leg raise (SLR) (inclinometer)	There were no statistically significant interactions for LBP ($p = .325$), leg pain ($p = .172$), and trunk flexion ($p = .818$) between the groups, but SLR showed a significant difference in favor of the neuroplasticity explanation ($p = .041$). Additionally, the neuroplasticity group were 7.2 times (95% confidence interval = 1.8-28.6) more likely to improve beyond the MDC on

						the SLR than participants in
						the mechanical group
Ibrahim et al (2018), Motor	Patient education + Motor	Chronic low back pain	To assess the feasibility of	Pilot randomized clinical	Feasibility was assessed	Many patients were willing to
control exercise and patient	control exercise versus	Chrome low back pain	implementing MCE and PE	trial	through recruitment rate,	participate in the study and the
education program for low	patient education versus		program in the management	ti ki	treatment compliance,	recruitment rate was 77%.
					-	
resource rural community	motor control exercise		CLBP in a low resource rural		retention/dropout rate, report	Treatment compliance in
dwelling adults with			Nigerian community		of adverse events, perceived	all the three groups were $>$
chronic low back pain: a pilot					helpfulness, overall	65% for supervised treatment
randomized clinical trial,					satisfaction, and clinical	sessions and $\leq 50\%$ for
Nigeria [56]					outcome of pain (numeric pain	prescribed home program.
					rating scale) and functional	Retention rate was high and
					disability (Oswestry Disability	greater overall satisfaction with
					Index)	the interventions was reported.
						Compared with the baseline,
						all the three groups improved

Other clinical trial (4)

Author, year of publication,	Intervention and	Study population	Objective(s)	Design	Outcome measure(s)	Important result(s)
title and country of the study	comparator					
Mansouri et Kostur (2018),	PNE booklet +	Chronic low back pain	To assess the relevance of	Non randomized controlled	Each patient, on inclusion in	Concerning the simple verbal
Preliminary descriptive study	multidisciplinary program		PNE education in the	trial	the study (D0), performed two	scale, a better score is observed
on pain neurophysiology	versus multidisciplinary		physiotherapy management of		assessments. The first assesses	for the experimental group on
education for patients with	program alone		chronic low back pain patients		the intensity of his lumbar pain	the intensity of the pain and its
chronic low back pain: Use of			and to assess whether its		using the simple verbal scale	functional repercussions in the
an illustrative brochure, France			illustration by a brochure is an		(EVS): at the present moment	chronic low back pain patient
[73]			appropriate tool for the		(EVS 1), usually for the last	who received a brochure
			transmission of knowledge to		eight days (EVS 2) and the	compared to those who did no
			this patient.		most intense for the eight last	receive one, in the short and
					days (EVS 3). The second	long terms. Discussion The
					evaluates the functional impact	small population sample
					of lumbar pain in activities of	included does not allow
					daily living (ADL) through the	conclusions to be drawn on the
					Dallas questionnaire which	effects of the illustration by a
					goes beyond the limitations of	brochure on NPD for the
					the patient's physical abilities	population with chronic low
					by exploring daily activities	back pain compared to those
					(Dallas 1), occupations and	not receiving the brochure
					hobbies (Dallas 2), anxiety and	during their rehabilitation, on
					depression (Dallas 3) and	the intensity of the pain and it
					sociability (Dallas 4).	functional repercussions in the
						activities of daily living, in the
						short, medium and long term.
Moseley (2004), Evidence for	individual PNE or	Moderately disabled	To determine if a relationship	Quasi experimental study	Multiple regression analysis	There was a strong relationshi
a direct relationship between	individual spine physiology	chronic low back pain	exists between change in pain		evaluated the relationship	between cognitive change and
cognitive and physical change			cognitions and change in		between change in pain	change in straight leg raise

during an education			physical performance when		cognitions measured by the	(SLR) and forward bending
intervention in people with			chronic LBP patients		survey of pain attitudes	(r=0:88 and 0.79, respectively,
chronic low back pain,			participate in a single one-to-		(SOPA) and the pain	P < 0.01), mostly explained by
Australia [46]			one education intervention		catastrophizing scale (PCS)	change in the conviction that
			during which they have no		and change in physical	pain means tissue damage and
			opportunity to be active		performance, measured by the	catastrophizing
					straight leg raise (SLR) and	
					standing forward bending	
					range	
Rufa et al (2018), The use of	PNE	A 3 month or greater	The primary purpose of this	Quasi-experimental	Subjects' perception of PNE	Subjects consistently reported a
pain neuroscience education in		history of lower back (>	quasi experimental feasibility	feasibility study	was measured after the second	positive experience with PNE.
older adults with chronic back		50%) and/or lower	study was to determine		session and gait speed, pain	There were statically significant
and/or lower extremity pain,		extremity pain	whether adults over the age of		disability, and fear of	positive improvements in gait
USA [31]			65 with chronic pain find PNE		movement were measured pre-	speed, pain disability, and fear
			to be understandable, relevant		and post-PNE	of movement after the
			and helpful			intervention
Louw et al (2017), De-educate	de-education session	Chronic low back pain	To determine if a brief de-	Case serie	Prior to and immediately after	Significant changes were found
to re-educate: Aging and low	regarding aging and low		education session regarding		the education pain ratings for	in positive shifts with LBP (p =
back pain, USA [37]	back pain (LBP) (part of		aging and low back pain		LBP and leg pain (numeric	0.002), leg pain (p = 0.042),
	PNE)		(LBP) can shift pain ratings,		pain rating scale-NPRS), fear-	FABQ-physical activity
			fear-avoidance beliefs, beliefs		avoidance (fear avoidance	subscale ($p = 0.004$) and active
			regarding aging and LBP, and		belief questionnaire - FABQ),	trunk forward flexion (p <
			limited active trunk flexion		beliefs regarding aging and	0.001)
					LBP (Likert scale) and active	
					trunk flexion were measured	

Systematic review (4)

Author, year of publication,	Intervention and	Study population	Objective(s)	Design	Outcome measure(s)	Important result(s)
title and country of the study	comparator					
Clarke et al (2011), Pain	PNE	Chronic low back pain	To investigate the evidence for	Systematic review with	The main outcome measures	Two moderate quality RCTs
neurophysiology education for			PNE in the management of	meta-analysis	were pain, physical-function,	(n=122) were included in the
the management of individuals			patients with CLBP		psychological-function, and	final review. According to the
with chronic low back pain:					social-function	CBRG criteria there was very
systematic review and meta-						low quality evidence that PNE
analysis. United Kingdom [5]						is beneficial for pain, physical-
						function, psychological-
						function, and social-function.
						Meta-analysis found PNE
						produced statistically
						significant but clinically small
						improvements in short-term
						pain of 5 mm (0, 10.0 mm)
						[mean difference (95%CI)] on
						the 100 mm VAS

Nurrephysiological Pain OR usual care with CLDP neurephysiological pain meta-analysis summarized in a random trial studies (6 low and 1 hg Education for Patients With education (NPE) for patients effect meta-analysis for pain, quality were included. Chronic Low Back Pain A with chronic low back pain disability, and belavioral Statistically significant Systematic Review and Meta- (CLBP) anticudes differences in pain, in favor Analysis. Danemark.[6] .	Neurophysiological Pain Education for Patients With Chronic Low Back Pain: A		-				Seven randomized-controlled
Education for Patients With education (NPE) for patients effect meta-analysis for pati, disability, and behavioral Statistically significant Systematic Review and Meta- (CLBP) attitudes differences in pain, in forer. Analysis, Danemark.[6] NPE, were found after treatment, NMD=1.03 059 confidence interval [Cl], -0.2 to -1.52), and after 3 methy NPE, were found after treatment, NMD=1.09 (05% CL], -0.2 to -1.52), and after 3 methy Systematic Review and Meta- (CLBP) attitudes inferences in pain, in forer. Analysis, Danemark.[6] NPE, were found after treatment, NMD=1.09 (05% CL], -0.2 to -1.52), and after 3 methy inferences in pain, in forer. Systematic Review and Meta- (CLBP) attitudes inferences in pain, in forer. Analysis, Danemark.[6] NPE, were found after inferences in pain, in forer. Systematic Review and Meta- (CLBP) attitudes inference in forer. Gound in the NPE group after treatment, SMD=-0.3 (05% CL) in favor of NPE in reduction in favor of NPE in reduction Tampa Seake of Kinesiophob was not statistically signifierat WMD=-5.0 (05% CL), -1.3 disperience VMD=-5.0 (05% CL), -1.3 disperience in favor of NPE in reduction in favor of NPE in reduction Tampa Seake of Kinesiophob was not statistically sign	Education for Patients With Chronic Low Back Pain: A	OR usual care	with CLBP	neurophysiological pain	meta-analysis	summarized in a random	trial studies (6 low and 1 high
Chronic Low Back Pair: A with chronic low back pain disability, and behavioral Statistically significant Systematic Review and Meta- (CLBP) attitudes differences in pain, in favor Analysis: Danemark.[6] NPE, were found after reatment, WMD-1.03 (95) confidence interval [C1], -02 io -1.52, and after 3 months WMD-1.00 (95) confidence interval [C1], -02 io 1.52, and after 3 months WMD-1.00 (95) WMD-1.00 (95) confidence interval [C1], -02 io 1.52, and after 3 months WMD-1.00 (95) WMD-1.00 (95) confidence interval [C1], -02 io 1.52, and after 3 months WMD-1.00 (95) WMD-1.00 (95) confidence interval [C1], -02 io 1.52, and after 3 months Sund in the NPE group after is ginficant lower disabilitys io and its 3 wonths is ginficant lower disabilitys io and its 3 wonths in favor of NPE in reduction reatment, SMD0.38 (95) in favor of NPE in reduction ramp Scale of Kinesispheb was of statistically significant wither 3 months WMD0.94 (95) ci, -1.20 in favor of NPE in reduction ramp Scale of Kinesispheb was of statistically significant wither 3 months WMD0.94 (95) ci, -1.20 wither 3 months	Chronic Low Back Pain: A				-		that studies (0 low and 1 lligh
Systematic Review and Meta- Analysis. Danemark. [6] the set of the				education (NPE) for patients		effect meta-analysis for pain,	quality) were included.
Analysis. Danemark.[6] NPE, were found after treatment, WMD1.03 (95) confidence interval [C1], -0.2 to -1.52), and after 3 month WMD1.09 (95% C1, -2.17 0.00). Furthermore statistical significant lower disability we found in the NPE group after treatment, SMD0.47 (95) C1, -0.80 to -0.13) and after months months SMD0.38 (95% C1 -0.47 (95) C1, -0.80 to -0.13) and after months SMD0.38 (95% C1 months SMD0.37 (95% C1, -1.56) 0.74 to -0.02). The difference in favor of NPE in reduction Tampa Scale of Kinessiophob wean to statistically signifieration WMD-=5,73 (95% C1, -1.56) vo.2.14) and after 3 months WMD=-0.94 (95% C1, -6.28) WMD=-0.94 (95% C1, -6.28) 4.40)				with chronic low back pain		disability, and behavioral	Statistically significant
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confidence interval [C1], -0.2 to -1.52), and after 3 months WMD1.09 (95% C1, -2.17 0.00). Furthermore statistical significant lower disability w found in the NPE group afte treatment, SMD0.47 (95% C1, -0.80 to -0.13) and after months SMD0.38 (95% C1 -0.74 to -0.02). The differenc in favor of NPE in reduction Tampa Scale of Kinesiophob was not statistically significan WMD=-5.73 (95% C1, -13.6 to 2.14) and after 3 months WMD0.94 (95% C1, -6.28 4.40)	Analysis. Danemark.[6]						NPE, were found after
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0.74 to -0.02). The difference in favor of NPE in reduction Tampa Scale of Kinesiophob was not statistically significan WMD=-5.73 (95% CI, -13.6 to 2.14) and after 3 months WMD=-0.94 (95% CI, -6.28 4.40)							CI, -0.80 to -0.13) and after 3
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to 2.14) and after 3 months WMD=-0.94 (95% CI, -6.28 4.40)							
WMD=-0.94 (95% CI, -6.28 4.40)							
4.40)							
Wood et al (2019). A PNE could be delivered in Chronic low back pain To evaluate randomized Systematic review with The outcome measures of pain A total of 6,767 papers were							4.40)
	Wood et al (2019). A	PNE could be delivered in	Chronic low back pain	To evaluate randomized	Systematic review with	The outcome measures of pain	A total of 6,767 papers were
systematic review and meta- isolation or in combination controlled trials comparing the meta-analysis and disability were included found, eight were included (n	systematic review and meta-	isolation or in combination		controlled trials comparing the	meta-analysis	and disability were included	found, eight were included (n =
analysis of pain neuroscience with other forms of effectiveness of PNE on pain for this review. The principal 615). Meta-analysis for shore	analysis of pain neuroscience	with other forms of		effectiveness of PNE on pain		for this review. The principal	615). Meta-analysis for short-
education for chronic low back physiotherapy treatment: and disability in CLBP summary method was mean term pain (n = 428)	education for chronic low back	physiotherapy treatment:		and disability in CLBP		summary method was mean	term pain $(n = 428)$
pain: Short-and long-term versus waitlist controls, difference between-groups demonstrated a WMD of 0.7	pain: Short-and long-term	versus waitlist controls,				difference between-groups	demonstrated a WMD of 0.73
outcomes of pain and physiotherapy, other assessed at short term (<12 (95%CI -0.14, 1.61) on a ter	outcomes of pain and	physiotherapy, other				assessed at short term (<12	(95%CI -0.14, 1.61) on a ten-
	-						point scale of PNE against no
	[,]						
						-	
treatment term follow-up. Adverse PNE (GRADE analysis low						-	
treatment term follow-up. Adverse PNE (GRADE analysis low events were also captured evidence). When PNE						where mentioned	
treatment term follow-up. Adverse PNE (GRADE analysis low events were also captured evidence). When PNE where mentioned alongside physiotherapy							
treatment term follow-up. Adverse PNE (GRADE analysis low events were also captured evidence). When PNE where mentioned alongside physiotherapy interventions were grouped for							
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treatmentterm follow-up. AdversePNE (GRADE analysis lowevents were also capturedevidence). When PNEwhere mentionedalongside physiotherapyinterventions were grouped fpain (n = 212), a WMD of 1was demonstrated (95% CI							1.08, 1.56, $p < 0.00001;$
treatmentterm follow-up. AdversePNE (GRADE analysis lowevents were also capturedevidence). When PNEwhere mentionedalongside physiotherapyinterventions were grouped fpain (n = 212), a WMD of 1was demonstrated (95% CI							GRADE analysis moderate
treatment term follow-up. Adverse PNE (GRADE analysis low events were also captured evidence). When PNE where mentioned alongside physiotherapy interventions were grouped f pain (n = 212), a WMD of 1. was demonstrated (95% CI 1.08, 1.56, p < 0.00001;							evidence). Short-term
treatmentterm follow-up. AdversePNE (GRADE analysis lowevents were also capturedevidence). When PNEwhere mentionedalongside physiotherapyinterventions were grouped fpain (n = 212), a WMD of 1was demonstrated (95% CI1.08, 1.56, p < 0.00001;							disability (RMDQ) meta-
treatment term follow-up. Adverse PNE (GRADE analysis low events were also captured evidence). When PNE where mentioned alongside physiotherapy interventions were grouped f pain (n = 212), a WMD of 1. was demonstrated (95% CI 1.08, 1.56, p < 0.00001; GRADE analysis moderate evidence). Short-term							analysis demonstrated a WMD
treatment term follow-up. Adverse PNE (GRADE analysis low events were also captured evidence). When PNE where mentioned alongside physiotherapy interventions were grouped f pain (n = 212), a WAD of 1. was demonstrated (95% CI 1.08, 1.56, p < 0.00001; GRADE analysis moderate evidence). Short-term disability (RMDQ) mea-							of 0.42 (95%CI 0.28, 0.56; p <
treatment term follow-up. Adverse PNE (GRADE analysis low events were also captured evidence). When PNE where mentioned alongside physiotherapy interventions were grouped f pain (n = 212), a WMD of 1. was demonstrated (95% CI 1.08, 1.56, p < 0.0001); GRADE analysis moderate evidence). Short-term disability (RMDQ) meta- analysis demonstrated a WM							0.00001; n = 362; GRADE
treatment term follow-up. Adverse PNE (GRADE analysis low events were also captured e vidence). When PNE where mentioned alongside physiotherapy interventions were grouped f pain (n = 212), a WAD of 1. vas demonstrated 95% CI 1.08, 1.56, p < 0.0001; GRADE analysis moderate evidence). Short-term disability (RMDQ) meta- analysis demonstrated a WM of 0.42 (95%CI 0.28, 0.56; p							analysis moderate evidence);
treatment term follow-up. Adverse PNE (GRADE analysis low events were also captured evidence). When PNE where mentioned alongside physiotherapy interventons were grouped for pain (n = 212), a WMD of 1. Was demonstrated (95% CI - 1.08, 1.56, p < 0.0000); GRADE analysis moderate evidence). Short-term disability (RMDQ) meta- analysis demonstrated a WM of 0.42 (95%CI 0.28, 0.56; p - 0.0001; n = 362; GRADE							
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tratment emblow-up. Adverse emblow-up. Adverse emblow-up. Adverse events were also captured evidence). When PNE where mentioned alongside physiotherapy interventions were grouped f pain (n = 212), a WMD of 12 was demonstrated (95% CT 1.08, 1.56, p < 0.0001); GRADE analysis moderate evidence). Short-term disability (RMDQ) meta- analysis demonstrated a WM of 0.42 (95%CT 0.28, 0.56; p 0.0000); n = 362; GRADE analysis moderate evidence) interventions were grouped f could (95%CT 0.28, 0.56; p 0.0000); n = 362; GRADE analysis moderate evidence) whereas the addition of PNE							whereas the addition of PNE to physiotherapy interventions
treatment tem follow-up. Adverse PNE (GRADE analysis low events were also captured e vidence). When PNE where mentioned alongside physiotherapy interventions were grouped for phin (n = 212), a WMD of 1.2 was demonstrated (95% CI = 1.08, 1.56, p < 0.0001); a start of evidence). Short-term with the start of the start of evidence interventions were evidence interventions and start of evidence interventions were grouped for evidence interventions were gr							
treatment tem follow-up. Adverse PNE (GRADE analysis low events were also captured e vidence). When PNE where mentioned alongside physiotherapy interventions were grouped for phin (n = 212), a WMD of 1.2 was demonstrated (95% CI = 1.08, 1.56, p < 0.0001); a start of evidence). Short-term with the start of the start of evidence interventions were evidence interventions and start of evidence interventions were grouped for evidence interventions were gr							physiotherapy interventions demonstrated a WMD of 3.94

						0.00001; GRADE analysis
						moderate evidence
Barbari et al (2020),	communicative and	Chronic low back pain	To investigate the	Systematic review	evaluate at least one of the	24 randomized controlled trials
Effectiveness of	educative strategies		effectiveness of		three following outcomes: 1)	which intervention included
communicative and educative	(including PNE) aimed at		communicative and educative		maladaptive behavior	communicative and educative
strategies in chronic low back	increasing compliance with		strategies on 1) patient's low		modification,	strategies were
pain patients : a systematic	exercise, modifying		back pain		2) compliance with exercise or	selected. Most of the studies
review, Italy [14]	patient's maladaptive		awareness/knowledge, 2)		3) patient's LBP	were judged as low risk of bias
	behavior or LB		maladaptive behavior		awareness/knowledge assessed	and Cohen's Kappa was
	awareness/knowledge		modification and 3)		with objective measures,	excellent (=0.822).
	versus Waiting lists, usual		compliance with exercise in		patient-reported	Interventions
	care, placebo, no		patients with chronic low back		questionnaires or other	addressed were cognitive
	intervention, active or		pain		modalities	behavioral therapy as unique
	passive treatments, other					treatment or combined with
	educative interventions					other treatments (multimodal
						interventions), coaching,
						mindfulness, pain science
						education, self-management,
						graded activity and graded
						exposure

Other literature review (1)

Author, year of publication,	Intervention and	Study population	Objective(s)	Design	Outcome measure(s)	Important result(s)
title and country of the study	comparator					
Malfiet et al (2019), Best	The best evidence non-	Chronic low back pain	Therefore, this paper provides	Up-to-date evidence from	pain and function as outcomes	Most physically inactive
Evidence Rehabilitation for	invasive rehabilitation		a state-of-the-art overview of	systematic reviews, meta-	for chronic low back pain	therapies should not be
Chronic Pain Part 3: Low	(including PNE)		the best evidence non-invasive	analysis and available	management	considered for CLBP
Back Pain, Belgium [22]			rehabilitation for CLBP	treatment guidelines		management, except for pain
				(nonsystematic review)		neuroscience education and
						spinal manipulative therapy if
						combined with exercise
						therapy, with or without
						psychological therapy.
						Regarding active therapy, back
						schools, sensory discrimination
						training, proprioceptive
						exercises, and sling exercises
						should not be considered due
						to low-quality and/or
						conflicting evidence

Qualitative study (1)

Author, year of publication,	Intervention and	Study population	Objective(s)	Design	Outcome measure(s)	Important result(s)
title and country of the study	comparator					
King et al (2018), Pain	PNE	Chronic low back pain	To explore the extent, and	Qualitative study with	semi-structured interview	We observed varying degrees
Reconceptualisation after Pain			nature, of patients'	thematic analysis (inductive		of (1) degrees of
Neurophysiology Education in			reconceptualization of their	& deductive)		reconceptualization, (2)
Adults with Chronic Low			chronic low back pain (CLBP)			personal relevance, (3)
Back Pain: A Qualitative			following PNE			importance of prior beliefs, and
Study, United Kingdom [48]						(4) perceived benefit of PNE.
						We observed varying degrees
						of reconceptualization from
						zero to almost complete, with
						most participants showing
						partial reconceptualization.
						Personal relevance of the
						information to participants and
						their prior beliefs were
						associated with the degree of
						benefit they perceived from
						PNE. Where benefits were
						found, they manifested as
						improved understanding,
						coping, and function

Case studies / series (6)

Author, year of publication,	Intervention and	Study population	Objective(s)	Design	Outcome measure(s)	Important result(s)
title and country of the study	comparator					
Agarwal et al (2020),	PNE in conjunction with	75-year-old female with	This case study demonstrates	Case sutdy	Pre-intervention and post-	The attending physician
Physician-Delivered Pain	monitored tapering of	chronic low back pain	the delivery of PNE by an		intervention Yellow Flags	tapered opioids and other
Neuroscience Education for	opioids and other		internal medicine physician to		Questionnaire (YFQ) scores	medicine associated with her
Opioid Tapering: A Case	medication.		a patient with chronic pain and			CLBP, depending on her
Report, USA [23]			opioid use			responses. Over the 12-week
						period (89 days since
						discharge from the hospital),
						the patient's CLBP decreased
						from 7/10 to 0/10 (Figure 1)
						and opioids and
						antidepressants were
						completely abolished (Figure
						2). Both of these positive
						results were still intact at the
						final six-month follow-up
Anandkumar et al (2018),	PNE (with individualized	two patients, aged 35 and	This report is a potential first-	Case series	EA Inventory (EAI), Numeric	At discharge, they were pain-
Effect of physical therapy	curriculum), mindfulness,	45 years, respectively, who	time description of the		Pain Rating Scale (NPRS);	free and fully functional,

management of nonspecific	breathing, quota-based	presented with chronic	successful physical therapy		Short Form 36 (SF-36) health	which was maintained at a s
low back pain with exercise	reduction in exercises and	nonspecific low back pain	management of EA in chronic		survey and Global Rating of	month follow-up
addiction behaviors: A case	modification of exercises	having exercise addiction	NSLBP using PNE (by		Change (GROC) scale were	
series, Canada [24]	into social participation,	behaviors	adopting a pain curriculum),		used as outcome measures.	
	pleasure activities and		mindfulness, breathing, quota-		Both EAI and NPRS were	
	hobbies		based reduction in exercises		measured at baseline and	
			and activity modification		anticipated to be used at the	
					beginning of each treatment	
					session with the GROC	
					measured after the first therapy	
					session. SF-36 was measured	
					at baseline and intended to be	
					used after discharge from	
					treatment	
Louw et al (2012), Use of an	Treatment consisted of an	A 64-year-old female with	First, the report aims to show	Single case study	A physical examination, the	She attended twice a week
abbreviated neuroscience	abbreviated PNE approach,	history of chronic low back	how a NE session can be		Numeric Pain Rating Scale	4 weeks, or 8 visits total. I
education approach in the	exercises (range of motion,	pain	applied in a time frame that is		(NPRS), Oswestry Disability	NE, the patient reported NI
reatment of chronic low back	stretches, and		clinically reasonable		Index (ODI), Fear-Avoidance	= 9/10; ODI = 54%; FABQ
oain: a case report, USA [58]	cardiovascular), and aquatic		(approximately 1 hour).		Beliefs Questionnaire (FABQ),	= 25/42,; FABQ-PA = 20/
	therapy		Second, it aims to describe the		and Zung Depression Scale	and Zung = 58. Immediate
			clinical reasoning process used		were assessed during her initial	following the 75-minut
			in determining that a NE		physical therapy visit,	evaluation and NE session,
			session was needed for this		immediately after her first	patient reported improvem
			patient. Third, the exact		physical therapy session, and	in all four outcome measu
			content of NE sessions are		at 7-month follow-up	most notably a reduction ir
			poorly described, and this case			FABQ-W score to 2/42 and
			report aims to provide			FABQ-PA to 1/24. At a
			clinicians with a concise and			month follow-up, all outco
			effective way to deliver this			measures continued to b
			intervention to a patient with			improved.
			central sensitization, using			
			strategies, metaphors, and			
			descriptions			
Moseley (2005), Widespread	PNE	A thirty-six year old female	Using a single case design, we	Case report	The abdominal drawing-in	Before education there w
brain activity during an		with a history of chronic	were interested in whether		task, which involves a gentle	widespread brain activit
abdominal task markedly		disabling low back pain (~	pain physiology education had		drawing-in of the lower	during performance of the
duced after pain physiology		4.5 years since onset with a	an effect on the pattern of		abdomen, was used for the	including activity in corti
lucation: fMRI evaluation of		fall at work) and with no	brain activity during		study. Accurate performance	regions known to be invol
single patient with chronic		neural signs	performance of this abdominal		can be verified by a trained	in pain, although the task
ow back pain, Australia [44]			task. Changes in cortical		physiotherapist and confirmed	not painful. After educat
			activation should provide		using real-time ultrasound. The	widespread activity was ab
			insight into the nature of the		following self-report measures	so that there was no bra
			effect of pain physiology		were also used: McGill Pain	activation outside of the
			education on motor tasks		Questionnaire, Roland Morris	primary somatosensory co
					Disability Questionnaire, Fear	
					Avoidance Beliefs	
					Questionnaire, physical	

					Efficacy Questionnaire.	
Peterson et al (2019), Physical	Targeted exercise approach	Three non-consecutive	To describe the physical	Case series	Outcome measures were	By discharge, all patients ha
Therapy Management of	of hip abductor	patients with chronic low	therapy management and		administered at baseline, 4	made clinically important
Patients With Chronic Low	strengthening + additional	back pain: -a 77-year-old	outcomes of 3 patients with		weeks, discharge, and 3-month	improvements in pain (3- to
Back Pain and Hip Abductor	treatments including heel	man, a 78-year-old woman,	CLBP matching a previously		follow-up. Outcome measures	point reduction on the Nume
Weakness, USA [29]	lift and pain neuroscience	and an 85-year-old woman	identified subgroup		were : NPRS, Global Rating of	Pain Rating Scale), functio
	education when indicated		characterized by substantial		Change (GROC), Owestry	(10- to 16-point change on t
			hip abductor weakness		Disability Index (ODI) and	Modifi ed Oswestry Disabil
					Fear Avoidance Beliefs	Index), and perceived
					Questionnaire (FABQ).	improvement (6-7 on Glob
						Rating of Change Scale).
						Lumbar range of motion w
						painless, and hip abductor
						strength was improved from
						+/5 to 3 +/5 in all 3 patient
						These gains were maintain
						at 3-month follow-up
Toomey et al (2020), How	12-week multimodal	A 44-year-old female	This case study will discuss	Case report	Outcome measure Score	The patient had a significant
manual therapy provided a	approach consisting of	presented to physiotherapy	the physiotherapy management		Numerical Pain Rating Scale at	reduction in the Numerica
gateway to a biopsychosocial	manual therapy, exercise	with a 13-year history of	of a 44-year-old woman with		rest 5/10 Oswestry Disability	Pain Rating Scale (NPRS),
management approach in an	rehabilitation, and PNE	persistent pain, having had	chronic postsurgical low back		Index 42% Fear-Avoidance	Oswestry Disability Inde
adult with chronic post-		a spinal fusion 12 years	pain using manual therapy,		Beliefs Questionnaire 18/24	(ODI) and the Fear Avoidated
surgical low back pain: a case		prior, following a skiing	pain neuroscience education			Belief Questionnaire Physic
report, New Zealand [34]		accident	and a progressive exercise			Activity Subscale (FABQ-F
			program			scores following the
						intervention. She returned
						running and cycling, reporti
						that pain was something sl
						would 'work with instead
						against'

Expert opinion (14)

Author, year of publication,	Intervention and	Study population	Objective(s)	Design	Outcome measure(s)	Important result(s)
title and country of the study	comparator					
Nijs et al (2014), A modern	Modern neuroscience	Chronic spinal pain	This perspective paper	Perspective paper	N / A	N / A
neuroscience approach to	approach (including PNE)		explains why and how such an			
chronic spinal pain: combining			approach to CSP can be			
pain neuroscience education			applied in physical therapy			
with cognition-targeted motor			practice			
control training, Belgium [26]						
Nijs et al (2011), How to	PNE	Unexplained chronic	In what follows the reader is	Practice guideline	N / A	N / A
explain central sensitization to		musculoskeletal pain	provided with a brief overview			

patients with 'unexplained'			of the clinical evidence of pain			
chronic musculoskeletal pain:			physiology education in			
practice guidelines, Belgium			patients with chronic			
[31]			musculoskeletal pain. The			
			largest part of the paper is			
			dedicated to practice			
			guidelines on how to apply			
			pain physiology education in			
			patients with chronic			
			musculoskeletal pain			
Van Wilgen et Keizer (2012),	the sensitization model	Chronic pain	To provide a clinical education	Review article	N / A	N / A
The sensitization model to			model for nurses who work			
explain how chronic pain			with patients with chronic pain			
exists without tissue damage,						
The Netherlands [59]						
Moseley and Butler (2015),	PNE	People in pain	1) To describe the historical	Critical review	N / A	N / A
Fifteen years of explaining			context and beginnings of EP,			
pain: the past, present, and			suggesting that it is a			
future, Australia.[2]			pragmatic application of the			
			biopsychosocial model of pain,			
			but differentiating it from			
			cognitive behavioral therapy			
			and educational components of			
			early multidisciplinary pain			
			management programs; 2) To			
			address common			
			misconceptions of EP that			
			have emerged over the last 15			
			years, highlighting that EP is			
			not behavioral or cognitive			
			advice, nor does it deny the			
			potential contribution of			
			peripheral nociceptive signals			
			to pain; 3) To explain that EP			
			is grounded in strong			
			theoretical frameworks, that its			
			targeted effects are			
			biologically plausible, and that			
			available behavioral evidence			
			is supportive; 4) To update			
			available meta-analyses with			
			results of a systematic review			
			of recent contributions to the			
			field and propose future			
			directions by which we might			
			enhance the effects of EP as			
			part of multimodal pain			
			rehabilitation			

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Louw et al (2017), The	PNE	Chronic pain	The article systematically	Perspective article	N / A	N / A
clinical application of teaching			discusses key elements of PNE			
people about pain, USA [36]			including examination,			
			educational content, and			
			delivery methods, merging of			
			PNE with movement, goal			
			setting, and progression			
Moseley (2003), A pain	A pain neuromatrix	Chronic Pain	This paper presents an	Masterclass	N / A	N / A
neuromatrix approach to	approach (including PNE)		approach to rehabilitation of			
patients with chronic pain,			pain patients			
Australia [42]						
Diener et al (2016)	The interview process	Chronic musculoskeletal	This article highlights the	Perspective article	N / A	N / A
Listening is therapy : patient	focusing on a pain science	pain	interview process focusing on	r enspective uttole	10772	10774
interviewing from a pain	perspective (before PNE)	pani	a pain science perspective as it			
science perspective, USA [12]	perspective (before FINE)					
science perspective, USA [12]			relates to screening patients,			
			establishing psychosocial			
			barriers to improvement, and			
			pain mechanism assessment			
Wijma (2016). Clinical	clinical biopsychosocial	Chronic pain	To describe the use of the Pain	Perspective article	N / A	N / A
biopsychosocial physiotherapy	assessment is recommended		- Somatic factors - Cognitive			
assessment of patients with	prior to PNE		factors - Emotional factors -			
chronic pain: the first step in			Behavioral factors - Social			
pain neuroscience education,			factors - Motivation - model			
Belgium [55].			(PSCEBSM-model) during the			
			intake, as well as a pain			
			analysis sheet			
Louw (2017) A Clinical	PNE and manual therapy	Chronic pain	To explore the notion of PNE	Perspective article	N / A	N / A
	r NE and manual therapy	Chronic pain	-	r erspective article	N/A	N/A
Perspective on a Pain			and manual therapy co-			
Neuroscience Education			existing			
Approach to Manual Therapy,						
USA [60]						
Lotze & Moseley (2015),	Key aspects of modern pain	Chronic pain	A brief overview is provided	Narrative review	N / A	N / A
Theoretical Considerations for	rehabilitation (including		of the key aspects of modern			
Chronic Pain Rehabilitation,	PNE)		pain rehabilitation and the			
Germany [40]			considerations that should lead			
			our interaction with patients			
			with chronic pain			
Blieckenstaff & Pearson	Conceptual framework of	Chronic pain	This article will introduce a	Perspective article	N / A	N / A
(2016), Reconciling	kinesthetic education that is	Chronic pain	conceptual framework of	r enspective article	177 A	177 A
movement and exercise with	consistent with and		kinesthetic education that is			
pain neuroscience education:	reinforces pain		consistent with and reinforces			
A case for consistent	neuroscience education		pain neuroscience education.			
education, Canada [38]			This article will also provide			
			some specific guidance for			
			integrating pain neuroscience			

			education with exercise and			
			movement in a more congruent			
			manner			
Nijs et al (2017), In the spine	This approach includes	Chronic low back pain	To show that CLBP is also	Narrative review and expert	N / A	N / A
or in the brain? Recent	cognitively preparing		characterized by differences in	opinion		
dvances in pain neuroscience	patients for exercise therapy		the morphology and			
pplied in the intervention for	using (therapeutic) pain		functionality of the brain.			
ow back pain, Belgium [27]	neuroscience education,		Understanding these brain			
	followed by cognition-		changes in CLBP improves			
	targeted functional exercise		our understanding not only of			
	therapy		pain symptoms, but also of			
			prevalent CLBP comorbidities			
			like sleep disturbances and			
			fear avoidance behaviour. The			
			second part of the paper			
			explains how clinicians can			
			apply our current			
			understanding of			
			contemporary pain			
			neuroscience to improve care			
			for people with CLBP.			
Puentedura et Flynn (2016),	PNE and manual therapy	Chronic low back pain	To review the literature	Narrative review	N / A	N / A
Combining manual therapy			supporting the inclusion of			
with pain neuroscience			manual therapies in the			
education in the treatment of			therapeutic management of			
chronic low back pain: A			CLBP by re-thinking the value			
narrative review of the			of "hands-on" interventions in			
literature, USA [30]			combination with PNE and			
			supervised exercise			
Nijs et al (2015) Exercise	Integrating pain	Chronic pain	Here the authors explain how	Perspective article	N / A	N / A
therapy for chronic	neuroscience education		musculoskeletal therapists can			
musculoskeletal pain:	with exercise therapy		alter pain memories in patients			
Innovation by altering pain			with chronic musculoskeletal			
memories. Belgium [39]			pain, by integrating pain			
			neuroscience education with			
			exercise interventions. The			
			latter includes applying graded			
			exposure in vivo principles			
			during exercise therapy, for			
			targeting the brain circuitries			
			orchestrated by the amygdala			
			(the memory of fear center in			
			the brain)			

Author, year of publication,	Intervention and	Study population	Objective(s)	Design	Outcome measure(s)	Important result(s)
title and country of the study	comparator					
Pain In Motion, Pourquoi ma	PNE	Chronic pain	This brochure is intended to	Educational booklet for	N / A	N / A
douleur persiste-t-elle ?			explain to you how the human	patient		
Belgium [63]			body's pain processing system			
			works. Let's start by describing			
			the nervous system. The			
			changes that occur when the			
			pain becomes chronic and the			
			phenomenon of increased			
			sensitivity to pain will also be			
			reprimanded. You will			
			understand what can explain			
			your pain by means of the			
			most recent scientific			
			discoveries. You will also find			
			in this brochure advice that			
			could be useful to you in your			
		daily life.				
Retrain Pain [65] Pl	PNE	Chronic pain	This free online course teaches	Educational website for	N / A	N / A
			you an evidence-based	patient		
			approach to reducing your			
			symptoms and getting back to			
			the life you want.			
Louw et al (2015) Everyone	PNE	Chronic low back pain	Our ultimate goal with this	Educational book for	N / A	N / A
has back pain, USA [10]			back book is to teach you how	patient		
			pain works and provide you			
			with strategies for lessening			
			the pain. In order to do this,			
			we must address some of the			
			misinformation and myths that			
			exist about back pain.			
Lehman (2017), Recovery	PNE	Chronic pain	This booklet will give you	Educational booklet for	N / A	N / A
Strategies, Canada [66]			some knowledge about pain,	patient and practitioner		
			which, ideally, will allow you			
			to manage it better.			
Butler et Mosley (2013),	PNE	Chronic pain	1)To provide a conduit from	Educational booklet for	N / A	N / A
Explain Pain, Australia [8]	TINE	Chilonic pain	the world of basic	patient and practitioner	N/A	N/A
Explain 1 ani, Australia [0]			neuroscience to clinicians and	patient and plactitioner		
			to their patients; 2) To enable			
			people in pain to understand			
			more about their situation and			
			to become less frightened of			

Educational support (patient) (8)

			their pain; 3) To assist people			
			in pain, and those involved			
			with them, to make the best			
			choices about their			
			management; 4) To outline			
			modern models of			
			management and provide the			
			management essentials for			
			overcoming pain and returning			
			to normal life.			
Moseley et Butler (2015), The	PNE	Chronic pain	Because understanding your	Educational book for	N / A	N / A
Explain Pain Handbook :			pain is the single most	patient		
Protectometer, Australia [9]			important thing you can do to			
			start on the road to recovery.			
Laurent al (2012) With D. J.	DAID	Changi i	To be improved as a state of the	Education - 1 have been	N / A	N / A
Louw et al (2013), Why Do I	PNE	Chronic pain	It is important to know that	Educational book for	N/A	N/A
Hurt ? USA [64]			persistent pain is more due to	patient		
			the nervous system sensory			
			and how the brain processes			
			information from the body and			
			the environment. This book			
			was written to teach you how			
			the nervous system and the			
			brain processes information			
			and contributes to your painful			
			experience. The latest research			
			shows that the more you know			
			about pain and its functioning,			
			the better off you will be. This			
			include moving and moving			
			better, to feel less pain and to			
			have a growing interest and			
			ability to do more exercise and			
			movement that can benefit			
			your health. That knowledge is			
			essential for your recovery.			
Moseley (2007), Painful	PNE	Chronic pain	First, I hope you find the	Educational storybook book	N / A	N / A
Yarns: Metaphors and Stories			stories as interesting and as	for patient		
to Help Understand the			fun as I do. Second, I hope the			
Biology of Pain, Australia [68]			stories help you understand the			
			biology of pain.			

Educational support for clinician

Author, year of publication, Inte	ervention and	Study population	Objective(s)	Design	Outcome measure(s)	Important result(s)
title and country of the study c	comparator					

Louw et al (2018), Pain	PNE	Chronic pain	Our original goal for writing	Educational support for	N / A	N / A
Neuroscience Education, USA			this text was to create a single,	practitioners		
[13]			user-friendly resource for			
			clinicians and students			
			learning to apply pain			
			neuroscience education in the			
			treatment of patients with			
			chronic musculoskeletal pain			
			complaints.			
Moseley et Butler (2017),	PNE	Chronic pain	Explain Pain Supercharged is	Educational support for	N / A	N / A
Explain Pain Supercharged,			for all health professionals	practitioners		
Australia [18]			treating pain			
			and indeed anyone teaching			
			people about pain. In this			
			brand new book,			
			with entirely original content,			
			Moseley and Butler apply their			
			unique style to take the			
			neuroimmune science of pain			
			further and deeper, enriching			
			your core knowledge while			
			providing immediately			
			applicable education			
			strategies, conceptual change			
			science, curriculum			
			development and hundreds of			
			ready to use clinical metaphors			
			and therapeutic narratives.			

N / A : Not applicable.