

Nordic Association for Disability and Oral Health
 Aug. 27-29, 2015

Pain - a bio-psycho-social perspective

- with special focus on hypnosis

Robert Zachariae
 Professor, DMSc.

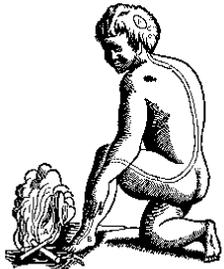
UNIT FOR PSYCHOONCOLOGY AND HEALTH PSYCHOLOGY
 DEPT. OF ONCOLOGY - AARHUS UNIVERSITY HOSPITAL - DEPT. OF PSYCHOLOGY - AARHUS UNIVERSITY

Topics

- Pain – a bio-psycho-social perspective
- Interactive dimensions of pain
- Psychosocial modulators of pain
- What is hypnosis – brief historical overview
- Modern theories of hypnosis
- Hypnotic modulation of pain
- Placebo and hypnotizability
- Pain coping and behavioral traits
- Clinical effectiveness

Classical specificity theory

The roots of the classical biomedical view on pain lie in Descartes' mechanical model, which argues for a simple, direct relationship between the sensory stimulus of a pain receptor and the behavioral response, mediated by a neural transmission mechanism



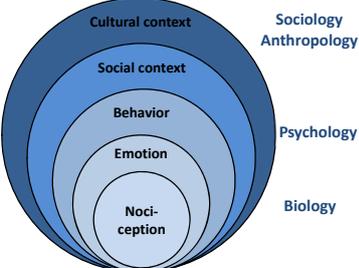
Shortcomings of the specificity theory

- Did not sufficiently explain several pain-related phenomena, e.g.
 - The association between the severity of the tissue damage and pain perception is not linear – but often weakly correlated
 - Differences in pain ratings of soldiers wounded in battle and civilians with the same level of tissue damage
 - Phantom limb pain
 - Pain in the absence of underlying organic disease or damage
 - Placebo – nocebo responses

Beecher, 1946; Melzack & Wall, 1965

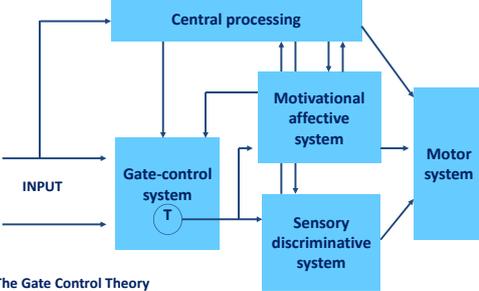
A bio-psycho-social view

The modern view recognizes the complexity of pain experience and the underlying mechanisms. Pain is regarded as a result of complex interactions between biological, psychological, and socio-cultural factors. Pain research must therefore rely on interdisciplinary approaches.



Gate control theory

Melzack & Wall 1965



The Gate Control Theory

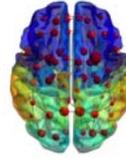
Shortcomings of the GCT

- The GCT has led to fruitful pain research
- Several details have been shown to be inaccurate
- Oversimplifications in the presentation of the neural architecture of the spinal cord and the hypothesized modulatory system
- Does not fully explain differences between various types of pain (cutaneous, visceral, muscular, neuropathic) and does not explain phantom limb pain

Melzack, 1992; Moayed & Davis, 2013

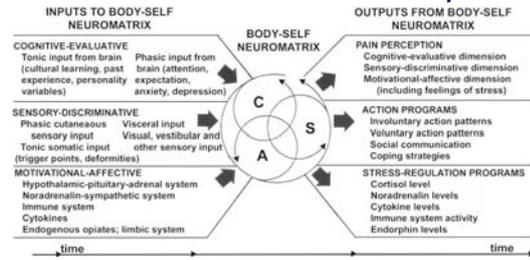
Neuromatrix theory

Melzack, 1992



- Neuromatrix is believed to be a neuronal network composing the anatomical substrate of the physical self
- The output is a **neurosignature** determining the perception of physical "self" and "non-self"
- The neuromatrix is thought to be (partly) genetically hard-wired: Individuals born without limbs may still experience phantom sensations
- Experience is thought to "add, delete, or strengthen existing synapses"
- Lack of sensory feedback from amputated limbs and/or excessive firing from damaged nerves activates/alters the output pattern from the neuromatrix – and is felt as pain
- Pain memories (as a part of the neuromatrix) prior to amputation may play a role

The neuromatrix theory



"Neuromatrix" a general concept not specific to pain – rather than a specific pain matrix, the evidence suggests a multi-modal network related to the detection of salient sensory input (which includes pain – but not pain only).

The multidimensional definition of pain (IASP)

"an unpleasant sensory and emotional experience associated with actual or potential tissue damage, or described in terms of such damage"

INTERACTIVE DIMENSIONS OF PAIN

DIMENSION	ASPECTS
SENSORY-DISCRIMINATIVE	Intensity, location, quality, duration
AFFECTIVE-MOTIVATIONAL	Unpleasantness, fight-flight response
COGNITIVE-EVALUATIVE	Appraisal, cultural norms, context, cognitive state

Melzack & Casey, 1968

McGill Pain Questionnaire

Psychological modulators of pain perception

COGNITIVE PROCESSES

- Attention
- (Dis)attention
- Dissociation
- State-dependency
- Perceived control

Psychological modulators of pain perception

EMOTIONAL PROCESSES

- Active anger
- Passive anger
- Anxiety (fear of pain)
- Anxiety (pain unrelated)
- Depression
- Happiness

Psychophysiological modulators of pain perception

AROUSAL

- Relaxation
- Tension

Psychological modulators of pain perception

“PLACEBO”

- Conditioning?
- Expectation?
- Response-appropriate sensation?
- Anxiety-reduction?
- “Non-specific” inter-personal factors?
- Imagery

Psychological modulators of pain perception

PAIN COPING STRATEGIES

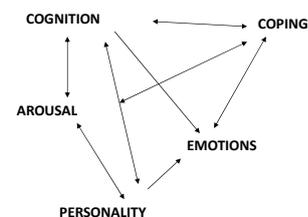
- Catastrophizing
- Re-interpretation
- Distraction
- Perceived control

Psychological modulators of pain perception

PERSONALITY TRAITS

- Locus of control
- Emotional repression
- Augmenters/reducers
- Social desirability
- Neuroticism
- Hypnotizability

interactions



EPOS

Hypnotic modulation of pain



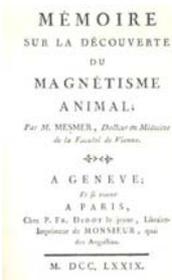
ZACHARIAE

What is hypnosis?



ZACHARIAE

Historical debates



Animal magnetism (Mesmer)

Or

Imaginative involvement (Royal Commission)

ZACHARIAE

Historical debates



Psychopathological (Hysteria) (Charcot)

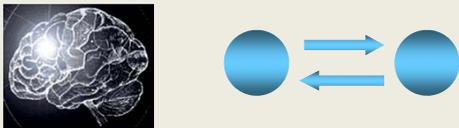
or

Normal phenomenon (Bernheim)

ZACHARIAE

Historical debates

- State (Hilgard)
 - Psychophysiological correlates
 - High hypnotizable vs. simulator-designs
 - Etc.
- Role-taking (Spanos)
 - Expectation
 - Order of presentation designs
 - Etc.



ZACHARIAE

What is hypnosis?

Hypnotic trance is a phenomenon involving **attention** and **concentration**, which may be induced by several types of events.

Hypnosis

- Religious ceremonies
- Traumatic events
- Intense involvement

➔

HYPNOTIC TRANCE

- Absorption
- Dissociation
- Suggestibility

ZACHARIAE

Principles in hypnotic communication

- Rapport
- Motivation
- Attention
- Repetition
- Positive suggestions
- Creating a "yes-set"
- Embedding
- Truisms
- Handling non-responses
- Implied directive
- Confusion
- Double bind
- Metaphors
- Direct vs indirect sugg.
- Timing and pacing
- Reversible effect
- Utilization
- Dissociative suggestions

The phenomenology of hypnosis

How do we know when someone is hypnotized?



- Response to ideomotor suggestions
- Perceived involuntariness
- Response-inhibition
- Changes in cognition (time-distortion etc.)
- Physiologic/behavioral correlates (eye-movements, catalepsy)
- Hallucinations (positive & negative)
- Posthypnotic amnesia
- Response to post-hypnotic suggestions
- Hypnotic analgesia
- High hypnotizable vs. simulator design

Everyday forms of hypnosis



Modern theories on hypnosis

- Dissociation theory of hypnosis (Hilgard)
 - The control of behavior is normal
 - The individual's experience – monitoring – of the behavior is impaired
- Neo-dissociation theory (Bowers)
 - Dissociated control – the underlying control of behavior is changed (executive function)

Modern theories on hypnosis

- Non-state social-psychological theory of hypnosis (TX Barber, Spanos)
 - Views hypnosis as role-taking – not as an altered dissociated state
 - Behaviors and experiences associated with hypnosis are acted out by the hypnotized individual in accordance with the social context and expectations of the hypnotic setting
 - People control their hypnotic experience by acting how they believe they are supposed to act during hypnosis
 - c.f. Martin Orne's real-simulator experiment

Modern theories on hypnosis

- Interactionist theory (Sheehan, McConkey)
 - Hypnosis depends on the qualities the individual brings to the hypnotic setting, and the ability of the hypnotist to establish the conditions that favor the persons ability to create and believe the suggested experiences
 - The hypnotic subjects are motivated and actively involved in hypnotic suggestions to develop a strong commitment to the communications of the hypnotist
 - Although they are actively involved, they still experience their response as outside their control and subjectively real

Modern theories on hypnosis

- Cold-control theory (Dienes & Perner)
 - Take offset in theories of cognition:
 - First-order and second-order states
 - Consciousness depends on our being conscious of our consciousness – e.g. I am conscious of my mental state
 - First order: e.g. imagining a cat or intending to lift my arm
 - Second order: I am aware of imagining or my intention
 - Example: blind-sight patients lack awareness of being in the mental state of seeing
 - The hypnotic experience is not compromised executive function but being without higher order consciousness of the behavior

Modern theories on hypnosis

- Discrepancy-attribution theory (Barnier & Mitchell)
 - The hypnotic experience is due to discrepancy between the cognitive process and its attribution
 - Normal: I am imagining a cat, and the resulting imagined image of a cat is attributed to that: imagination
 - Hypnosis: I am imagining a cat, but the resulting image is not attributed to the process of imagining
 - Attribution to imagination is related to the effort associated with imagination
 - The shift to a different attribution is due to the slight changes in the ease of producing images in hypnosis

Neuropsychophysiology of hypnosis

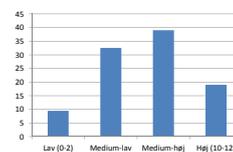
- Hypnotizability a reliable and stable personality trait measurable by tests e.g. SSHS and HGSHS
- A heritability component in hypnotizability
- Correlations between hypnotizability and absorption and ability to partition attention
- EEG-differences between highs and lows – e.g. highs generate more high theta-power



Morgan, 1973; Crawford, 1990; Lichtenberg et al. 2000; Zachariae et al. 2000

Hypnotic susceptibility

- Measured with standardized tests, e.g., The Harvard Group Scale of Hypnotic Susceptibility
- Normally distributed
- Stable over time (e.g., 25 year test-retest)



Measuring hypnotizability

Distribution of Hypnotizability
After Hilgard (1965)



And "virtuoso" hypnotizables

Neuropsychophysiology of hypnosis

- Highs show greater hemispheric specificity/ asymmetry than lows
- Highs have higher preference for right-hemispheric activity
- Hypnosis associated with increased cerebral metabolism in certain brain regions
- Hypnotizability associated with increased frontal lobe activity and highs have more effective and flexible attentional and inhibitory (dis-attentional) systems
- Highs more able to suppress or enhance event-related brain potentials

Crawford, 1989; Gruzelier, 1999; De Benedittis & Longostreui, 1988; Zachariae & Bjerring, 1994

Neurobiology of hypnosis

• Stronger associations between areas in the brain related to the ability of focused attention in high than low hypnotizable individuals.

Hoefl et al. 2012.

Neurobiology of hypnosis

Hypnotisk "lammelse"

Cojan et al. 2009

Neurobiology of hypnosis

Greater blood flow in areas associated with color perception when instructed to perceive grey as color, but only in hypnotic condition

Kosslyn et al. 2000

Neurobiology of hypnosis

- Taken together, the evidence suggests
 - That hypnosis is supported by neural activity that allows for
 - highly focused attention
 - Ability to manipulate sensory input
 - Reduction in self-consciousness
 - Involves intense cognitive activity that is experienced as effortless
 - May be a developed ability to alter attention, inhibit arousal, reduce conflict monitoring in response to threats (stress management)

Spiegel, 2008

The Domain of Hypnosis

Hilgard, 1973

An altered state of consciousness

- Altered perception
- Altered attention
- Dissociation
- Trance
- Memory

An interpersonal relationship

- Hypnotic role-taking
- Suggestive influence
- Therapeutic alliance

Hypnosis

A (neuro)physiological phenomenon

- Altered cerebral blood flow
- Altered EEG-patterns
- Altered reflexes

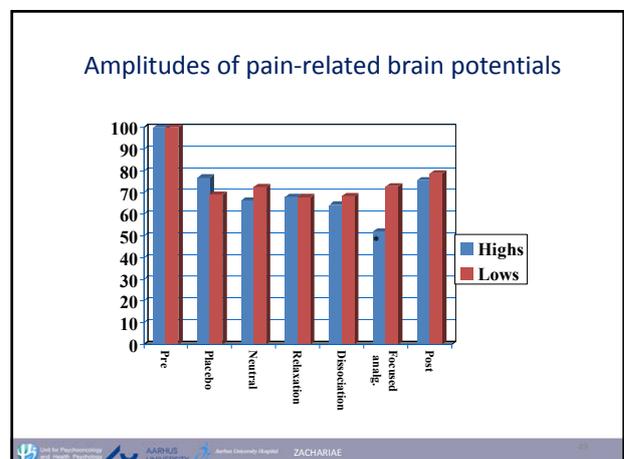
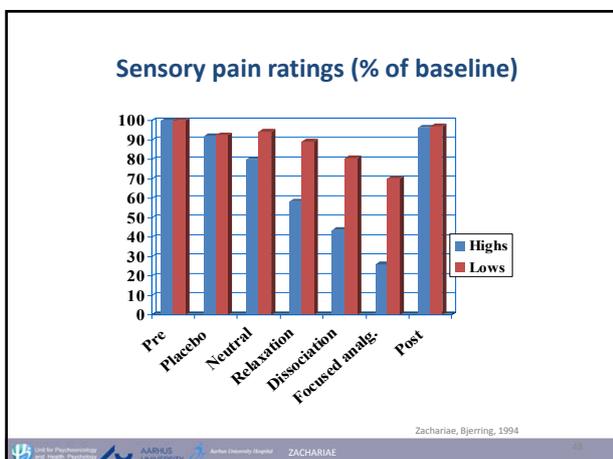
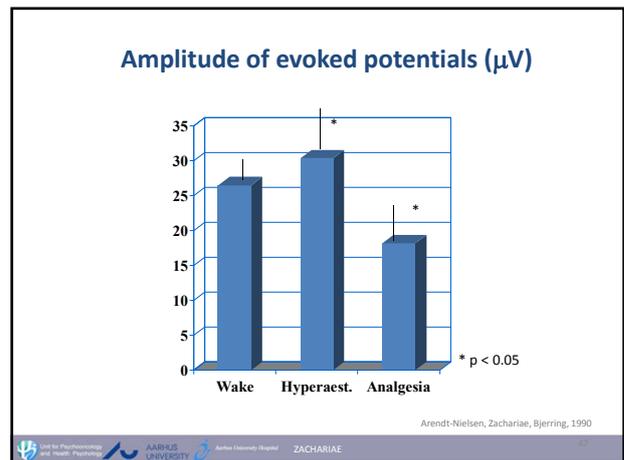
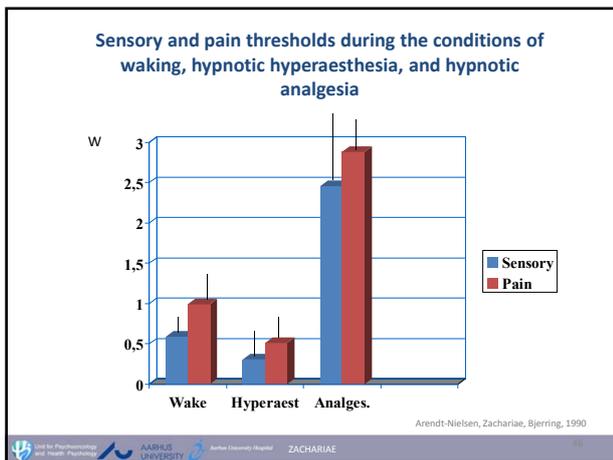
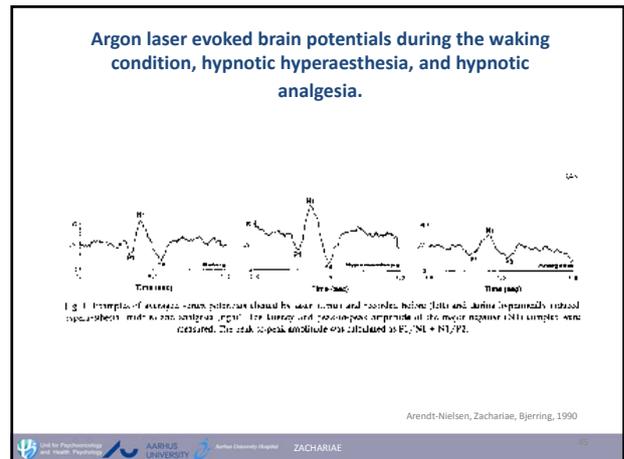
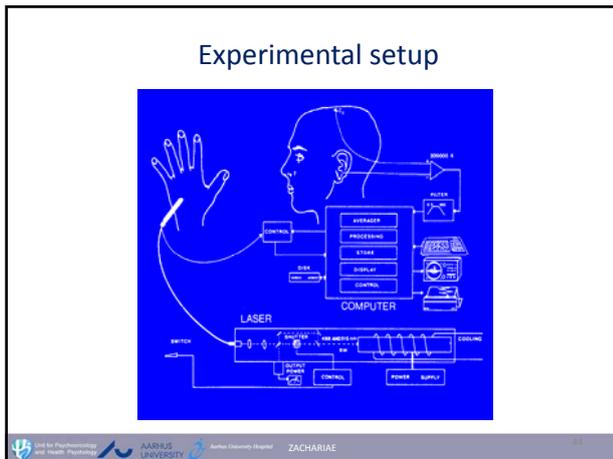
Behavioral response

- Increased suggestibility
- Altered motivation
- Altered expectations
- Perceived involuntarism

ZACHARIAE

Hypnotic modulation of sensory pain perception

ZACHARIAE



Hypnotic analgesia

- Patients with temporomandibular disorder

Chiropr J Volume 27, Number 4, May 2011

FIGURE 3. Effect of treatment with hypnosis or control on MRI scores of the characteristic pain intensity (mean \pm SEM values) in patients with temporomandibular disorder. Each row was scored 1 times daily 7 days before treatment (black bars) and after treatment (white bars). * indicates significant lower MRI scores in the hypnotic group after treatment (P < 0.001). NBS indicates numbers of rating scale.

Abrahamsen, Baad-Hansen, Zachariae, Svensson, 2011
Abrahamsen, Dietz, Lodahl, Roepdorff, Zachariae, Østergaard, Svensson, 2010

Nociceptive reflexes (%)

Significant effects of both condition ($p < .01$) and hypnotic susceptibility. ($p < .05$)

Zachariae, et al. 1998

Percent change of pain intensity and reflexes during conditions of analgesia with naloxone and with saline compared to analgesia without injection

* $p < .05$

Hypnotic analgesia, pain-related brain potentials, and histamine flare reaction during hypnotic analgesia

- Lesions of the sensory nerves are associated with reduced histamine flare reaction and unilateral remission of psoriasis (Lynn, 1988; Farber, 1990)
- Local analgesics associated with reduced histamine flare reaction (Bjerring & Arendt-Nielsen, 1990)
- Ten high hypnotizable subjects had argon laser induced pain-related brain potentials and histamine flare reactions measured during:
 - A pre-hypnotic baseline
 - During suggestions to experience hypnotic analgesia
 - After hypnosis

Pain-related brain potentials

Ratio between histamine flare reactions in the analgesic and the control arm

100 = 1:1

Hypnotically induced inflammation

- Hypnotists claim that they are able to induce inflammatory skin reactions by asking patients to imagine that they have been burned by a cigarette (Ullman, 1947)
- In a study, nurses were given suggestions in hypnosis that they were burned by boiling fat while frying bacon. One woman responded and told that she had once had a similar experience (Johnson & Barber, 1976)






 ZACHARIAE

Hypnotically induced inflammation



En tv-producer (LS-O), der tidligere havde haft en forbrænding på hånden, fik fremkaldt en rødmen i den samme hånd i hypnose ved at genkalde sig situationen (Zachariae, DR2, 2013)




 ZACHARIAE



EEG




 ZACHARIAE






 ZACHARIAE

Emotions and pain

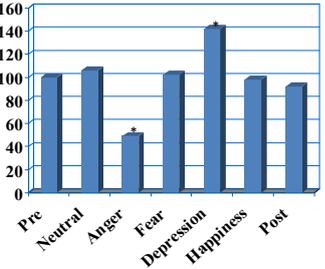







 ZACHARIAE

Pain-evoked brain potentials during hypnotically induced emotions



Emotion	Brain Potential (%)
Pre	100
Neutral	105
Anger	55*
Fear	105
Depression	145
Happiness	100
Post	95

*) p < .05

Zachariae, et al. 1991




 ZACHARIAE

EPOS

Placebo and hypnosis



62

Placebo

- Proposed placebo mechanisms
 - Conditioning
 - Expectancy (e.g., “priming”)
 - “Response-appropriate sensation”
 - Reduction of anxiety/tension
 - “Non-specific” inter-personal factors

63

Study name	Outcome	Statistics for each study			Correlation and 95% CI	
		Lower limit	Upper limit	Z-Value p-Value		
Cassileth et al. 1985 +	Occurrence	0.070	-0.270	0.394	0.397	0.692
Jacobsen et al. 1988	Combined	0.245	-0.052	0.502	1.621	0.105
Haut et al. 1991	Combined	0.531	0.245	0.732	3.400	0.001
Andrykowski & Gregg 1992	Combined	0.151	-0.097	0.380	1.195	0.232
Rhodes et al. 1995 +	Occurrence	0.169	0.062	0.272	3.081	0.002
Watson et al. 1998	Occurrence	0.070	-0.147	0.260	0.631	0.528
Montgomery & Bovbjerg 2000	Occurrence	0.303	0.033	0.532	2.190	0.029
Roscoe et al. 2000a	Severity	0.424	0.068	0.684	2.308	0.021
Roscoe et al. 2000b	Severity	0.224	0.006	0.422	2.012	0.044
Mississotte et al. 2002	Combined	0.050	-0.195	0.260	0.415	0.678
Roscoe et al. 2004	Severity	0.067	-0.075	0.206	0.927	0.354
Oliver et al. 2005 +	Severity	0.224	0.017	0.413	2.113	0.035
Zachariae et al. 2007	Severity	0.150	-0.050	0.338	1.473	0.141
Higgins et al. 2007	Severity	0.147	-0.021	0.365	1.075	0.281
Booth et al. 2007	Occurrence	0.270	0.111	0.416	3.276	0.001
Colaguri et al. 2008	Combined	0.141	0.066	0.214	3.669	0.000
Shelke et al. 2008	Combined	0.237	0.131	0.338	4.315	0.000
		0.175	0.138	0.214	6.595	0.000

Colaguri & Zachariae: Patient Expectancy and post-chemotherapy nausea – a meta-analysis 2010

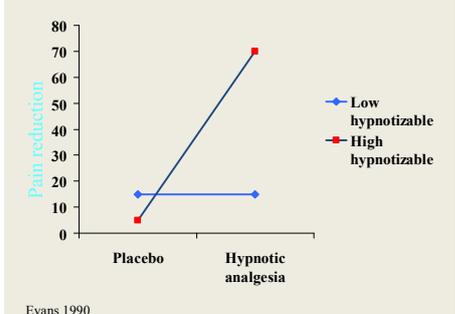
64

Hypnosis and placebo

- Is the placebo response related to suggestibility?
- Are high hypnotizables better placebo-responders?

65

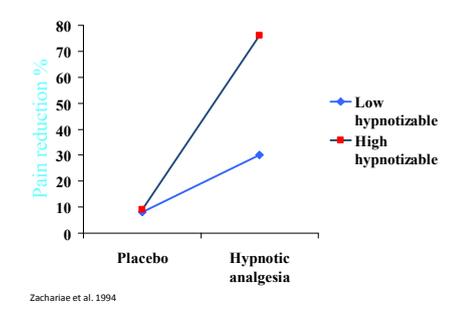
Hypnosis and placebo



Evans 1990

66

Hypnosis and placebo



Zachariae et al. 1994

67

EPOS

Pain coping and behavioral traits

10 9 8 7 6 5 4 3 2 1 0

100

Coping Strategies questionnaire

Rosenstiehl & Keefe, 1983; Swartzman et al. 1994

- **Distraction**
 - I think about pleasant previous experiences
- **Ignoring sensations**
 - I tell myself that my pain must not stop me from doing what I need to do
- **Reinterpretation**
 - I pretend that the pain is not a part of me
- **Catastrophizing**
 - I feel that I cannot cope any longer
- **Prayer and hope**
 - I put my trust in God

100

Pain catastrophizing

When I'm in pain ...

- I worry all the time about whether the pain will end.
- I feel I can't go on.
- It's terrible and I think it's never going to get any better.
- It's awful and I feel that it overwhelms me.
- I feel I can't stand it anymore.
- I become afraid that the pain will get worse.
- I keep thinking of other painful events.
- I anxiously want the pain to go away.
- I can't seem to keep it out of my mind.
- I keep thinking about how much it hurts.
- I keep thinking about how badly I want the pain to stop.
- There's nothing I can do to reduce the intensity of the pain.
- I wonder whether something serious may happen.

Dimensions:
Helplessness
Rumination
Intrusive thoughts

100

Pain catastrophizing

- Pain catastrophizing associated with pain intensity and unpleasantness in clinical (chronic tension-type headache) and healthy samples (students) – even when adjusting for depression and trait-anxiety
- Hypnotic suggestions for high and low pain catastrophizing associated with changes in clinical (chronic tension-type headache) and experimentally induced pain (intramuscular infusion of hypertonic saline)

Kjægg H, Zachariae R, Pheiffer-Jensen M, Kasch H, Svensson P, Jensen TS, Vase L. Pain frequency moderates the relationship between pain catastrophizing and pain. *Frontiers in Psychology*, 2014.

Kjægg H, Kasch H, Zachariae R, Svensson P, Jensen TS, Vase L. Experimental manipulations of pain catastrophizing influence pain levels in chronic pain patients and healthy volunteers (submitted manuscript)

100

Behavioral traits

- Locus of control
- Alexithymia
- Emotional repression
- Augmenting/reducing
- Social desirability
- Neuroticism (negative affectivity)
- Somato-sensory amplification
- Monitoring/blunting
- Hypnotizability

100

Somato-sensory amplification

Somatosensory Amplification Scale

1. When someone else coughs, it makes me cough too
2. I can't stand smoke, smog, or pollutants in the air
3. I am often aware of various things happening within my body
4. When I bruise myself, it stays noticeable for a long time
5. Sudden loud noises really bother me
6. I can sometimes hear my pulse or my heartbeat throbbing in my ear
7. I hate to be too hot or too cold
8. I am quick to sense the hunger contractions in my stomach
9. Even something minor, like an insect bite or a splinter, really bothers me
10. I have a low tolerance for pain

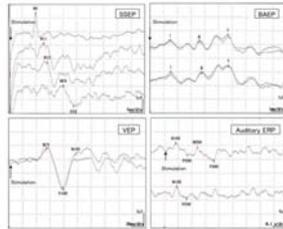
Total

Barsky, 1992

100

Somato-sensory amplification

- Studies suggests that individuals high on SSA process somatosensory stimuli differently than individuals low in SSA



Nakao & Barsky, 2007

Coping with threatening health-related information – two strategies

- "MONITORING"
 - High degree of attention towards bodily sensations and stimuli from the environment (hypervigilance)
 - High level of need for perceived control
 - High level of need for detailed information
- "BLUNTING"
 - Perceives sensations and stimuli as potentially threatening
 - Copes by suppressing unpleasant thoughts and emotions
 - Have limited needs for information about diagnosis and treatment

Miller, 1995

Coping with threatening health-related information – two strategies

- Example
 - In a stratified, randomized study, patients received either limited or extensive information prior to a surgical procedure.
 - Monitors, who received limited information, and blunTERS, who received extensive information, reported more post-surgical pain and experienced more complications than the remaining patients.

Miller, 1995

Take-home message



12

- Pain is a **multidimensional** phenomenon including sensory, cognitive, and affective processes
- Pain is influenced by **coping**, e.g. pain catastrophizing
- Individual differences in pain responses may be related to **behavioral traits**, e.g., somato-sensory amplification, hypnotizability
- Self-reported pain perception and neurobiological pain correlates can be modulated by communicative **intervention** strategies, e.g., placebo, hypnosis
- Pain is a **bio-psycho-social** phenomenon and clinicians should focus on all dimensions when treating pain

Clinical effectiveness of hypnosis

Both volunteers and patients experience significant pain relief from hypnotic analgesia

Significant effect on effect size (Cohen's d) of hypnotizability:

- High = 1.22
 - Medium = 0.64
 - Low = 0.10
- Effect size conventions (Cohen 1988):
- Large = 0.8
 - Medium = 0.5
 - Small = 0.2

Study	Participants	Type of Pain	n	d
Wakeman et al. (1978)	patients	burn	24	13.41
Wakeman et al. (1979)	patients	burn	18	17.42
Cassidy & Wood (1979)	students	cold pressure	20	-0.01
Cassidy & Wood (1979)	students	cold pressure	20	1.50
Cassidy & Wood (1979)	students	cold pressure	20	-0.45
Spencer et al. (1980)	students	cold pressure	20	0.12
Spencer et al. (1984)	students	cold pressure	45	0.47
Spencer et al. (1985)	students	cold pressure	42	0.94
Trigg & Marks (1986)	students	cold pressure	28	0.88
Stasi et al. (1987)	students	ischemic	45	0.12
Ellen et al. (1988)	students	ischemic	137	0.70
Edelson et al. (1988)	patients	chronic pain	14	0.94
Spencer et al. (1988)	students	focal pressure	96	0.29
Spencer et al. (1988)	students	focal pressure	60	0.37
Zohar et al. (1988)	students	cold pressure	37	1.19
Spencer & Katsanis (1989)	students	focal pressure	20	1.35
Spencer & Katsanis (1989)	students	focal pressure	20	1.81
Spencer et al. (1990)	students	focal pressure	30	1.67
Wakeman & Liu (1991)	patients	ostomy	32	0.65
Syjala et al. (1992)	patients	cancer	22	0.10
Falzon et al. (1992)	patients	burn	20	0.72
Baker et al. (1993)	students	cold pressure	20	0.30
Baker et al. (1993)	students	cold pressure	20	0.41
ter Kuile et al. (1994)	community	headache	29	-0.43
ter Kuile et al. (1994)	patients	headache	36	0.34
ter Kuile et al. (1994)	students	headache	26	0.42
Liang et al. (1996)	patients	radiological procedure	30	1.01

Note: Ischemic, cold pressure, and focal pressure are experimental pain stimuli.

Montgomery et al. 2000

