

# **Novel developments in assessment of Neuropathic pain and Breakthrough pain**

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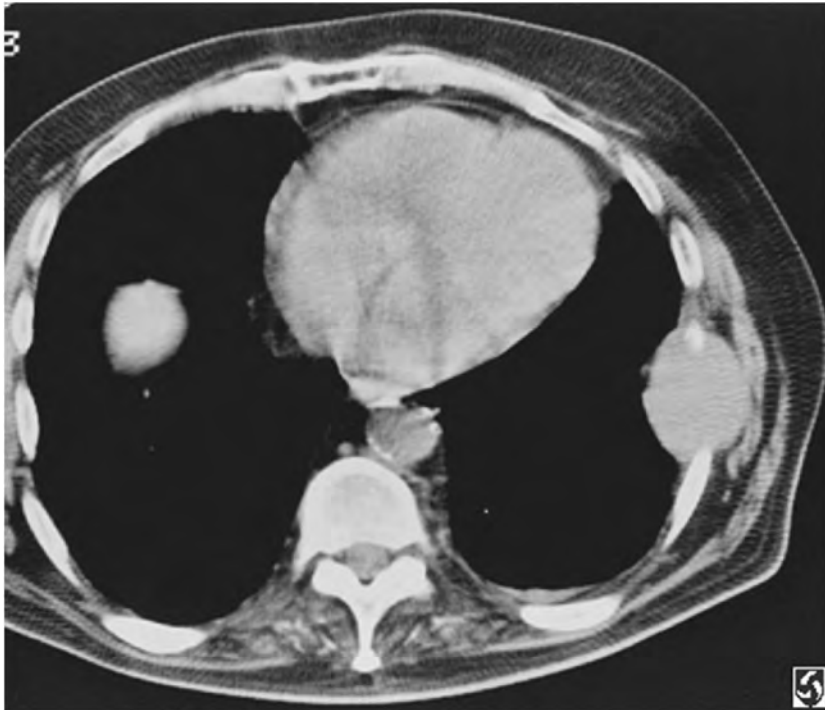
University of Leeds, UK

- **Neuropathic pain**
  - Current practice lacks precision
  - New guidance is promising but needs testing
  - I'll tell you the details
- **Breakthrough pain**
  - Frankly, it's a mess
  - Some serious work needed to simplify this area
  - I'll spare you the details



# Neuropathic pain in cancer patients

- **Standard neuropathic cancer pain**
- **New types of neuropathic pain in cancer:**



- In developed countries, increasingly larger population of older people with cancer





# Aetiology

- Neuropathic cancer pain?
  - directly caused by cancer
- ....or neuropathic pain in a cancer patient?
  - treatment neuropathies
  - co-morbid conditions
- Need to be clear for epidemiological, clinical and research purposes



“Is my patient’s  
cancer pain  
predominantly  
neuropathic or  
nociceptive in  
origin?”





IASP®

PAIN® 153 (2012) 359–365

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[www.elsevier.com/locate/pain](http://www.elsevier.com/locate/pain)

## Prevalence and aetiology of neuropathic pain in cancer patients: A systematic review

Michael I. Bennett<sup>a,\*</sup>, Clare Rayment<sup>b</sup>, Marianne Hjermstad<sup>c,d</sup>, Nina Aass<sup>e,f</sup>, Augusto Caraceni<sup>g</sup>,  
Stein Kaasa<sup>h</sup>

- 22 studies, 13,600 patients
- Pain type diagnosed by clinical judgement
- Estimated ‘conservative’ and ‘liberal’ prevalence



- Cancer patients have 2 distinct pains, on average:
- 20% of pains in cancer patients are neuropathic in origin
  - 18.7% - 21.4%
- Up to 40% of cancer patients are affected by neuropathic pain
  - 19% - 39.1%

- Take 5 cancer patients.....



- Aetiology of pain in cancer patients.....

	All cancer pain*	Neuropathic pain only**
Direct effect of cancer	76%	64%
Cancer treatment	11%	20%
Indirect effects	5%	4%
Co-morbid conditions	8%	12%

\* Grond et al, 1996

\*\*Bennett et al 2011

- Aetiology of pain in cancer patients.....

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\* Grond et al, 1996

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## **Neuropathic cancer pain: Prevalence, severity, analgesics and impact from the European Palliative Care Research Collaborative–Computerised Symptom Assessment study**

Clare Rayment, Marianne J Hjermstad, Nina Aass, Stein Kaasa, Augusto Caraceni, Florian Strasser, Ellen Heitzer, Robin Fainsinger, Michael I Bennett

*Palliative Medicine*

0(0) 1–8

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 SAGE

1051 patients assessed in 17 European centres



# Epidemiology

Compared to nociceptive cancer pain, NeuP patients had:

- More oncological treatment
- More opioids
- More adjuvant analgesia
- Poorer QOL, reduced performance status
- No overall differences in pain intensity

*Fainsinger et al 2010, Rayment et al 2011*



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**ORIGINAL ARTICLE**

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Diagnosing Neuropathic Pain in Patients with  
Cancer: Comparative Analysis of  
Recommendations in National Guidelines from  
European Countries

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Virginie Piano, MD<sup>\*,†</sup>; Stans Verhagen, PhD<sup>†,‡</sup>; Annelies Schalkwijk, MSc<sup>†</sup>;  
Jako Burgers, PhD<sup>§</sup>; Hans Kress, MD, PhD<sup>¶</sup>; Rolf-Detlef Treede, PhD<sup>||</sup>; Yechiel Hekster,  
PhD<sup>\*\*</sup>; Michel Lanteri-Minet, MD<sup>\*</sup>; Yvonne Engels, PhD<sup>†</sup>; Kris Vissers, MD, PhD<sup>†</sup>

Pain Practice 2013, 13(6):433-439

- 9 National guidelines identified
  - Italy, UK, France, Netherlands, Spain, Norway

## History

- Screening tools (n=2),
- Assessment of aetiology (n=8)
- Pain drawings (n=3)

## Examination

- Sensory abnormalities (n=4), QST (n=2)
- Diagnostic test (n=0)

Poor response to opioids (n=4)





- History:
  1. pain in a neuro-anatomically plausible distribution
  2. relevant lesion or disease
- Examination:
  3. abnormal function
    - bedside examination: numbness, allodynia, hyperalgesia
  4. abnormal structure
    - MRI or ENMG demonstrating site of the nerve lesion



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## Prevalence and aetiology of neuropathic pain in cancer patients: A systematic review

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Topical review

## How is neuropathic cancer pain assessed in randomised controlled trials?

Geana Paula Kurita<sup>a,b,c,\*</sup>, Angelika Ulrich<sup>d</sup>, Troels Staehelin Jensen<sup>e</sup>, Mads Utke Werner<sup>b</sup>, Per Sjøgren<sup>a</sup>



# Current practice

- **Summary of 31 studies (n= 13,951)**
- Met both history criteria
  - Distribution plus lesion= 18 / 31
- Met at least one examination criteria
  - Abnormal function = 20 / 31
  - Abnormal structure = 8 / 31
- Reached at least probable NP
  - 15 of 31 studies



# Current practice

- What is the effect of more rigorous assessment on prevalence estimates?
  - In 14 non-rigorous studies:
    - 20% (pure) to 39.1% (mixed)
  - In 8 rigorous studies:
    - 13.2% (pure) to 35.8% (mixed),  $p < 0.001$



# Confirming neuropathic pain in cancer patients: applying the NeuPSIG grading system in clinical practice and clinical research

- Matthew R. Mulvey
- Roman Rolke
- Pål Klepstad
- Augusto Caraceni
- Marie Fallon
- Lesley Colvin
- Barry Laird
- Michael I. Bennett

**PAIN 2014 (in press)**

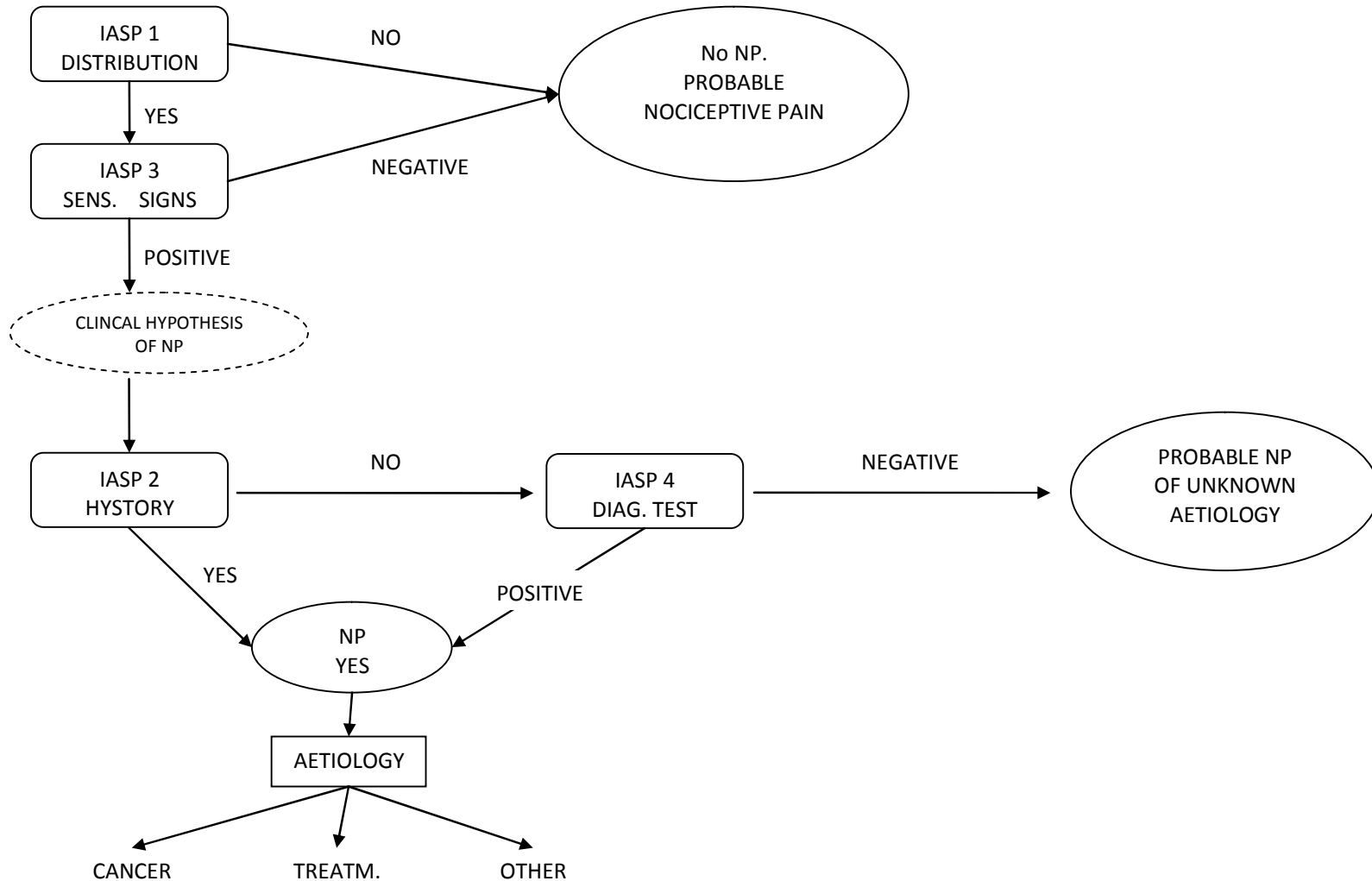


- Applying the grading system in clinical practice
  - The grading system is simple to adapt for clinical practice
- Some test serve a dual purpose, for example:
  - Delineating area of pinprick hyperalgesia determines the distribution of pain (criterion 1) and confirms sensory abnormalities (criterion 3)
  - CT scan showing tumour growth compressing a nervous structure demonstrates a history of a relevant lesion (criterion 2) and provides an additional objective confirmatory test (criterion 4).

# EAPC RN Delphi survey



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## Is there a role for verbal description or screening tools in the assessment process?

*Can the LANSS scale be used to classify pain in chronic cancer pain trials?*

*Support Care Cancer. 2013 Dec;21(12):3387-91*

*Hardy J, Quinn S, Fazekas B, Agar M, Currow D.*

When the clinical assessment was compared with the LANSS scale, the overall accuracy was 94 % (79/84).

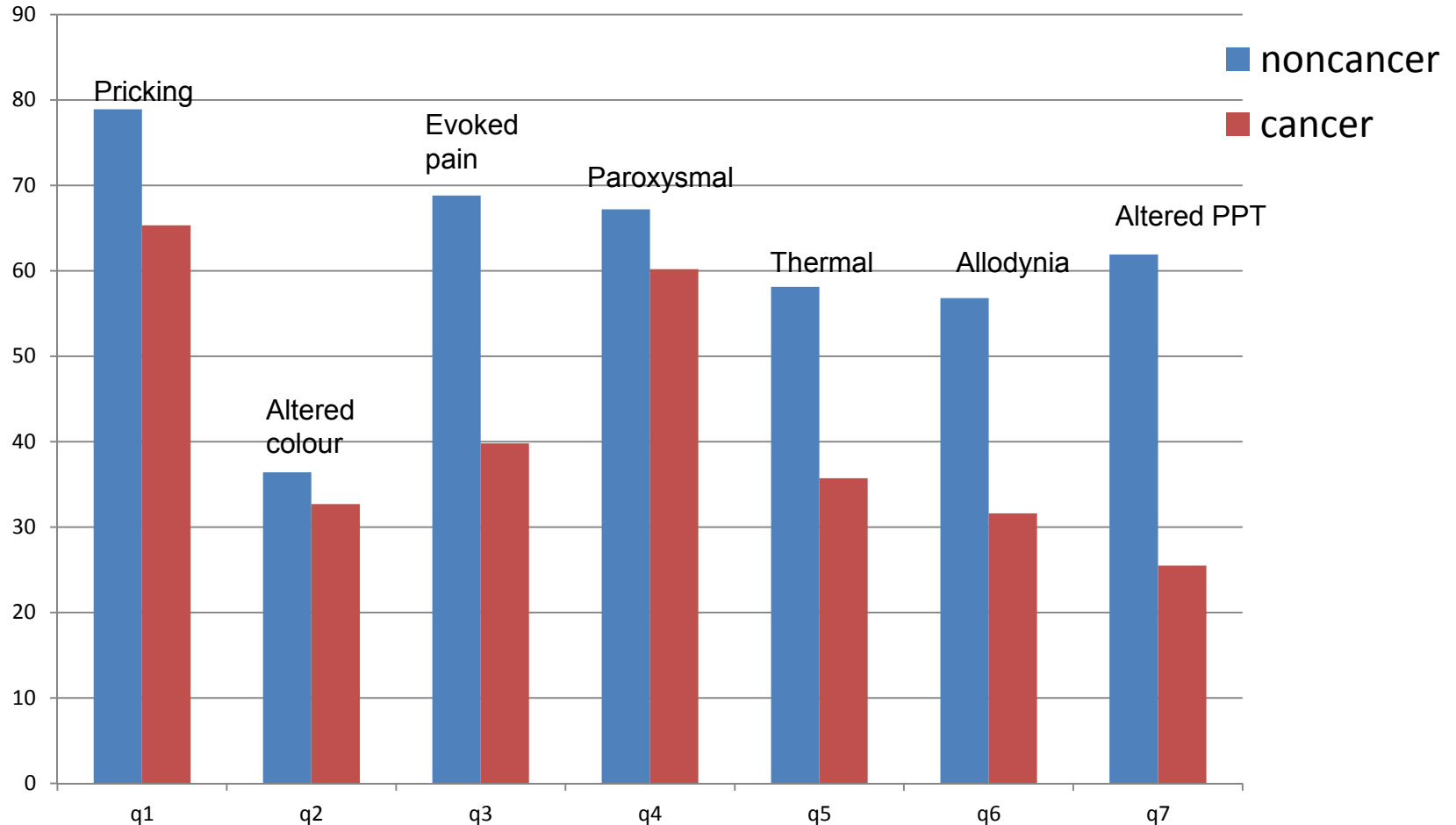


# Non-cancer NP v Cancer NP



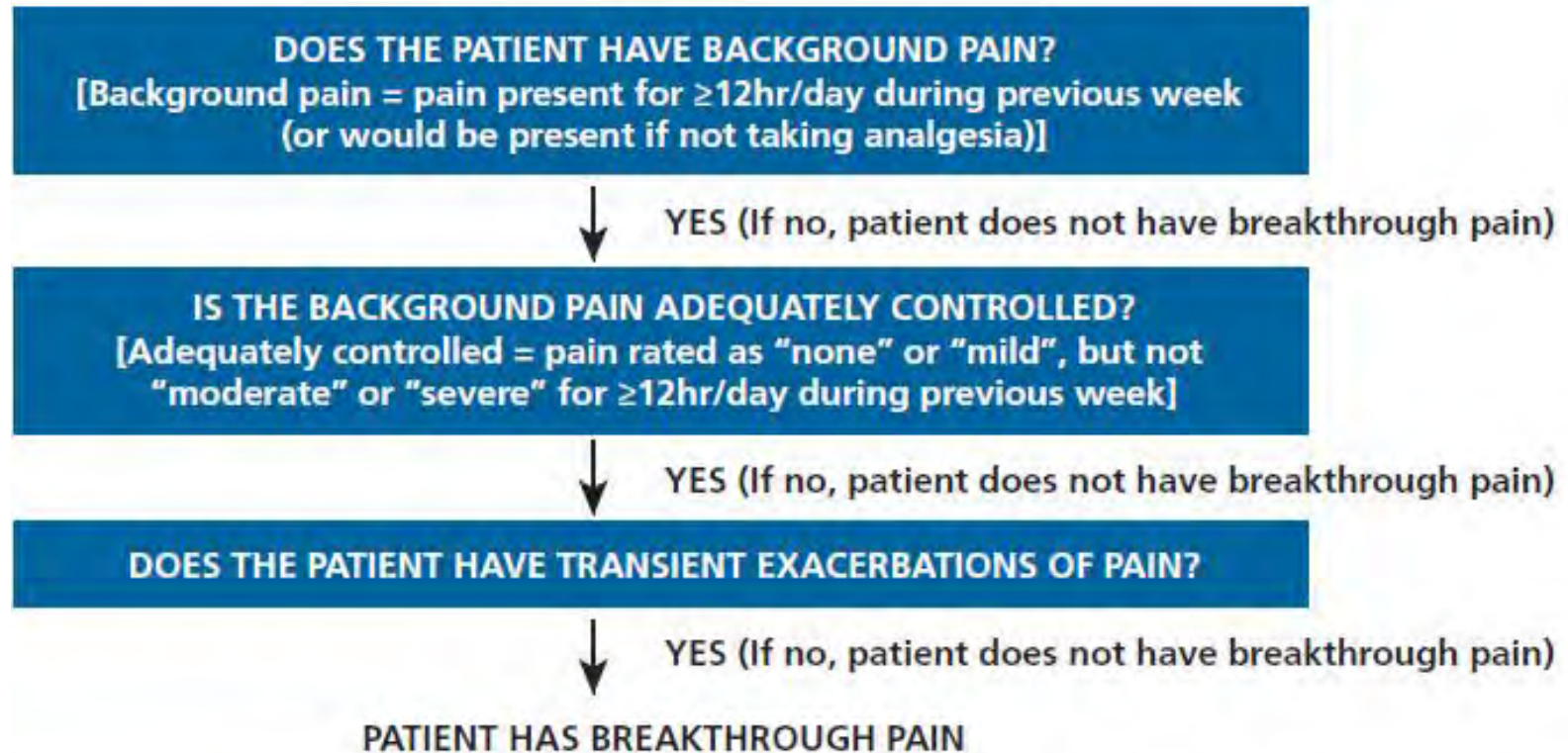
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## LANSS item frequency





# Breakthrough pain



**Figure 1** Algorithm for diagnosing patients with breakthrough pain. Reprinted from Davies AN, Dickman A, Reid C, et al. The management of cancer-related breakthrough pain: recommendations of a task group of the Science Committee of the Association for Palliative Medicine of Great Britain and Ireland. *Eur J Pain* 2009;13:331–338, with permission from Elsevier.



## Assessment and classification of cancer breakthrough pain: A systematic literature review

Dagny Faksvåg Haugen<sup>a,b,\*</sup>, Marianne Jensen Hjermsstad<sup>a,c</sup>, Neil Hagen<sup>d</sup>, Augusto Caraceni<sup>e</sup>, Stein Kaasa<sup>a,f</sup>,  
On behalf of the European Palliative Care Research Collaborative (EPCRC)



**Table 1**

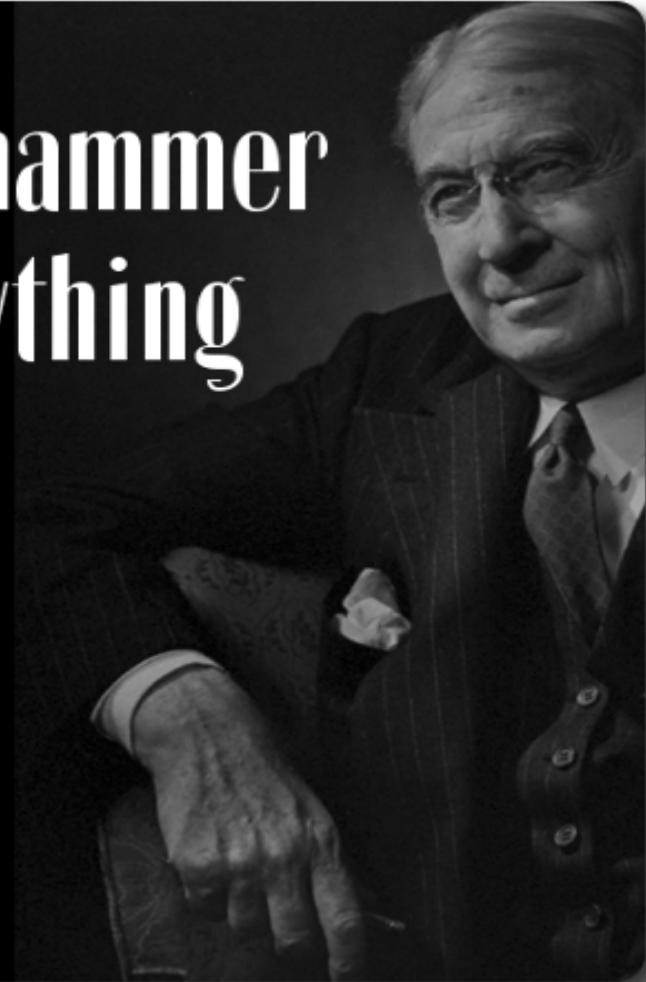
Classification of breakthrough pain: areas of high and low degree of consensus.

Area	Summary of consensus points
Term to be used	Breakthrough pain
Etiology	Caused directly by the cancer Caused indirectly by the cancer Unrelated to the cancer disease; caused by concurrent illness, or unknown/uncertain
Pathophysiological mechanism	Somatic  Visceral Neuropathic Mixed
Type or subtype	Incident pain (precipitated) Volitional (predictable) Non-volitional (predictable or unpredictable) Spontaneous/idiopathic pain (stimulus-independent, unpredictable) End-of-dose failure
Area	Main points of disagreement
Definition	Is opioid treatment a prerequisite for diagnosing BTP? Is controlled baseline pain a prerequisite for diagnosing BTP, and how should controlled baseline pain be defined? Should end-of-dose failure be included in BTP?
Formal classification system	(No formal classification system exists)



**If all you have is a hammer  
in the toolbox, everything  
looks like a nail.”**

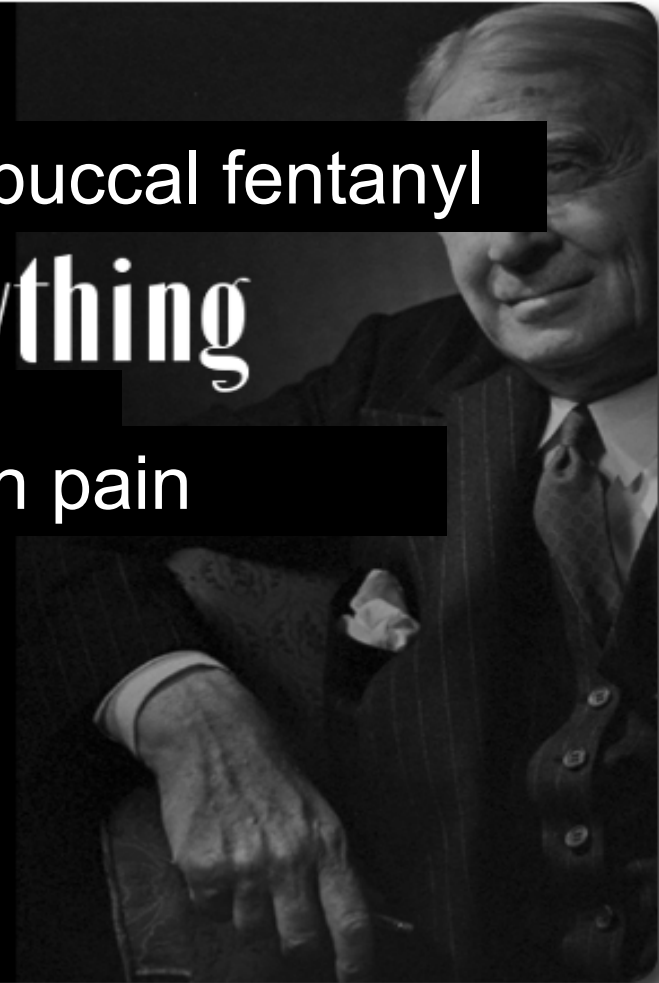
**- Bernard Baruch**





If all you have is a buccal fentanyl  
in the toolbox, everything  
looks like breakthrough pain

- Bernard Baruch





- EAPC RN project on breakthrough pain classification and assessment
  - More news next year!





**Thank you**

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