

# Anxiety and depression. Significance, assessment and treatment

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2021

KOMPETANSESENTER  
I LINDRENDE BEHANDLING  
MIDT-NORGE

# Adjustment

# Adjustment

# Anxiety

# Anxiety

- A category:

No anxiety

Anxiety

- A continuum:



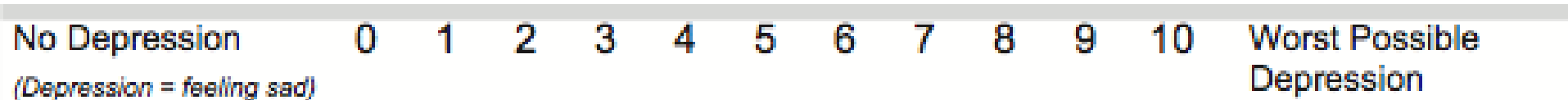
# Depression

- A category:

No depression

Depression

- A continuum:



# Anxiety and depression

One symptom

Sets of symptoms

No Depression <i>(Depression = feeling sad)</i>	0	1	2	3	4	5	6	7	8	9	10	Worst Possible Depression
No Anxiety <i>(Anxiety = feeling nervous)</i>	0	1	2	3	4	5	6	7	8	9	10	Worst Possible Anxiety

# ICD-11

International Classification of Diseases for  
Mortality and Morbidity Statistics

Eleventh Revision



World Health  
Organization

## DIAGNOSTIC AND STATISTICAL MANUAL OF MENTAL DISORDERS

FIFTH EDITION

# DSM-5

AMERICAN PSYCHIATRIC ASSOCIATION



# ICD-11. Sets of symptoms

## **Disorders specifically associated with stress**

### **Adjustment disorder**

A reaction to a stressor: e.g. illness

### **Anxiety in stress- and trauma (Traumatic stress)**

Acute stress reaction (A normal response to the stressor)

Post Traumatic Stress Disorder (Context: After extreme threat)

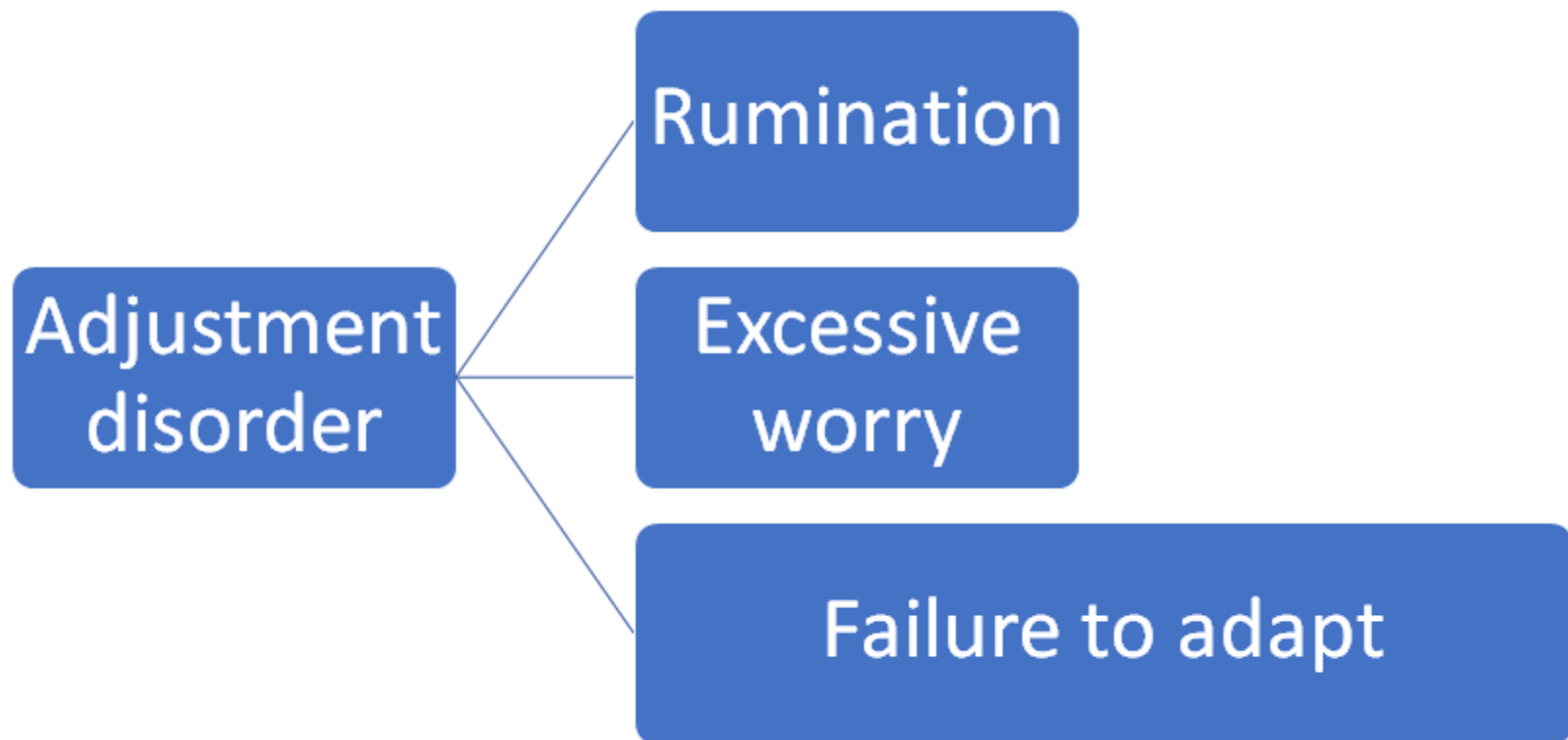
## **Mood disorders**

### **Major Depressive Episode**

# Adjustment Disorder (ICD-11)

Preoccupation with the stressor or its consequences, including **excessive worry**, recurrent and distressing thoughts about the stressor, or constant **rumination** about its implications

**Failure to adapt** to the stressor



# “Significance criterion”

ICD-11:

.....causes significant impairment in personal, family, social, educational, occupational or other important areas of functioning

DSM-5:

Causes clinically significant distress or impairment in social occupational, or other important areas of functioning

# Significance in palliative care?

Where is the cut-off for the need of intervention?

Symptom intensity

Significant

- distress
- psychological pain

Duration

Place in trajectory

Significant impact on

- quality of life
- function
- relationships

?

0 1 2 3 4 5 6 7 8 9 10

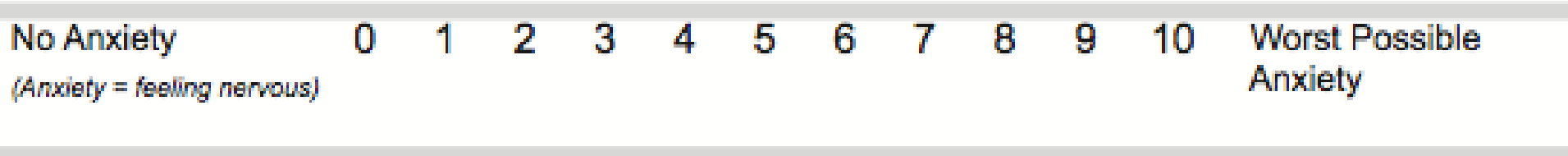
Anxiety

Stress and trauma related symptoms

# Stress and trauma related symptoms

## Anxiety





GAD-7 (Generalized Anxiety Disorder-7) Spitzer 2007



“Depressio” Latin

To press down

To be kept down

# Major Depressive Episode

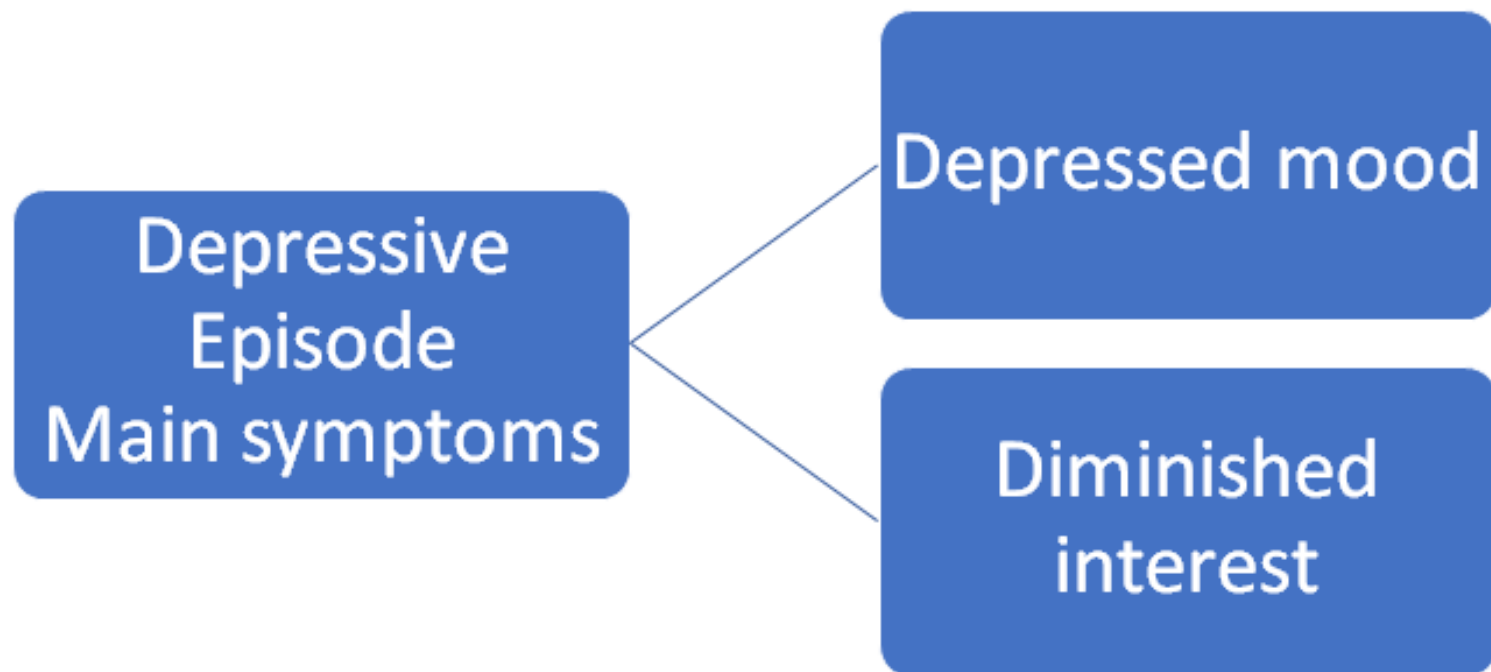
A set of symptoms

One or two of two main symptoms

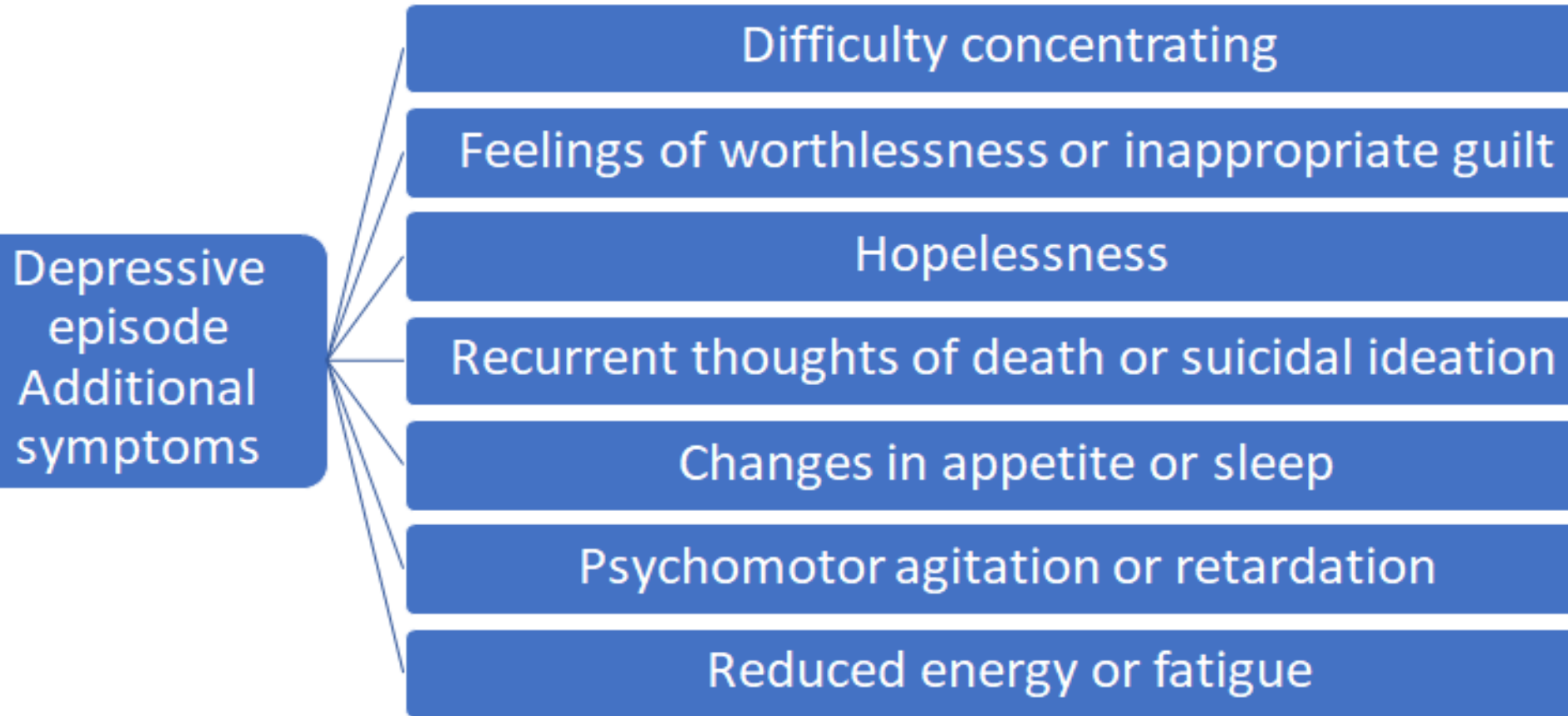
Duration of symptoms two weeks or more

Significance-criterion:

Impact on daily function



Depressive  
episode  
Additional  
symptoms



```
graph LR; A[Depressive episode Additional symptoms] --- B[Difficulty concentrating]; A --- C[Feelings of worthlessness or inappropriate guilt]; A --- D[Hopelessness]; A --- E[Recurrent thoughts of death or suicidal ideation]; A --- F[Changes in appetite or sleep]; A --- G[Psychomotor agitation or retardation]; A --- H[Reduced energy or fatigue];
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Difficulty concentrating

Feelings of worthlessness or inappropriate guilt

Hopelessness

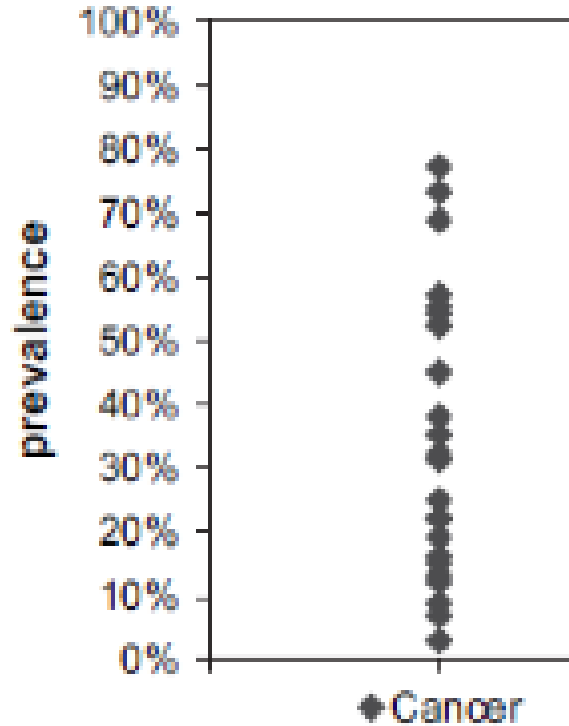
Recurrent thoughts of death or suicidal ideation

Changes in appetite or sleep

Psychomotor agitation or retardation

Reduced energy or fatigue

# Prevalence of depression



Solano et al 2006.

A dimensional phenomenon

Differences in assessment:

- Symptoms

- Ways to report

- Symptom degree, cut-off

- Duration of the symptoms

- Impact on daily function

# Prevalence

Teunissen 2007. Mitchell 2011. Grotmol et al 2016

## Depression

Assessed as a symptom  $\approx 40\%$

Depressive Episode  $\approx 15\%$

## Anxiety

Assessed as a symptom  $\approx 30\%$

Anxiety "Disorders"  $\approx 10\%?$

Depression and anxiety overlap and are strongly associated with reduced quality of life

# Assessment instruments

ESAS, Edmonton Symptom Assessment System

HADS, Hospital Anxiety and Depression Rating Scale

PHQ-2 - 4 - 9 , Patient Health Questionnaire

GAD-7, General Anxiety Disorder-7

MADRS, Montgomery Aasberg Depression Rating Scale

BDI, Beck Depression Inventory

# ESAS-r

## Edmonton Symptom Assessment System-revised

No Depression <i>(Depression = feeling sad)</i>	0	1	2	3	4	5	6	7	8	9	10	Worst Possible Depression
No Anxiety <i>(Anxiety = feeling nervous)</i>	0	1	2	3	4	5	6	7	8	9	10	Worst Possible Anxiety



During the past two weeks, how much (or how often) have you been bothered by the following problems:

1. Little interest or pleasure?
2. Feeling down, depressed or hopeless?
3. Feeling nervous, anxious, frightened,  
worried or on edge?
  1. Feeling panic or being frightened?
  2. Avoiding situations that make you anxious?

# Intervention

# Intervention

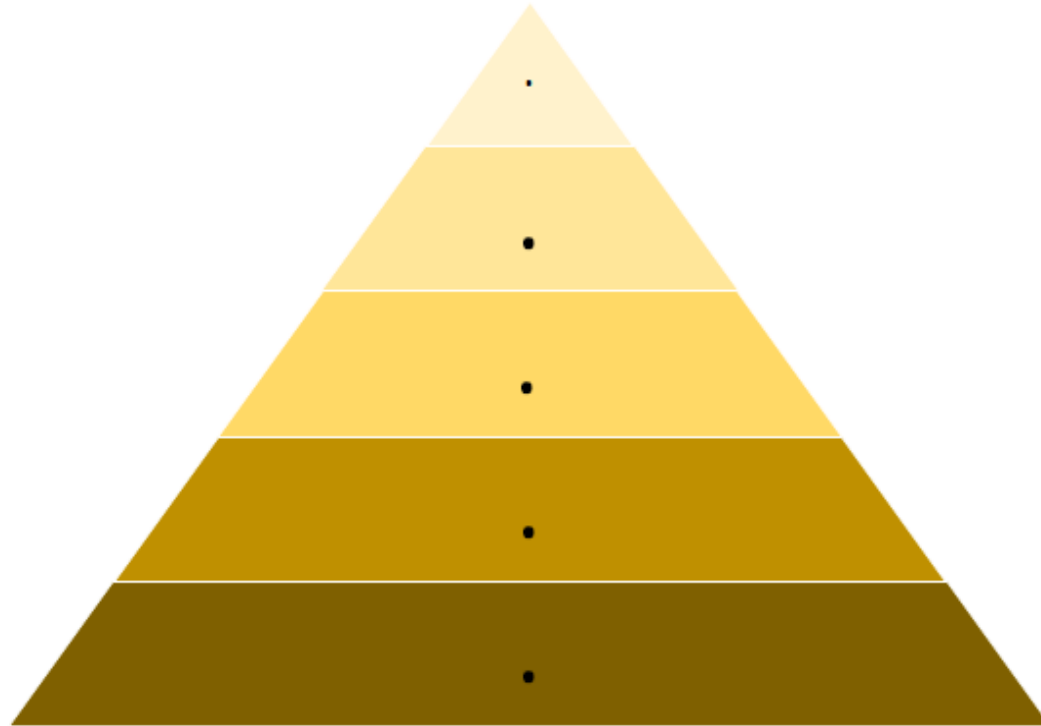
## Aims:

To alleviate psychological symptoms

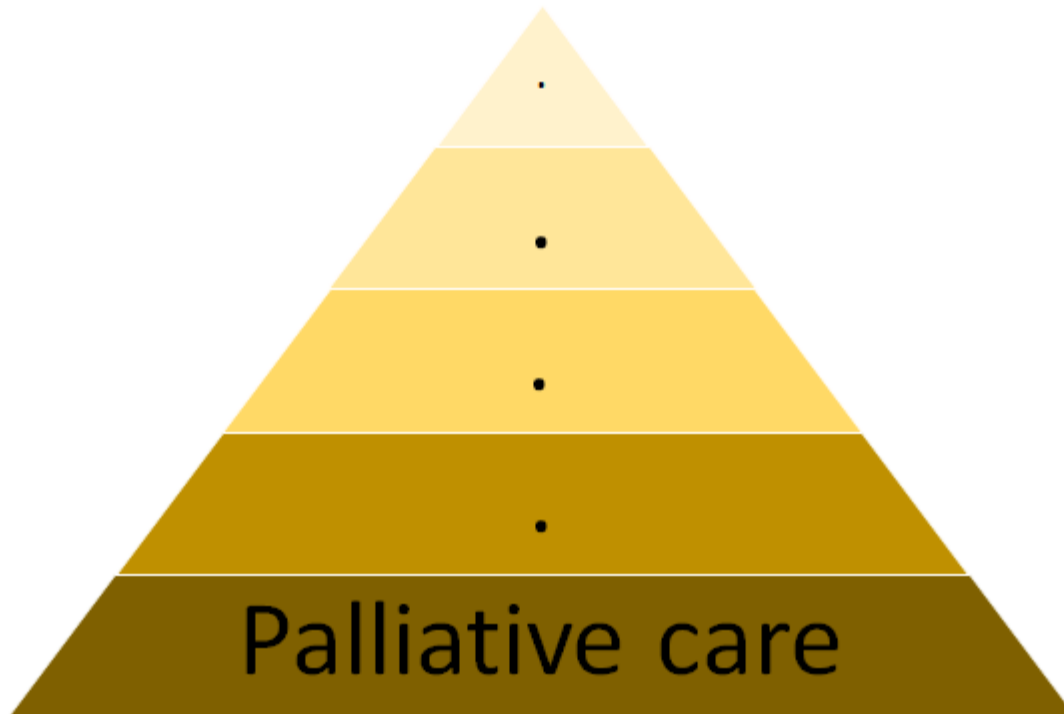
To facilitate adjustment

# Intervention

Rodin 2007. Rayner 2010



# Palliative care



# Palliative care

Temel 2010. Pirl 2012. Basch 2016

Patient perspectives and concerns. Goal and evaluation of treatment

Systematic Symptom assessment for early intervention.

Symptom control

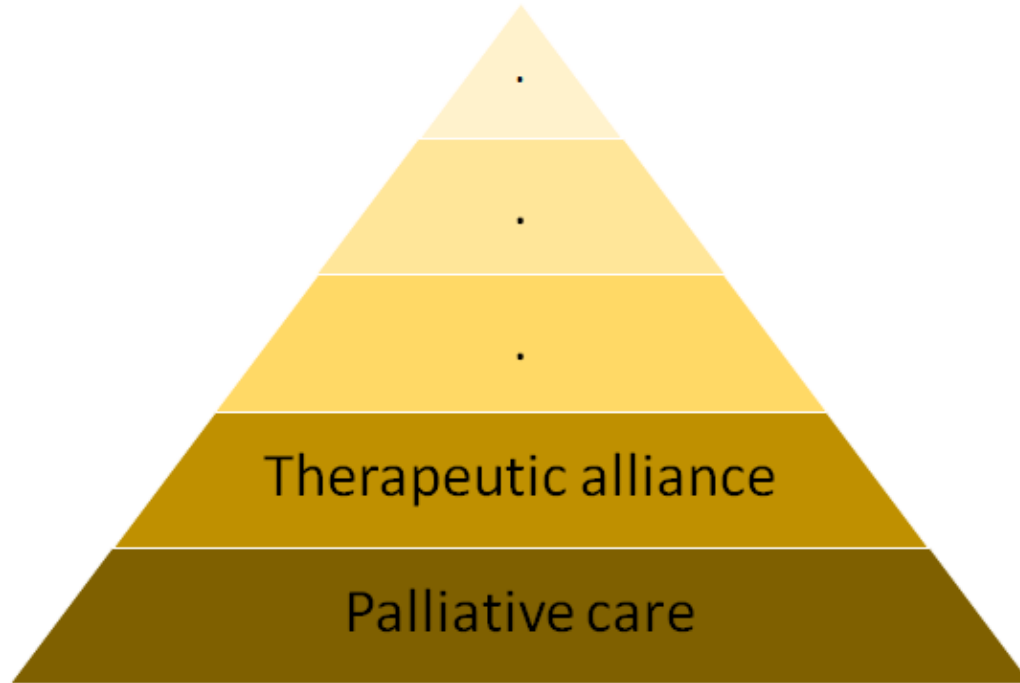
Function and roles

Coping and adapting

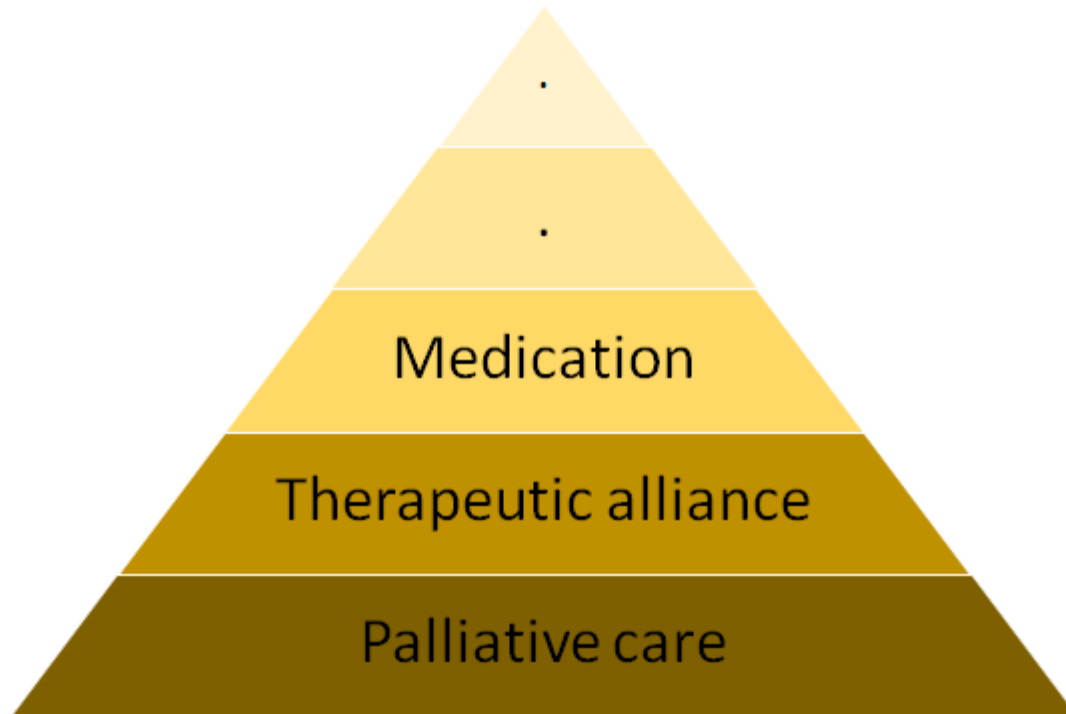
Plan and responsibility

Contact if needed

# Therapeutic alliance



# Medication





# Medication

Antidepressants

Benzodiazepines

# Antidepressants

Effects on

Depression

Anxiety

Sleep problems

In palliative care:

Meta-analysis OR 2,25 (1,38-3,67) Rayner 2014

Major Depressive Episode or Adjustment Disorder

Grassi and Rodin 2017

**Table 3. First-line ADs in cancer patients**

Generic name	Optimal indication	Standard adult dose	Level of evidence/grade of recommendation
Citalopram/Escitalopram	<ul style="list-style-type: none"> <li>Few CYP450 drug interactions</li> </ul>	Start: 10–20 mg o.d./ (5–10 mg q.h.s.) Goal: 20–40 mg/ (10–20 mg) Max: 40 mg o.d./ (20 mg q.h.s.)	<ul style="list-style-type: none"> <li>Level III<sup>a</sup> evidence</li> <li>Strong, moderate quality</li> </ul>
<b>SSRI</b>	<ul style="list-style-type: none"> <li>Escitalopram may have more rapid onset of action</li> </ul>		
Venlafaxine/Desvenlafaxine	<ul style="list-style-type: none"> <li>Optimal choice for patients on tamoxifen</li> <li>Consider for prominent hot flashes</li> </ul>	Start: 37.5–75 mg q.a.m./ (50 mg) Goal: 75–225 mg/ (50–100 mg) Max: 300 mg q.a.m./ (100 mg)	<ul style="list-style-type: none"> <li>Level III<sup>a</sup> evidence</li> <li>Strong, low quality</li> </ul>
<b>SSRI and SNRI</b>			
Bupropion XL	<ul style="list-style-type: none"> <li>Consider for prominent fatigue</li> </ul>	Start: 150 mg q.a.m. Goal: 150–300 mg Max: 450 mg q.a.m.	<ul style="list-style-type: none"> <li>Level III<sup>a</sup> evidence</li> <li>Strong, low-quality</li> </ul>
<b>NDRI</b>	<ul style="list-style-type: none"> <li>Aids sexual function</li> </ul>		
Duloxetine	<ul style="list-style-type: none"> <li>Separate indications for neuropathic and chronic pain</li> </ul>	Start: 30 mg q.a.m. Goal: 30–60 mg Max: 120 mg q.a.m.	<ul style="list-style-type: none"> <li>Level III<sup>a</sup> evidence</li> <li>Strong, low-quality</li> </ul>
<b>SNRI</b>			
Mirtazapine	<ul style="list-style-type: none"> <li>Consider for prominent insomnia, anorexia/ cachexia, diarrhoea</li> </ul>	Start: 7.5–15 mg q.h.s. Goal: 15–45 mg Max: 60 mg q.h.s.	<ul style="list-style-type: none"> <li>Level III<sup>a</sup> evidence</li> <li>Strong, low-quality</li> </ul>
<b>NaSSA</b>			

Noraderenerg. Selective Serotoninerg Antidepressant. Antihistaminerg

# Mirtazapine

Effect on

depression

anxiety

sleep problems: early-, middle- and late insomnia

Initial effect after 2 weeks (opposed to 3-4 weeks)

Increases appetite and weight

Antiemetic properties

Limited drug interactions

No increased risk of GIT bleeding

Tiredness a possible side effect

Orally dissolvable formulation

Grassi and Rodin 2017. Economos et al 2019

# Antidepressants. Limitations

Life expectancy

No parenteral administration

Adverse effects:

- Sedation

- Headache

- Increased risk of GIT bleeding (not mirtazapine)

- Cognitive change

- Emotional blunting

[Intervention Review]

# **Drug therapy for symptoms associated with anxiety in adult palliative care patients**

Susan Salt<sup>1</sup>, Caroline A Mulvaney<sup>2</sup>, Nancy J Preston<sup>3</sup>

2019

# Benzodiazepines. GABA agonists

No clear differences in profiles of effect and side effects between the different benzodiazepines

Routes of administration:

- peroral, sc, iv, rectal, intranasal

- sc continuous infusion (midazolam)

Adverse effects:

- Sedation

- Cognitive effects

# Other medication

- Collaboration with psychiatrists
  - Antiepileptics i.e. Lamotrigine, Valproate Geddes et al 2009. Smith et al 2010
  - Antipsychotics i.e. Quetiapine Albert et al 2016
- Anesthetics
  - Ketamine and Esketamine Glutamate receptor modulators  
Dean et al Cochrane review. September 2021.  
*Ketamine and other glutamate receptor modulators for depression in adults with unipolar major depressive disorder*  

24h: Ketamine iv once	SMD -0.87 (95% CI -1.26 to -0.48)
24h: Esketamine intranasal x2/week	SMD -0.31 (95% CI -0.45 to -0.17)
- Buspirone? Serotoninerger



# Other medication

## Psychostimulants. Dopaminerg

- Methylphenidate
- Dextramphetamine

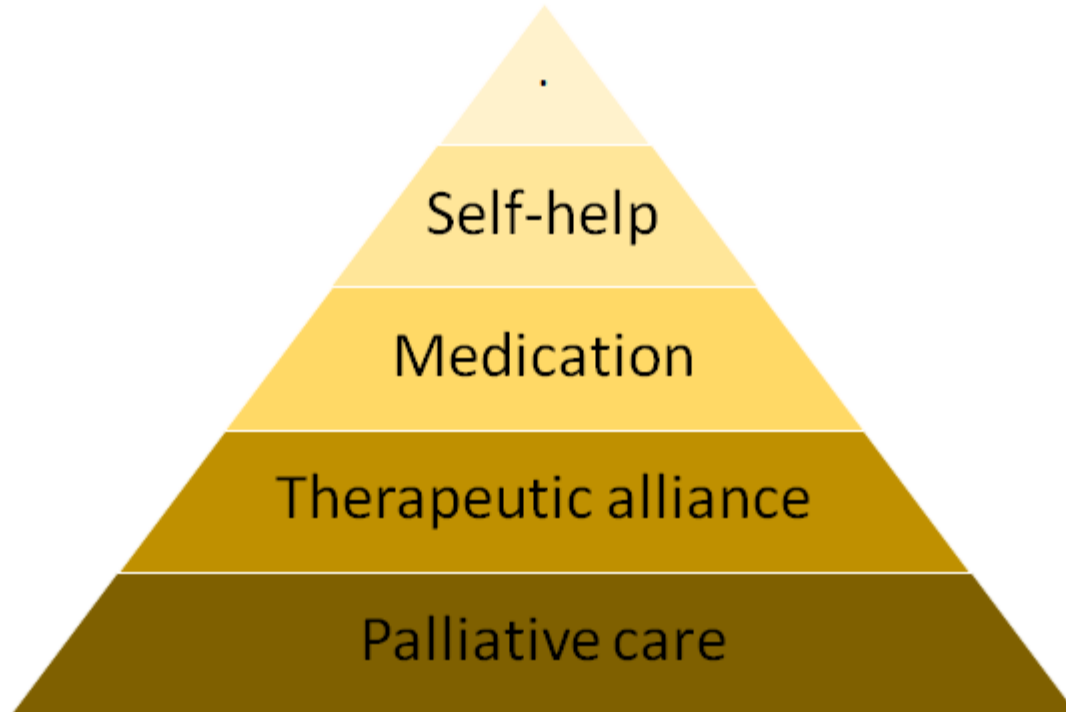
Mc Intyre et al 2017. *The Efficacy of Psychostimulants in Major Depressive Episodes. A Systematic Review and Meta-analysis.*

OR 1,41 (CI 95% 1,13-1,78)

*“....may be capable of reducing depressive symptom severity....A strong conclusion cannot be made.”*

More evidence is needed

# Self-help



# Guided self-help

Activation

Social contact. Relationships

Affiliation

Priorities. Personal goals

Meaning and purpose

Goal for a Self-help program:

Divert non-adaptive coping

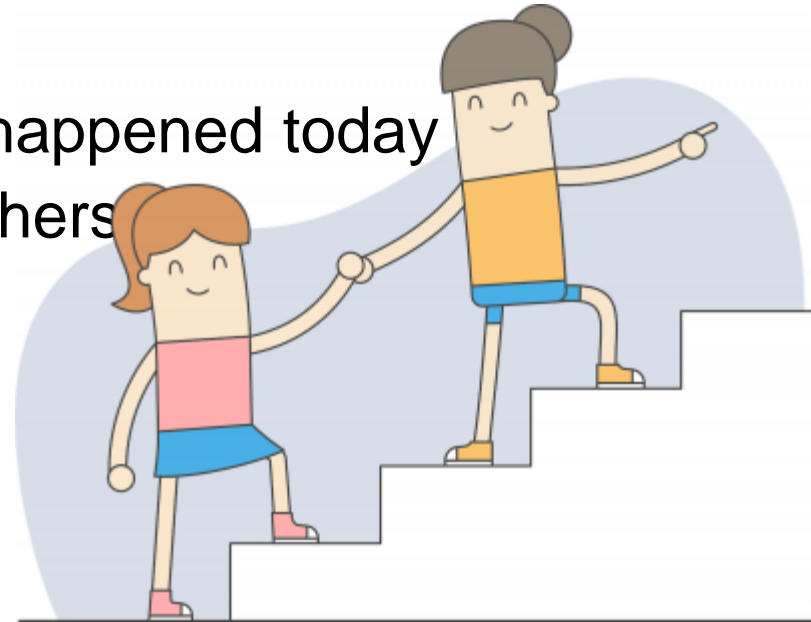
Support purposeful coping

Change focus

Life content

# Self-help

- Connect with other people. Keep in contact
- Make yourself a good day
- Pay attention to the present moment
- Pay attention to the small things
- Write down three good things that happened today
- Give to others. Do something for others
- Activity. Do something that you like
- Sleep enough



# Self-help program. Behavioral activation

Personal contact

Proper structure

Appointments

Follow-up



CrossMark

**Integrated collaborative care for major depression comorbid with a poor prognosis cancer (SMaRT Oncology-3): a multicentre randomised controlled trial in patients with lung cancer**

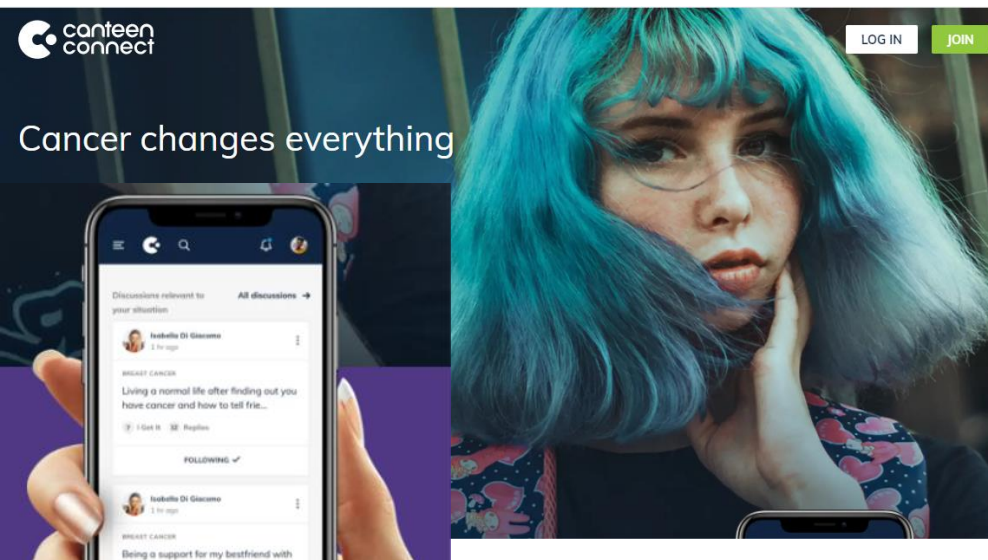
*Jane Walker, Christian Holm Hansen, Paul Martin, Stefan Symeonides, Charlie Gourley, Lucy Wall, David Weller, Gordon Murray, Michael Sharpe, for the SMaRT (Symptom Management Research Trials) Oncology-3 Team*

Walker et al 2014

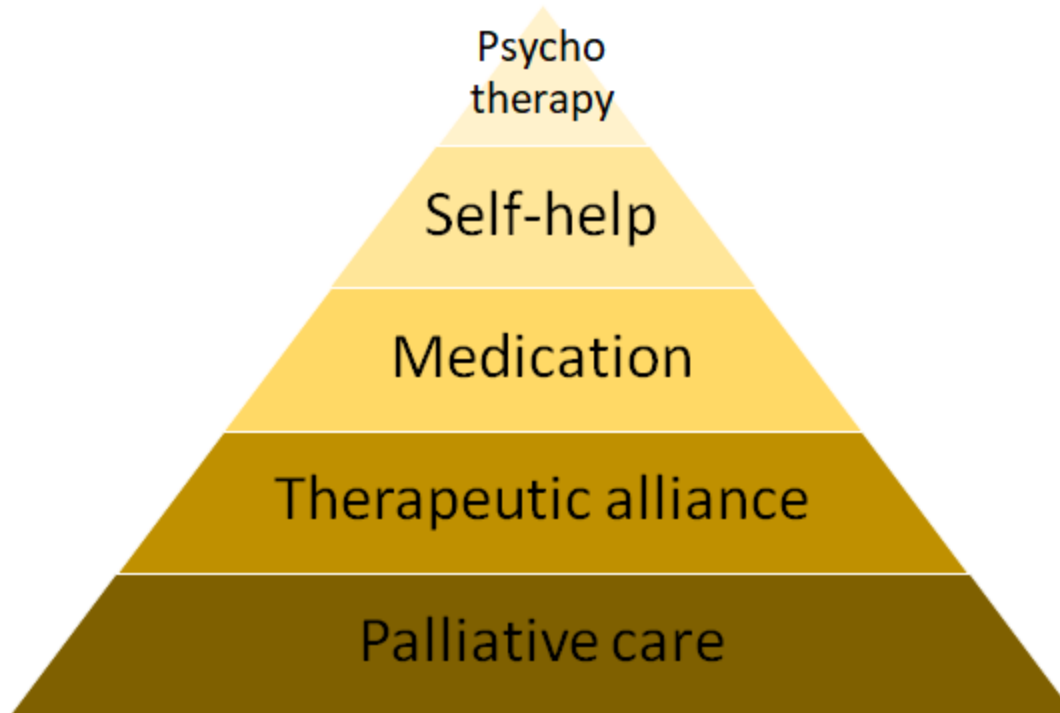
# Guided self-help

- Coach/ therapist
- Groups/ relationships
- Online program: ressources, social platform, follow-up

Veiledet Selvhjelp.no



# Psychotherapy



# Psychotherapy

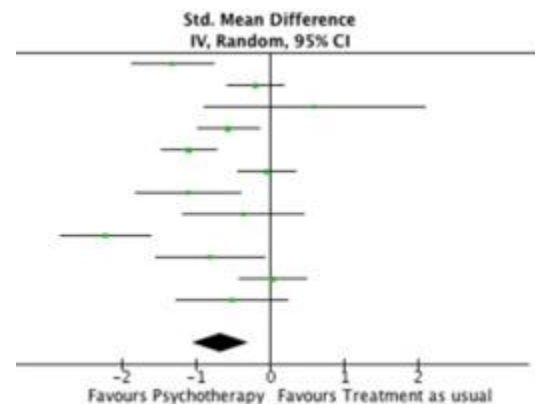
Effect in pall care patients

## Meta-analysis

Depression	SMD - 0,67	(-1,06 - -0,29)
Anxiety	SMD - 0.65	(-1,05 - -0,26)
Dep + Anx	SMD - 1,02	(-1,70 - -0.35)

Cannot differentiate effect between  
the different psychotherapeutic  
approaches.

Individual psychotherapy and group therapy





# CALM

## Managing Cancer and Living Meaningfully

4-6 sessions

# Elements from psychotherapeutic approaches and theories

Supportive psychotherapy

Modern grief theory

Mentalisation and Cognitive therapy

Existential therapy

# Supportive therapy

A supportive relationship

A secure base

Attention

Affirming: The person, reflections, insight, coping

Empathic listening

Curiosity

Co-thinking

A reflective space

# Modern grief theory

The dual process model

Double awareness

Alternate between two approaches to the situation

- To confront

- To divert

Time

Process of adjustment

# Mentalisation and Cognitive therapy

Mentalisation: Reflecting on thoughts and feelings

Cognitive therapy: Content of thoughts, changing thoughts

Verbalizing experience

- Formulating. Reformulating
- The naive observer

# Mentalisation and Cognitive therapy

Expand the experience

- Multiple views
- Not correcting
- Not false reassurance

Distinguishing feelings from facts

Metaphores

# Existential therapy

Fear of death as a condition of life

Leave traces behind

Purpose beyond oneself

Relationships

# Existential therapy

## Fear of death



# Psychotherapy

Availability of a therapist

Eventually combined with medication

Adjustment disorder or a Depressive episode

Adjustment of the therapy

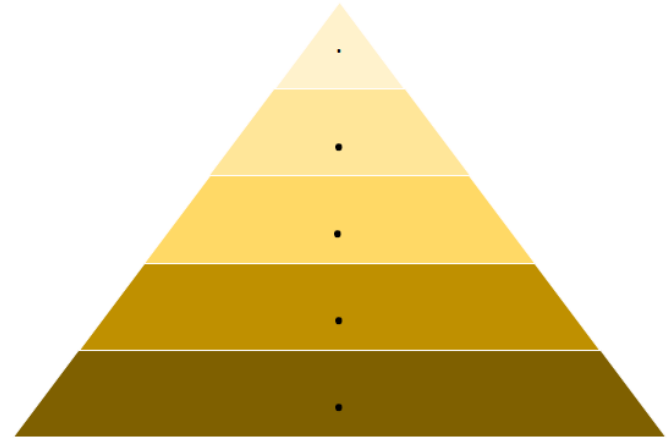
# Palliative care

The principles of self-help and psychotherapy are simplified integrated in palliative care

**Significance** in palliative care

Dimensional significance

Dimensional intervention



# Late palliative care setting

Good palliative care

Support

Family

# Suicidal ideation

Nuance the content

- Despair
- Concrete content or plans

The psychiatric team

# Summing up

