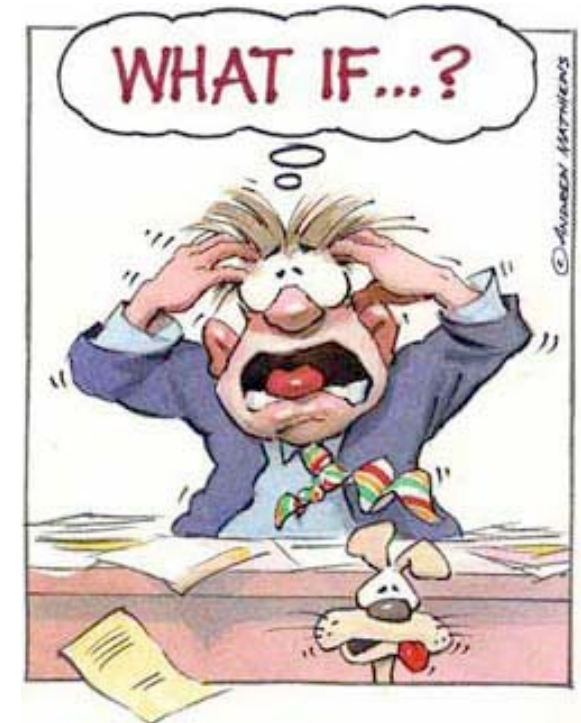


# Worry and Rumination: Concepts and Neurobiological correlates



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# Introduction

- Worry and rumination:
  - Definitions
  - Theoretical concepts/ information processing
  - Neural Correlates
- Clinical implications

# Definition: Worry

- **Repetitive**, relatively **uncontrollable** thoughts that emerge when a person faces a potential **threat**

(Beck, 1987)

- Association with **anxiety**: *state of anxious apprehension*
- Central feature of **Generalized Anxiety Disorder**



# Theoretical Concepts: GAD

- **Verbal cognitive activity** that allows the individual to **avoid the intrusion of more arousing material** such as fear-related **images** and their associated **autonomic responses**

(Borkovec 2004)

- Theoretical models of GAD: **AVOIDANCE OF INTERNAL EXPERIENCES**

- **Strategy for gaining emotional control:**

-> 'Short-term **protective role**'

-> Interferes with **functional emotional processing**

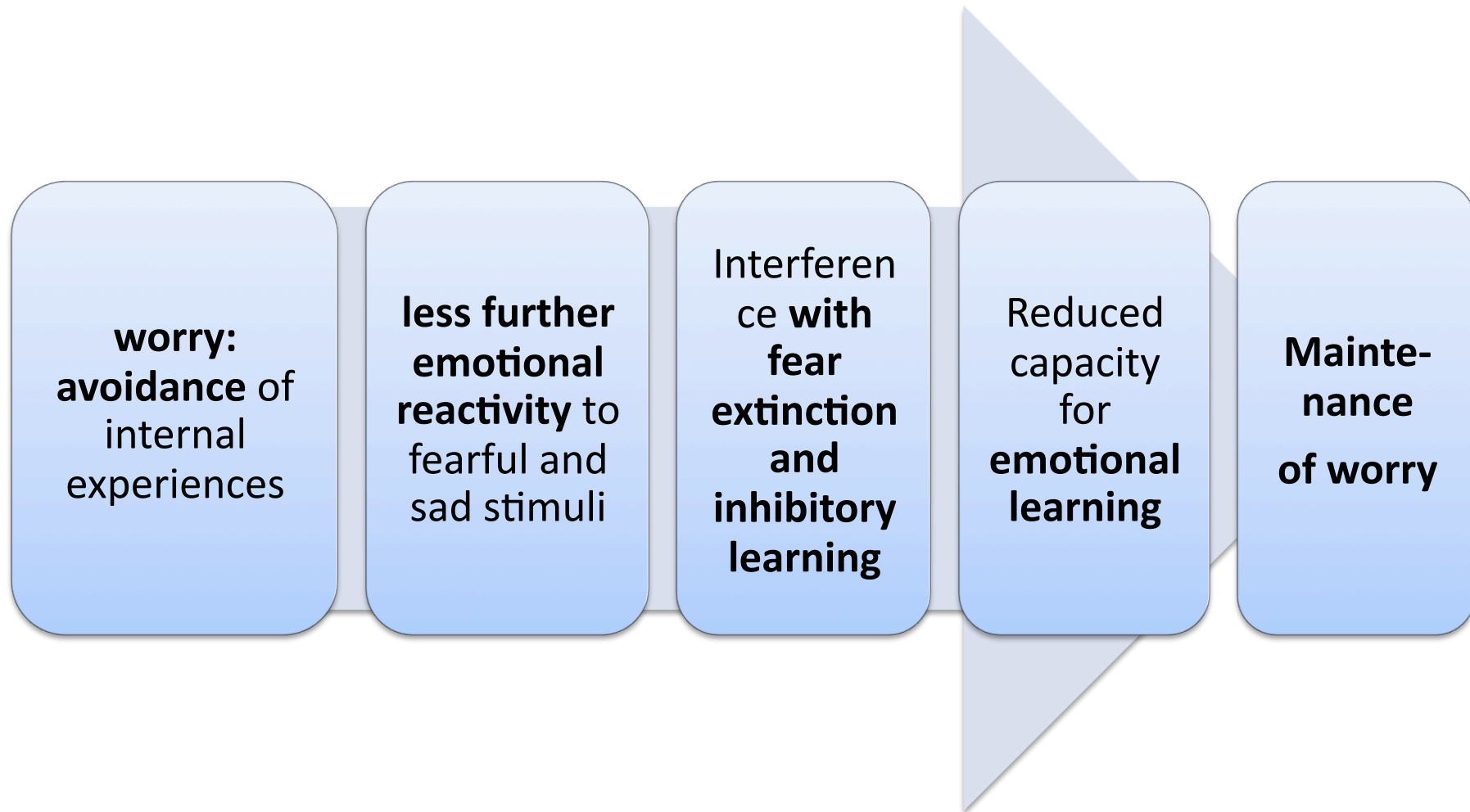
-> Prevents the extinction of **fear**

(Review Behar et al. 2009)



# EMOTIONAL INFORMATION PROCESSING: GAD

(Llera 2010; Newman 2011)



# Definition: rumination

- Repeated **self-focused** thoughts about **mood symptoms**, **negative aspects of one's self** and **one's life**
- Mostly related to **depression**

## **EMOTION FOCUSED RUMINATION**

Resons styles theory – in respons to mood (Nolen-Hoeksema 1998)

## **STRESS-REACTIVE RUMINATION:**

following stressful life events (Robinson and Aloy, 2003)



# Theoretical Concepts: Rumination:

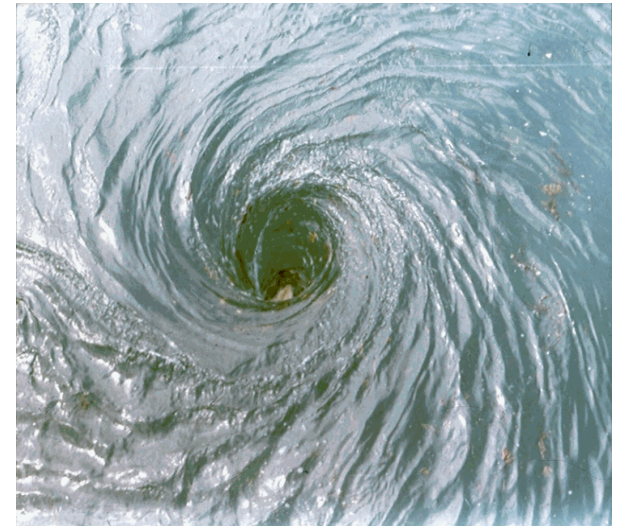
Problems in **'COGNITIVE TOP-DOWN CONTROL'** in **emotion regulation**

(Review Gotlib, 2010)

- **disengaging attention** from negative stimuli
- **negative autobiographical memories**
- **inhibitory control** of mood-congruent thoughts/memories
- ability for **Reappraisal**

**Content of WORKING MEMORY:**

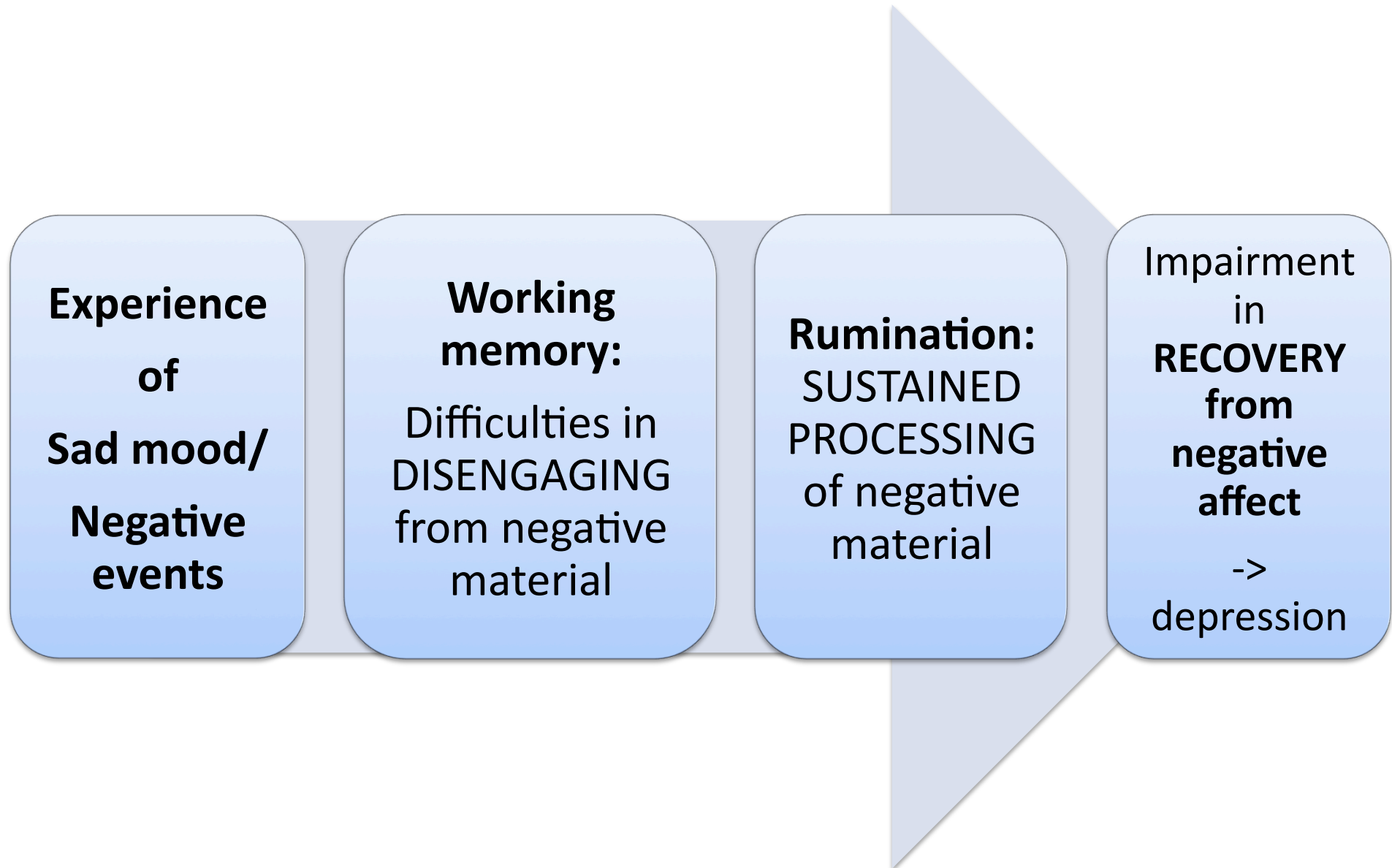
**Sustained processing of negative material**



# EMOTIONAL INFORMATION PROCESSING:

## Rumination

(Gotlib 2009, Smith 2010)





# Worry versus Rumination

(Smith 2010, Hong 2006; Watkins 2005, Segerstrom 2003, Wells 1999)

## WORRY

- Anxious apprehension, GAD
- **Content:** Future, problem-solving
- State of cognitive **over-control**
- **Avoidance** of internal experiences

## RUMINATION

- Depressive mood and symptoms
- **Content:** Past, loss
- Problems in **cognitive control** of working memory
- Problems in **disengaging attention** from negative material

Bridge between theoretical  
concepts and neurobiology?



# Neural correlates of Worry

- Anxiety disorders: **TWO CLASSES?**

(Review Berkowitz 2007)

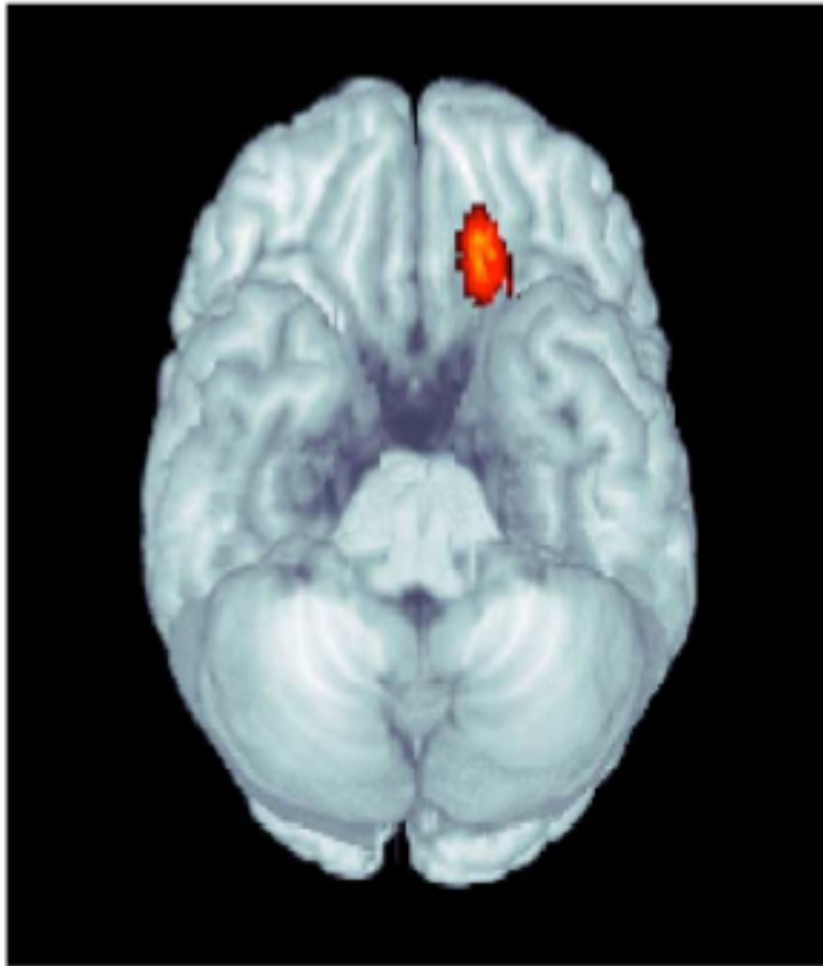
	Intense Fear/panic	GAD and OCD
Prefrontal cortex	↓	↑
Limbic system	↑	↓
Autonomic nervous system	↑	↓

- **Neural response to Emotional stimuli in GAD?**

Hyporesponsiveness in amygdala to faces with fearful expressions in GAD

(Blair 2008)

# Worry: non-clinical population



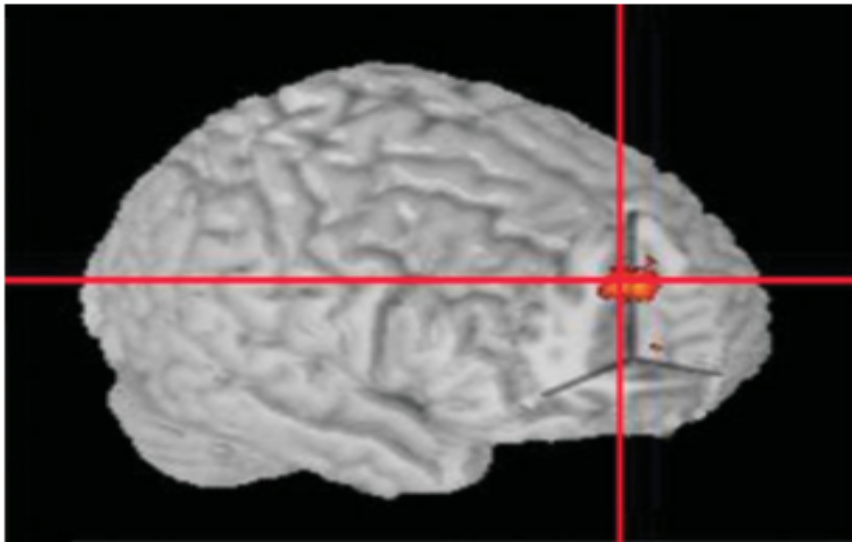
Negative correlation between  
activation of **left orbitofrontal  
cortex** and **amygdalae**

**PREFRONTAL OVERACTIVATION  
CAUSES LIMBIC INHIBITION?**  
≈ avoidance theory?

Fig. 3. Negative correlation between activated foci of the amygdalae and the left orbito-frontal cortex (height threshold:  $P=0.005$ ; extent threshold:  $k+100$  voxels).

# Worry: GAD

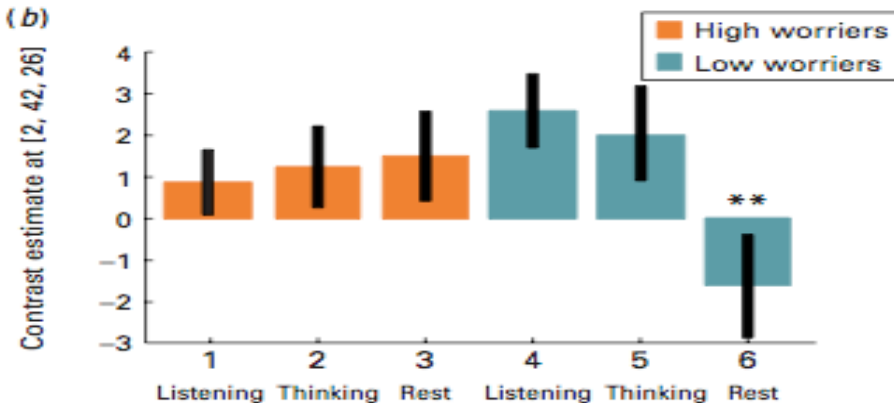
(a)



✓ **POST-WORRY RESTING STATE:**

**GAD: same regions remain active**

(b)



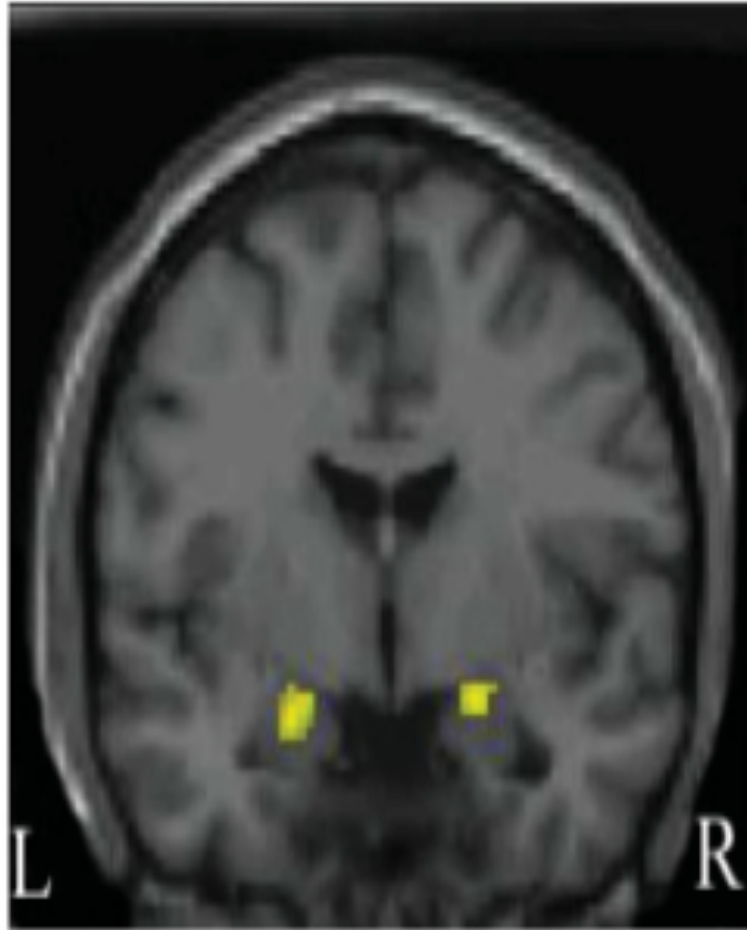
**QUANTITATIVE DIFFERENCE!**

**~ Prolonged preoccupation in GAD?**

(Paulesu et al. 2009)

# Rumination: nonclinical samples

(Ray and Ochsner 2005)



Trait rumination correlation with Amygdala activation in negative Reappraisal task

## TRAIT RUMINATION:

↓ Top-Down prefrontal control causing disinhibition amygdalae?

# Rumination: Clinical samples

- **RUMINATION INDUCTION TASK**

(Cooney 2010)

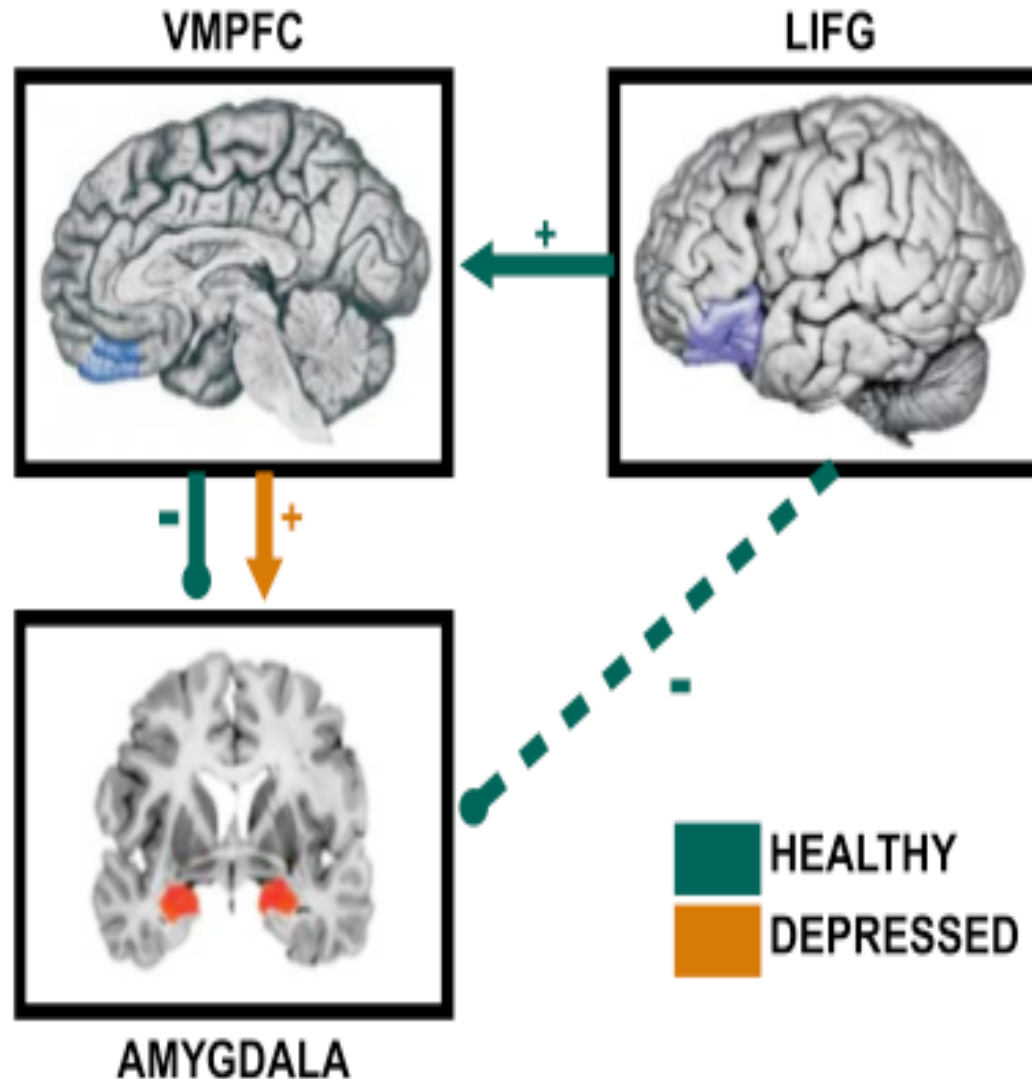
Hyperactivity in **limbic structures**, areas involved in **mediating self-referential content** (mPFC, ACG), retrieval of **mood-congruent autobiographical memories** (parahippocampus, PCG)

≈ **disinhibition limbic system**

≈ **activation areas self-referential thinking**

≈ **disengagement problems**

# Rumination: clinical samples



**FAILURE TO REGULATE  
MOOD IN DEPRESSION?**

**counterproductive  
recruitment in top-down  
prefrontal-subcortical**

(Johnstone 2007)



# SUMMARY

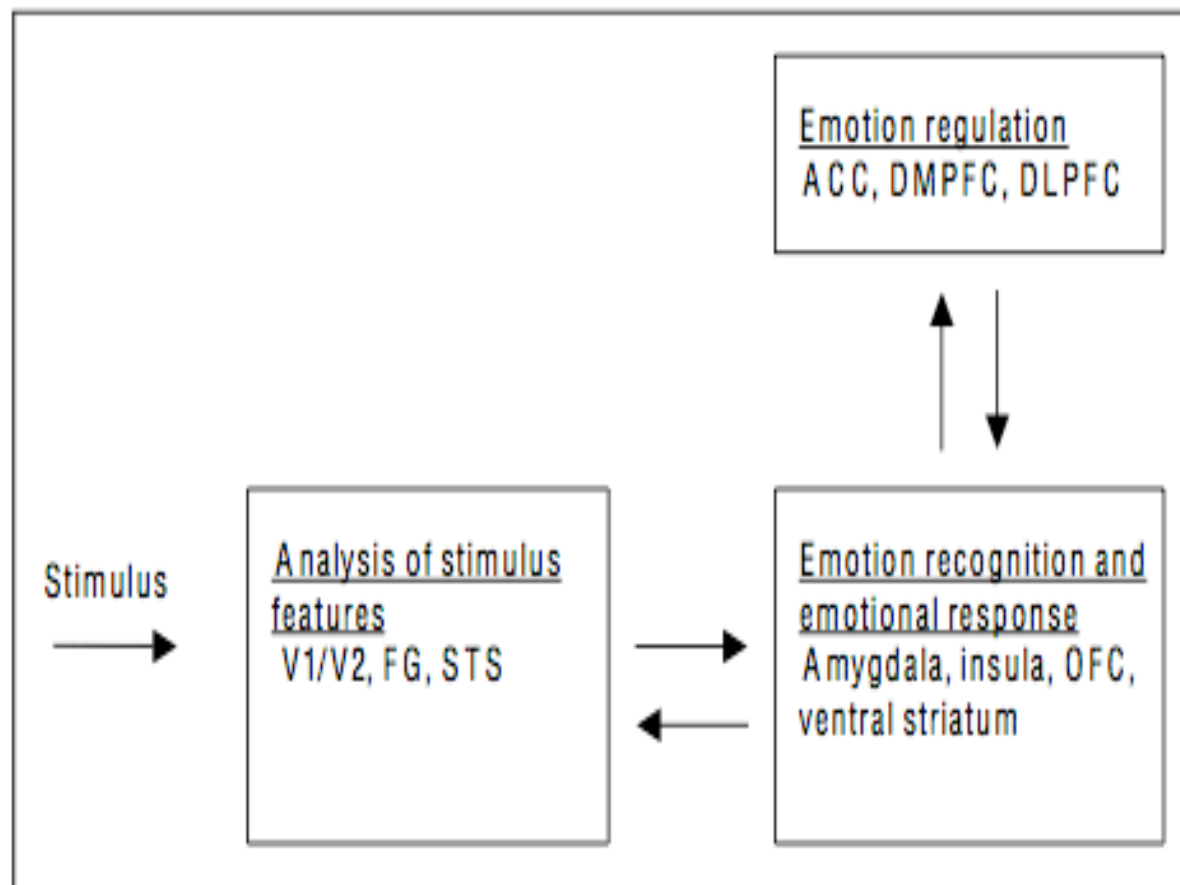
WORRY	RUMINATION
emotional <b>CONTROL</b> strategy <b>AVOIDANCE</b> of internal experiences	<b>TOP-DOWN</b> problems in control of content working memory difficulties <b>DISENGAGING</b> from negative material
Worry-induced <b>OVERACTIVATION OF PREFRONTAL REGIONS</b>	Attenuated <b>activation of top-down prefrontal control, sustained overactivation</b> of regions mediating <b>SELF-REFLECTION</b>
<b>SUPPRESSION OF LIMBIC REGIONS</b>	<b>DISINHIBITION OF LIMBIC REGIONS</b>

## Integrative research from an ‘emotional information processing’- point of view

(Lepänen 2006)

**Figure 2. A simplified diagram showing the psychological processes involved in the processing of (visual) emotional signals and the neural structures that subserve these processes**

Modified from models presented in [22–25]. ACC, anterior cingulate cortex; DLPFC, dorsolateral prefrontal cortex; DMPFC, dorsomedial prefrontal cortex; FG, fusiform gyrus; OFC, orbitofrontal cortex; STS, superior temporal sulcus.



# Clinical Relevance



- **Cognitive vulnerability factors**

**high risk populations** (offspring, partial remitted, recovered)

differences in **cognitive and neural processing of emotional stimuli** may play a **key role** in the development, maintenance or recurrence of mood and anxiety disorders

(Lepänen 2007)

- **Attention and cognitive interpretation training** program
- **Rumination-focused CBT** for residual depression

(Mathews 2004)

(Watkins 2007)

- **Outpatient CBT-based group program UZ Gent: 8 sessions**

Protocol: attention training, development of alternative strategies; effective engagement in tasks, functional thinking styles

## Rumination-focused cognitive–behavioural therapy for residual depression: phase II randomised controlled trial

Edward R. Watkins, Eugene Mullan, Janet Wingrove, Katharine Rimes, Herbert Steiner, Neil Bathurst, Rachel Eastman and Jan Scott

**Table 2** Mean (s.d.) scores on outcome measures at baseline assessment and post-intervention assessment for rumination-focused cognitive–behavioural therapy (RFCBT) group and treatment as usual (TAU) group

Measure	Baseline		Post-treatment		Mean difference in change scores <sup>a</sup> (95% CI)	Effect size <sup>b</sup>
	TAU	RFCBT	TAU	RFCBT		
Hamilton Rating Scale for Depression	12.19 (2.80)	13.29 (3.32)	9.05 (5.25)	5.48 (5.15)	4.67 (0.28–9.05)	0.94
Beck Depression Inventory-11	28.29 (7.63)	30.76 (8.17)	20.71 (10.84)	12.71 (11.37)	7.57 (1.86–19.08)	1.11
Rumination Scale of Response Style Questionnaire	59.17 (8.55)	58.45 (12.34)	54.38 (11.02)	44.50 (12.86)	9.16 (–3.4–21.73)	0.65

a. Mean difference in change scores: change on outcome measure from baseline to post-treatment for rumination-focused cognitive–behavioural therapy minus change in outcome measures from baseline to post-treatment for treatment as usual.

b. Between-group effect size for change in symptoms (Cohen's  $d$  where  $d = M_1 - M_2 / \sigma_{\text{pooled}}$ ,  $\sigma_{\text{pooled}} = \sqrt{[(\sigma_1^2 + \sigma_2^2)/2]}$ . ( $M_1$  is the mean change on outcome measure from baseline to post-treatment for rumination-focused cognitive–behavioural therapy;  $M_2$  is the mean change in outcome measures from baseline to post-treatment for treatment as usual). Large effect sizes were defined as  $\geq 0.80$ .

Response: 26% (TAU) v. 81% (rumination-focused CBT)  
 Remission: 21% (TAU) v. 62% (rumination-focused CBT)  
 Relapse: 53% (TAU) v. 9.5% (rumination-focused CBT)



*With aknowledgment to Dr Van den Abbeele, UZ Gent*

*Professor Lemmens, UZ Gent*