

ANXIETY DISORDERS

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Anxiety is abnormal fear that is out of proportion to any external stimulus, which is an unpleasant and unjustified sense of apprehension often accompanied by physiological symptoms.

What is anxiety?

Anxiety is a state of apprehension, tension, or uneasiness that occurs in anticipation of internal or external danger

The anxiety syndrome includes motor tension, autonomic hyperactivity, apprehensive expectation, and heightened vigilance

Anxiety occurs in a variety of neurological and medical disorders and can be precipitated by drugs

Normal and pathological

- Difference between pathological anxiety and anxiety as a normal or adaptive response
- /Classic flight and fight response...or feeling anxiety before exam, can be adaptive – e.g. in coping with stress/
- Pathological anxiety – in content and seriousness, can not be controlled - phobias, panic, ocd, ptsd....sense of fearfulness, terror, apprehension and vegetative symptoms



Heidegger, Kierkegaard in philosophy - the existential anxiety

ANXIETY AS A CONDITION OF LIVING

- **Existential anxiety** is the unavoidable result of being confronted with the “givens of existence” - death, freedom, choice, isolation, and meaninglessness. It arises as we recognize the realities of our mortality, our confrontation with pain and suffering, our need to struggle for survival, and our basic fallibility.
- **Normal anxiety** is an appropriate response to an event being faced. It is not a therapeutic goal to eliminate normal anxiety because it can be used as a motivation to change.
- **Neurotic anxiety** is anxiety about concrete things that is out of proportion to the situation. It is typically out of awareness, and it tends to immobilize the person.

Some historical data

- ..unlike depression, a syndrome recognized for centuries, the syndrome of anxiety has been recognized only in the relative recent times
- DaCosta –irritable heart 1871
- Freud's conceptualisation brought the patient's inner subjective feeling to the forefront, emphasizing the sense of fearfulness, terror panic

Neurotic symptoms

- Symptoms cause worrying and suffering for the patients
- Could not explained by organic abnormalities
- Chronic and recurrent (may be in other form)
- Not limited only external stresses
- Reality testing is intact
- Symptoms are not under intentional control
- The origin of symptoms is in the past history of personality
- The patient awares the illness – behaves like other patients, who suffers in somatic disorder



Neurotic disorders

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graph TD; ND([Neurotic disorders]); AD((Anxiety dis.)); SD([Somatoform d.]); ND --- AD; ND --- SD; SD --- S1([ ]); SD --- S2([ ]); SD --- S3([ ]); SD --- S4([ ]); SD --- S5([ ]); ND --- S6([ ]); ND --- S7([ ]);
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Anxiety dis.

Somatoform d.

ANXIETY DISORDERS

- panic disorder
- agoraphobia
- generalised anxiety disorder
- phobias (specific, social)
- obsessive-compulsive disorder
- posttraumatic stress disorder
- acute stress disorder

Anxiety disorders – diff.diagnosis the steps of decision tree

- 1. General medical conditions**
- 2. Substance abuse**
- 3. Response to stress? (acute stress disorder, PTSD, adjustment disorder)**

4. FORMS - DURATION - CONTEXT

Discrete, sudden anxiety attack ---- panic

- 5. Next steps – depending on the content of anxiety**

**The source of anxiety, what situations avoided?
(fear from the next panic attack, fear from a social adjustment, from a infection, contagion, e.g)**

--- GAD - residual category, the hard to be controlled chronic anxiety

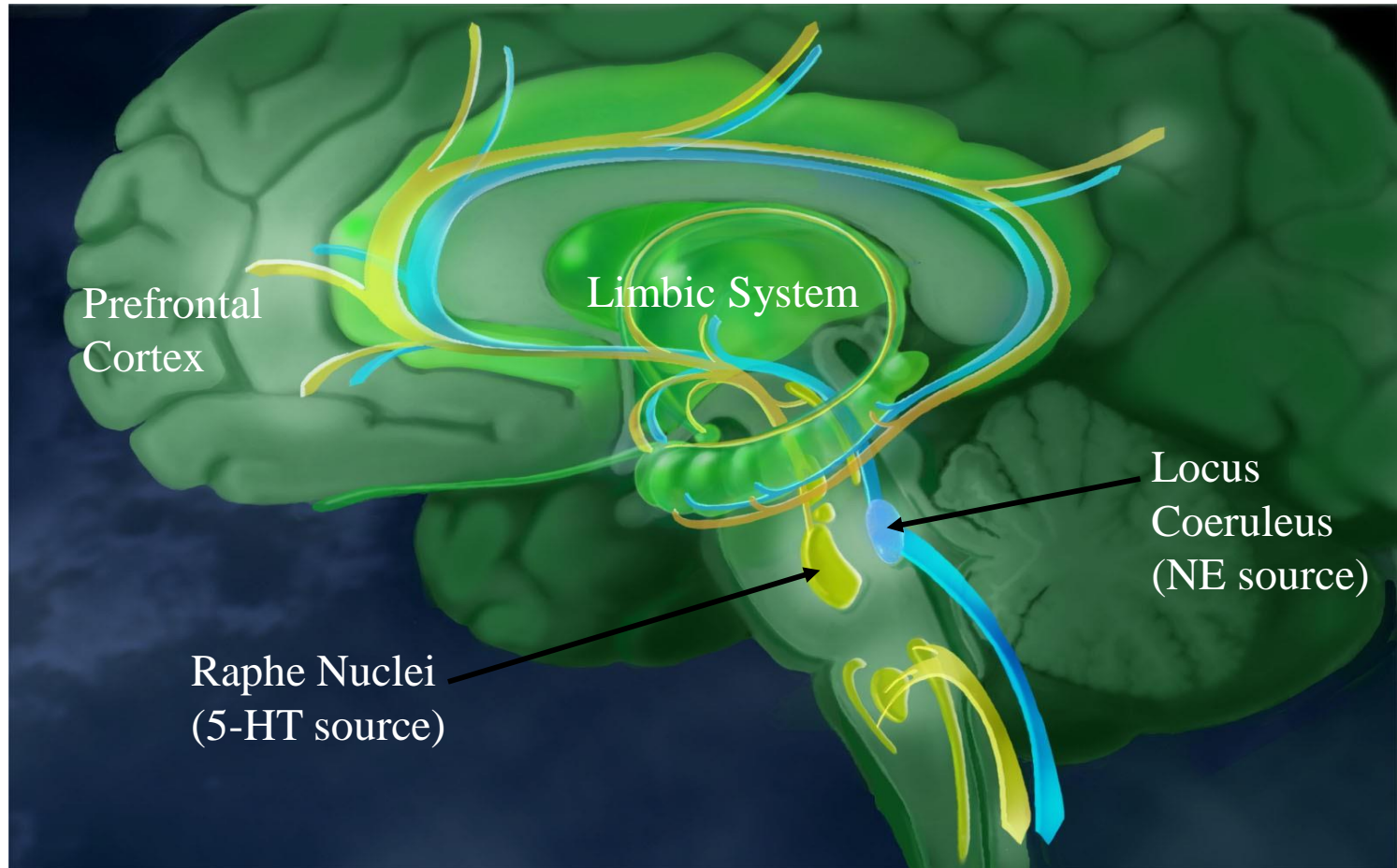
Table 10–8. Differential diagnosis of anxiety

Medical illness	Personality disorders
Angina	Adjustment disorder with anxious mood
Cardiac arrhythmias	Drugs
Congestive heart failure	Caffeine
Hypoglycemia	Aminophylline and related compounds
Hypoxia	Sympathomimetic agents (e.g., decongestants and diet pills)
Pulmonary embolism	Monosodium glutamate
Severe pain	Psychostimulants and hallucinogens
Thyrotoxicosis	Alcohol withdrawal
Carcinoid	Withdrawal from benzodiazepines and other sedative-hypnotics
Pheochromocytoma	Thyroid hormones
Menière's disease	Antipsychotic medication
Psychiatric illness	
Schizophrenia	
Mood disorders	

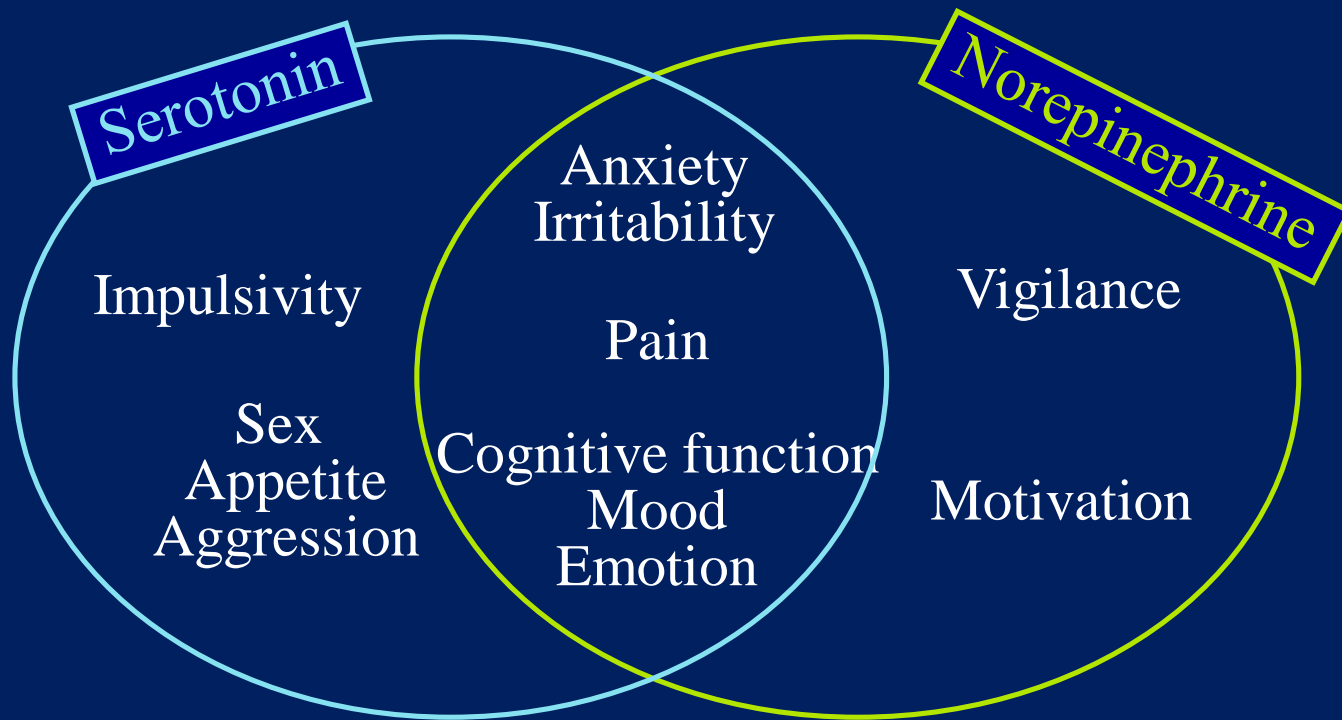
Etiological formulations I.

- **Genetic** - aggregation within family
SERT and other polymorphisms
Neurobiological -limbic system /amygdala,
prefrontal cortex, hippocampus (fear
responses, „limbic alert”) volumetric
measures
- **Biochemical** - role of 5HT and NE system,
GABA and glutaminergic system,
- **Dysregulation** of CRF and HPA axis -
excessive secretion of cortisol („stress
hormone”)

Serotonin and Norepinephrine pathways



Serotonin and Norepinephrine: Effect on Symptoms



➡ Dual action agents may provide the broadest spectrum of therapeutic effect across the full range of emotional and physical symptoms of depression

Etiological Formulation II.

- **Physical, psychiatric** disorders with anxiety, substance induced, or based on „general medical condition...”
- **Psychosocial** - learned response, conditioned in anxiety provoking situation
- **Psychoanalytic, dynamic** - related to unresolved unconscious conflicts

Etiology III. - Psychological approaches

1. Psychoanalytic

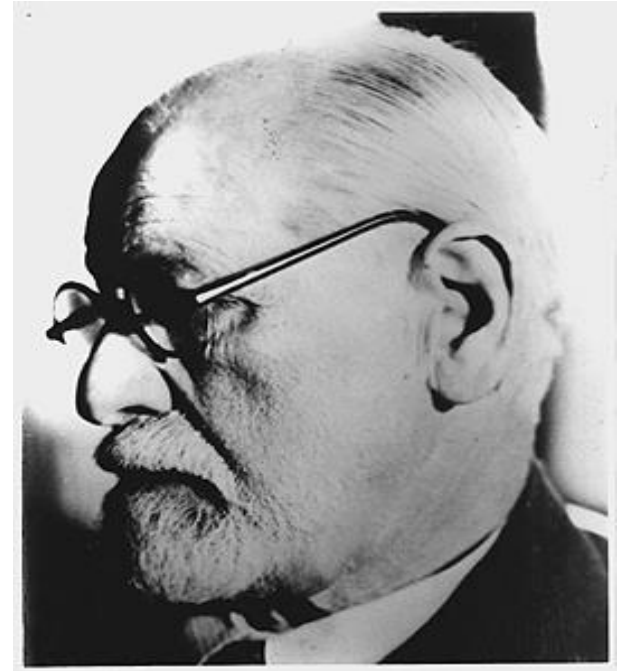
Freud - signal anxiety
developmental hierarchy:

- desintegrational anxiety
- paranoid
- separation
- castration “
- superego “

2. behavioral

- Learning theory
- Social learning
- Cognitive approach

3. Existential



ANXIETY DISORDERS

- panic disorder
- agoraphobia
- specific and social phobia
- obsessive-compulsive disorder
posttraumatic stress disorder
- acute stress disorder
- generalised anxiety disorder

- **PANIC DISORDER** with/without **AGORAPHOBIA**
 recurrent panic attacks (discrete periods of intense fear and discomfort)
 symptoms of panic attack: dispnoea; dizziness; palpitation;
 trembling/shaking; sweating; choking; nausea;
 depersonalization/derealization; paresthesias; chest pain/discomfort;
 fear of dying; fear of going crazy; - mitral valve prolapse may be
- **AGORAPHOBIA**
 fear of being in places or situations from which escape might not be available (**outside the** home alone, in a crowd, open place, bridge, in a bus, train or car)
- **SOCIAL PHOBIA**
 a persistent fear of one or more social situations (speaking in public; eating in front of others; unable to urinate in public lavatory; saying foolish things)
- **SIMPLE PHOBIA**
 specific phobias: animals (dogs), snakes, insects, mice, blood, closed place (claustrophobia), heights (agoraphobia) knife (aichmophobia), air travel
 such stimuli provoke anxiety response and avoidance behavior

OBSESSIVE-COMPULSIVE DISORDER

- recurrent obsessions and compulsions
 - ideas, thoughts, impulses
 - or actions (violence, contamination, doubt; stereotyped acts: hand-washing, counting, checking, touching)

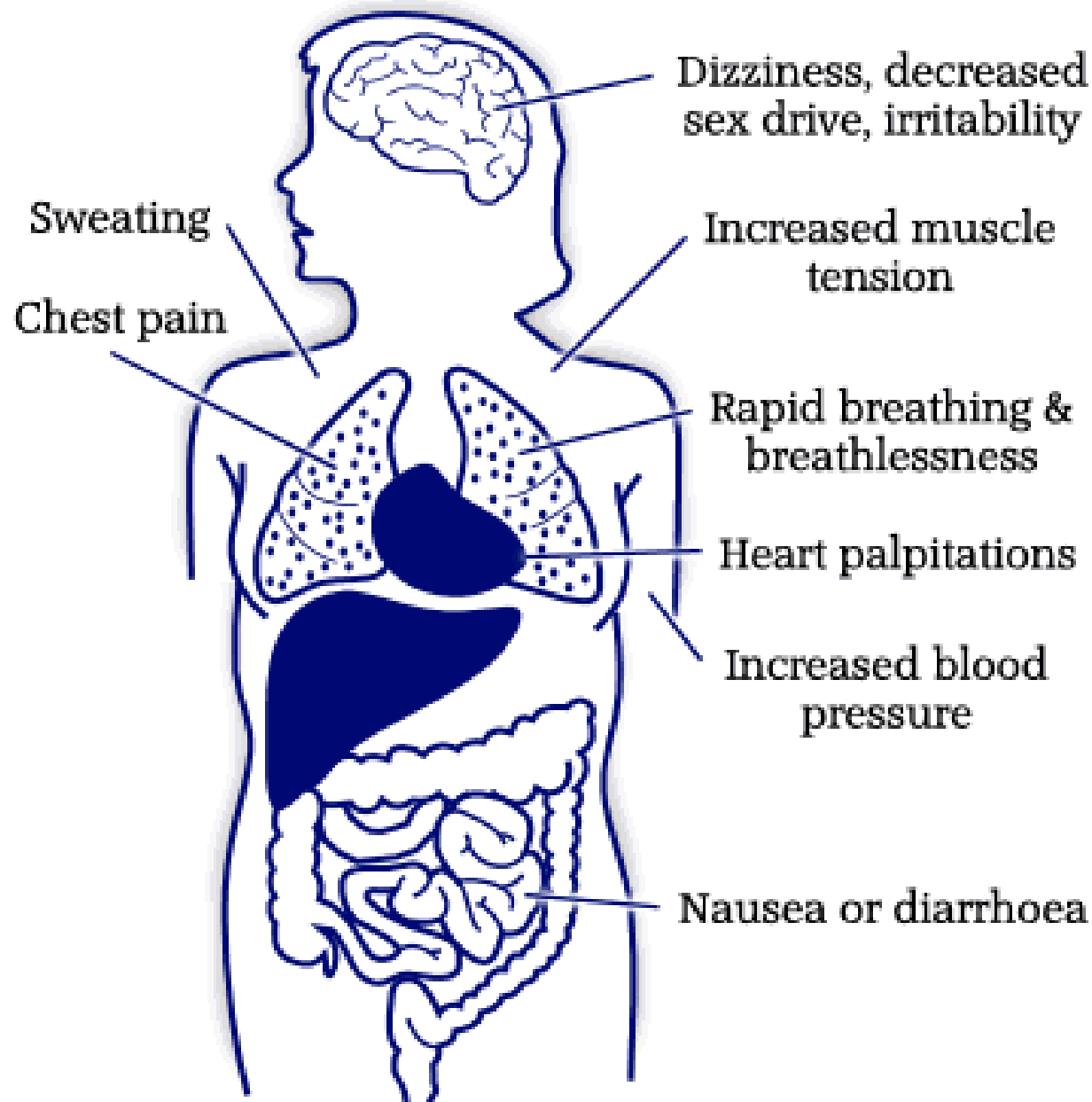
POST-TRAUMATIC STRESS DISORDER

- unusual, uncommon stressors (serious threat to life or physical integrity; sudden destruction of home, serious accident, violence, torture, kidnapping etc);
- natural disasters /= traumatic events/ provoke fear, terror, helplessness, re-experience and avoidance of the event

GENERALIZED ANXIETY DISORDER

- unrealistic or excessive anxiety and worry about two or more life circumstances / possible misfortune to one's child; worry about financial problems for no good reason

Physical Effects of Anxiety Disorders



Prevalence of the anxiety disorders (NCS)

	<i>LIFETIME PREVALENCE</i>	<i>1 MONTH PREVALENCE</i>
PANIC USA	3,5 % 5 – 2%	2,3% 3 - 1%
AGORAPHOBIA USA	1-5% 7 – 3,5%	5,3% 4 – 1,7%
SOC. PHOBIA USA	13,3% 15 – 11%	7,9% 9 – 6,6%
GAD USA	5,1% 6,6 – 3,6%	3,1% 4 – 2%

Table 10–2. DSM-IV criteria for a panic attack

A discrete period of intense fear or discomfort, in which four (or more) of the following symptoms developed abruptly and reached a peak within 10 minutes:

1. Palpitations, pounding heart, or accelerated heart rate
 2. Sweating
 3. Trembling or shaking
 4. Sensations of shortness of breath or smothering
 5. Feeling of choking
 6. Chest pain or discomfort
 7. Nausea or abdominal distress
 8. Feeling dizzy, unsteady, light-headed, or faint
 9. Derealization (feelings of unreality) or depersonalization (being detached from oneself)
 10. Fear of losing control or going crazy
 11. Fear of dying
 12. Paresthesias (numbness or tingling sensations)
 13. Chills or hot flushes
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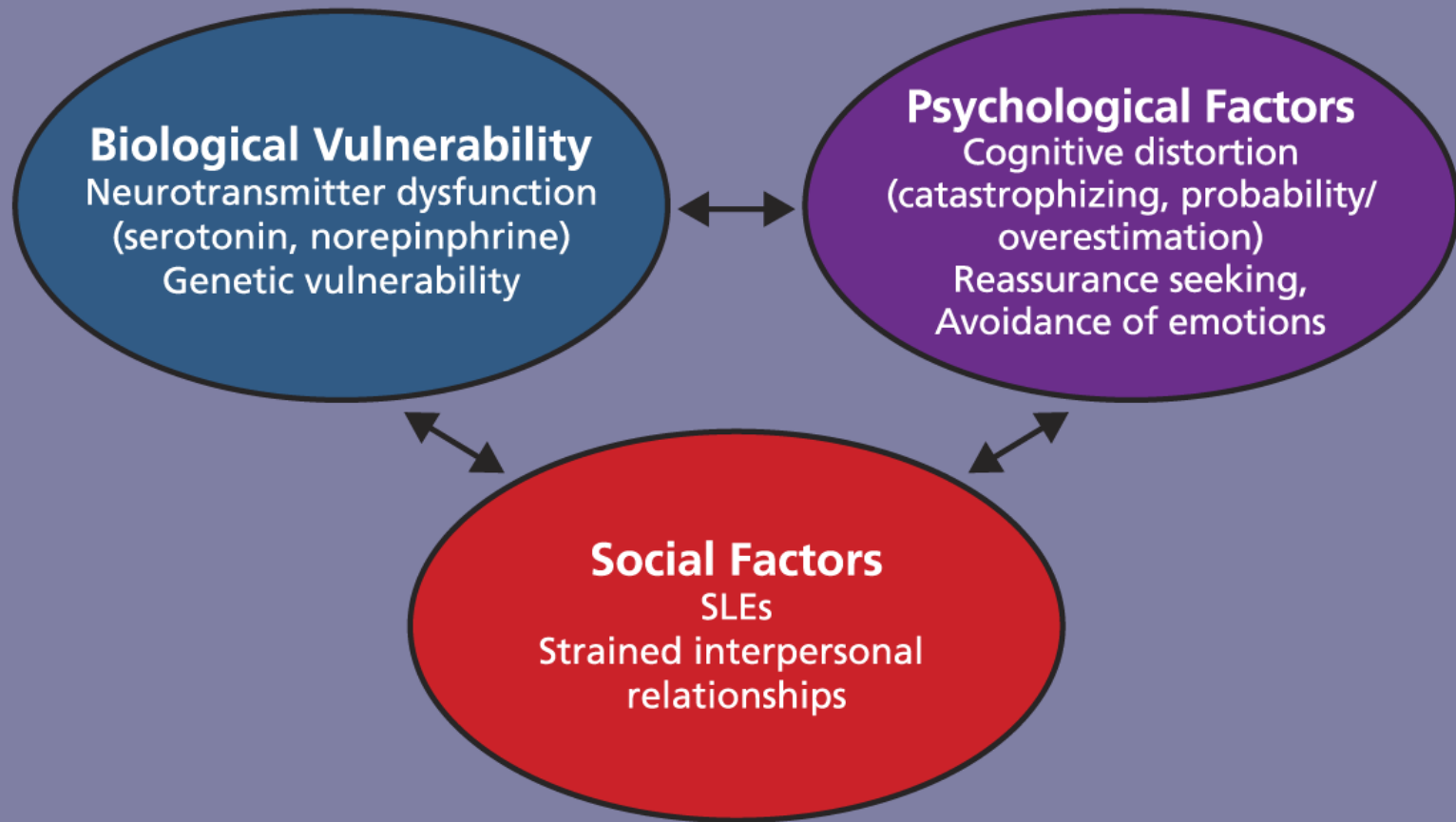
- **1. Psychological symptoms:**
- **apprehension, worry, fear, anticipation of misfortune sense of doom, or panic hypervigilance, irritability**
- **fatigue**
- **insomnia**
- **derealization, depersonalization**
- **difficulty concentrating**

- **2. Somatic complaints, Physical signs :**
- **Headache dizziness and lightheadedness**
- **palpitation and chest pain**
- **upset stomach and diarrhea**
- **frequent urination**
- **lump in the throat**
- **motor tension and restlessness**
- **shortness of breath**
- **Paresthesias, dry mouth**
- **cool, clammy skin**
- **tachycardia and arrhythmias**
- **flushing and pallor**
- **trembling, easy startle, and fidgeting, hyperreflexia**

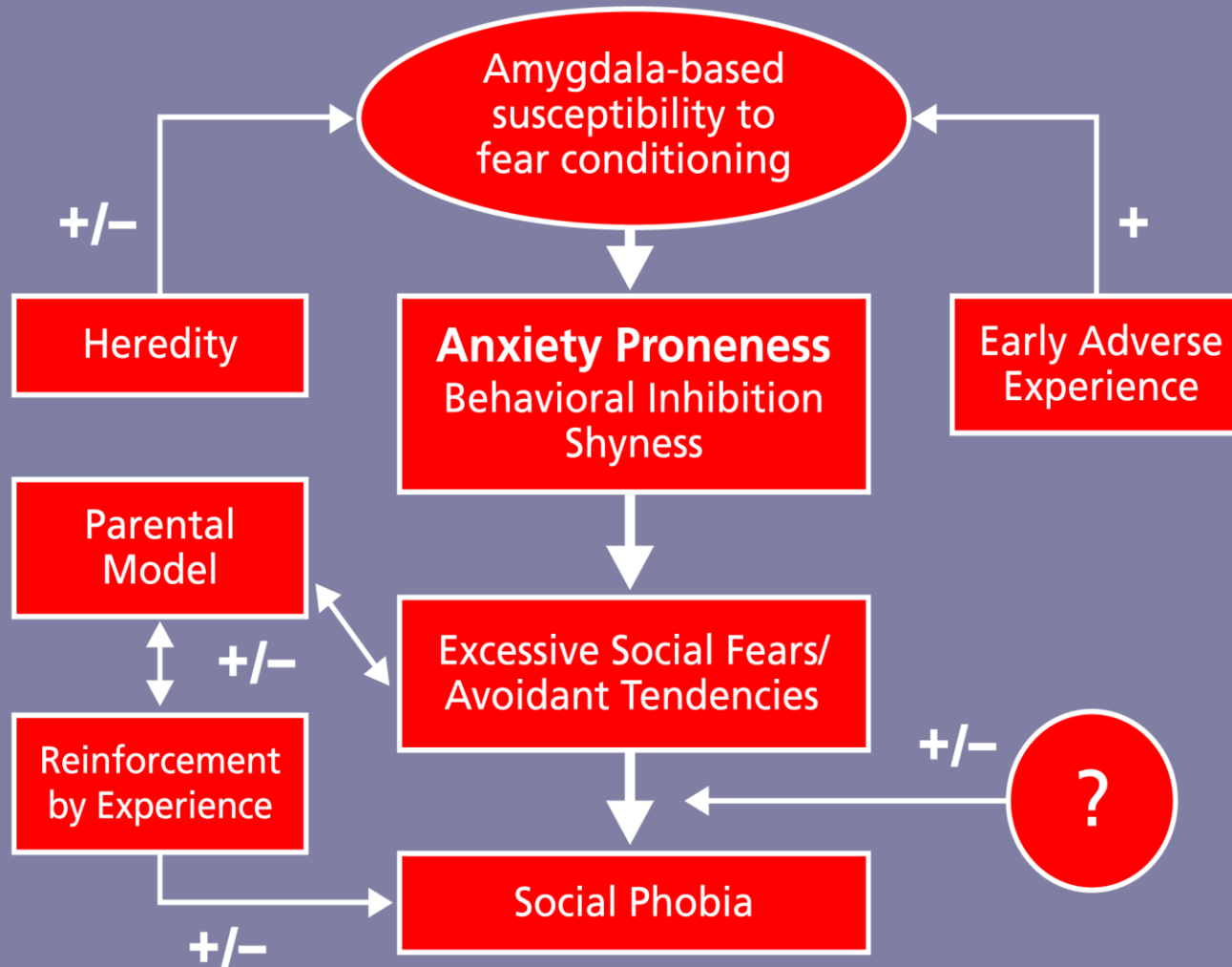
Symptoms of GAD

- **Excessive, ongoing worry and tension**
- **An unrealistic view of problems**
- **Restlessness or a feeling of being "edgy"**
- **Irritability**
- **Muscle tension**
- **Headaches**
- **Sweating**
- **Difficulty concentrating**
- **Nausea**
- **The need to go to the bathroom frequently**
- **Tiredness**
- **Trouble falling or staying asleep**
- **Trembling**
- **Being easily startled**

Biopsychosocial Model of Generalized Anxiety Disorder

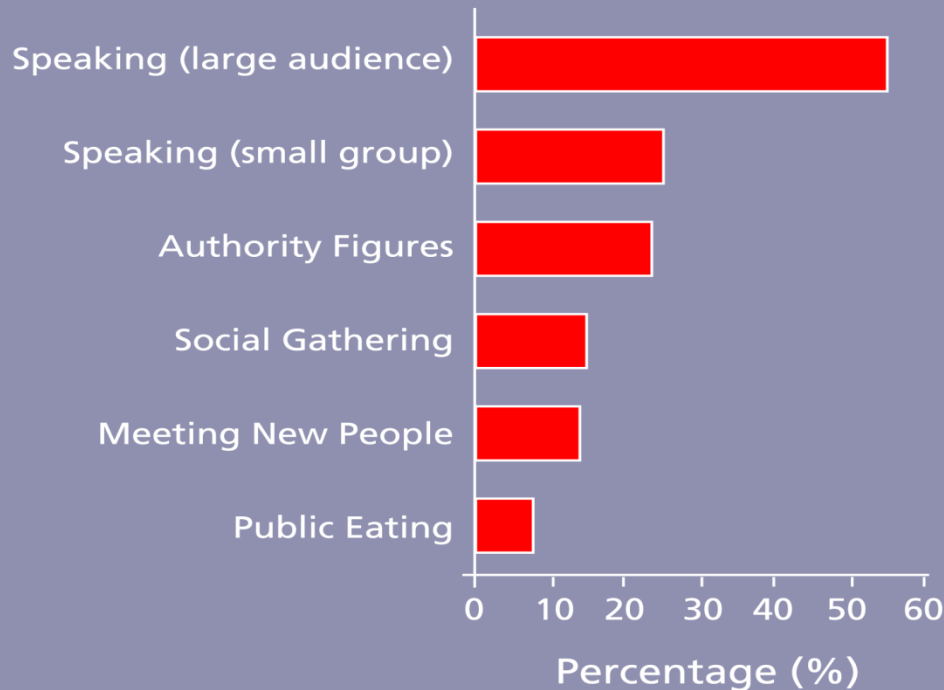


Impact of Genetics and Environment on Social Anxiety Disorder



Slide 1

Common Fears of Patients With Social Anxiety Disorder



- Incidence of social anxiety disorder in this population was 7%
- Most common fears expressed by social phobics
 - Public speaking
 - Speaking with strangers
 - Meeting new people
 - Dealing with authority figures
 - Eating or writing in public

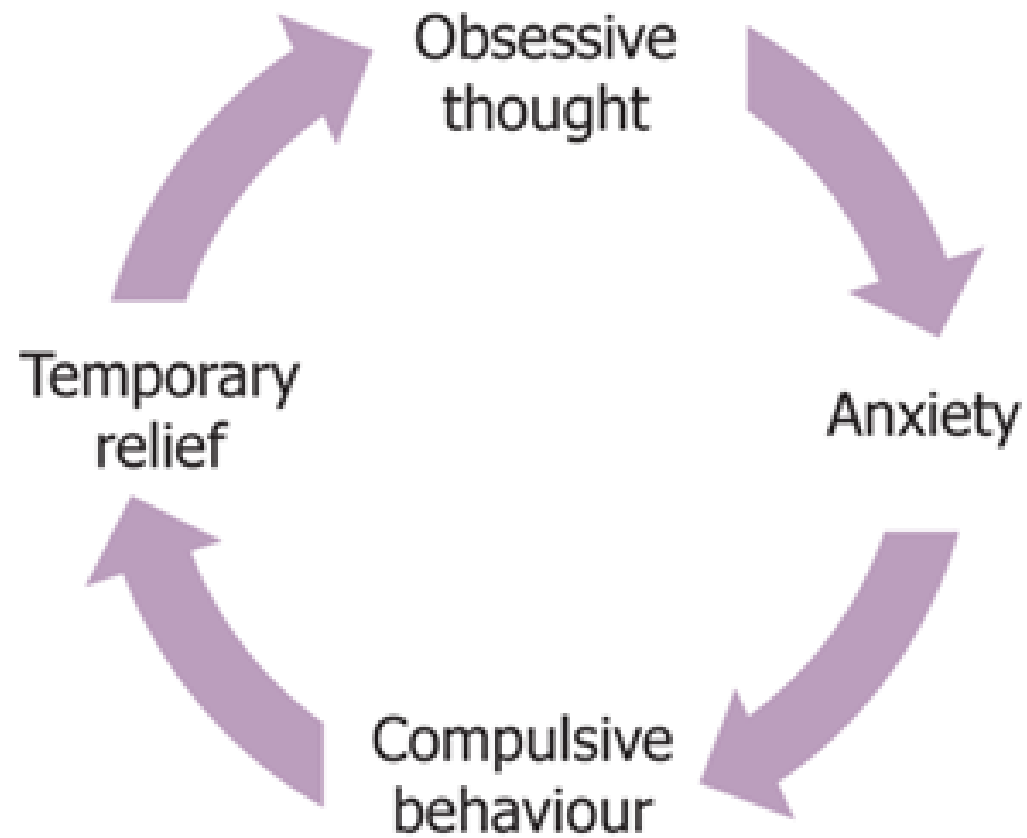


Table 1 - DSM-IV diagnostic criteria for OCD

The person exhibits either obsessions or compulsions

The person recognizes that the obsessions or compulsions are excessive or unreasonable*

The obsessions or compulsions cause marked distress, are time consuming (take more than 1 hour a day), or significantly interfere with the person's normal routine, occupational/academic functioning

If another axis I disorder is present, the content of the obsessions or compulsions is not restricted to it (e.g., eating disorder)

* Specify as poor insight; do not consider in children.



The most important symptoms of OCD

Obsessions

- ⌘ Contamination themes
- ⌘ Harm to self or others
- ⌘ Aggressive themes
- ⌘ Sexual themes
- ⌘ Scrupulosity/religiosity
- ⌘ Forbidden thoughts
- ⌘ Symmetry urges
- ⌘ Need to tell, ask, confess

Compulsions

- ⌘ Washing or cleaning
- ⌘ Repeating
- ⌘ Checking
- ⌘ Touching
- ⌘ Counting
- ⌘ Ordering/arranging
- ⌘ Hoarding
- ⌘ Praying

The main steps of diagnosis

- Anamnesis
 - Somatic disorders and treatment
 - Mental disorders – comorbidity: drugabuse, personality dis., affective dis., suicidality
- Careful physical and neurological examination – laboratory tests
- Psychiatric examination – interview – anxiety in the focus
- Development of the personality – coping mechanism, life circumstances

Diagnostic Tests

Evaluate the severity and follow up the treatment

Self ratings

- Beck Anxiety questionnaire
- Spielberger, State and Trait q.
- Yale Brown OCD scale /Y BOCS/

Rating by an evaluator

- Hamilton Anxiety Scale

General principles of therapy

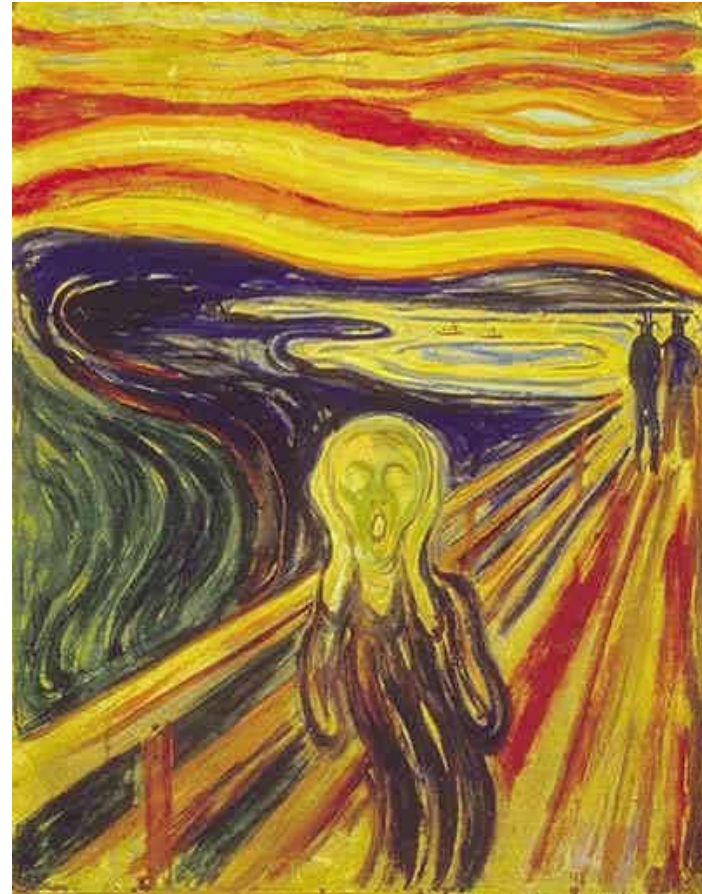
- Generally outpatient therapy
- Psychiatric hospitalisation needed:
 - Serious functional damage (comorbidity!)
 - Behavioural disturbance
 - Suicide ideation
- Pharmaco- and psychotherapy
- Follow up care
- Relapse prevention
- GP treatment - psychiatric consultation

The neurobiological basis of the effective treatment

- GABA-A RECEPTOR COMPLEX (omega 1-6)
 - benzodiazepine-agonists decrease anxiety
- SEROTONERGIC SYSTEM (dorsal raphe nucleus) 5-HT antagonists decrease anxiety
- NORADRENERGIC SYSTEM (locus coeruleus)
 - autoreceptor stimulants (clonidine) decrease anxiety,
- Dopaminergic system's role is also likely

The viewpoints of ideal anxiolytic medication?

- Effective in broad spectrum
- No sedation
- No effect on cognitive functions
- No tolerance - dependence
- No withdrawal symptoms and rebound anxiety
- Overdose is not life threatening



ANXIETY DISORDERS DRUG THERAPY

- BENZODIAZEPINES
- SEROTONERGIC (5-HT_{1A}) RECEPTOR PARTIAL AGONISTS
- ANTIDEPRESSANTS
- NORADRENERGIC DRUGS (beta-receptor blockers)

Modern antidepressants

α_2 - antagonist + 5 HT₂ antagonism:

- Remeron

RIMA:

- moclobemide

5-HT reuptake inhibition + 5-HT₂ antagonism:

- trazodone
- nefazodone

SSRI's:

- paroxetine
- fluoxetine
- sertraline
- Citalopram and escitalopram
- fluvoxamine

SNRI's, dual action

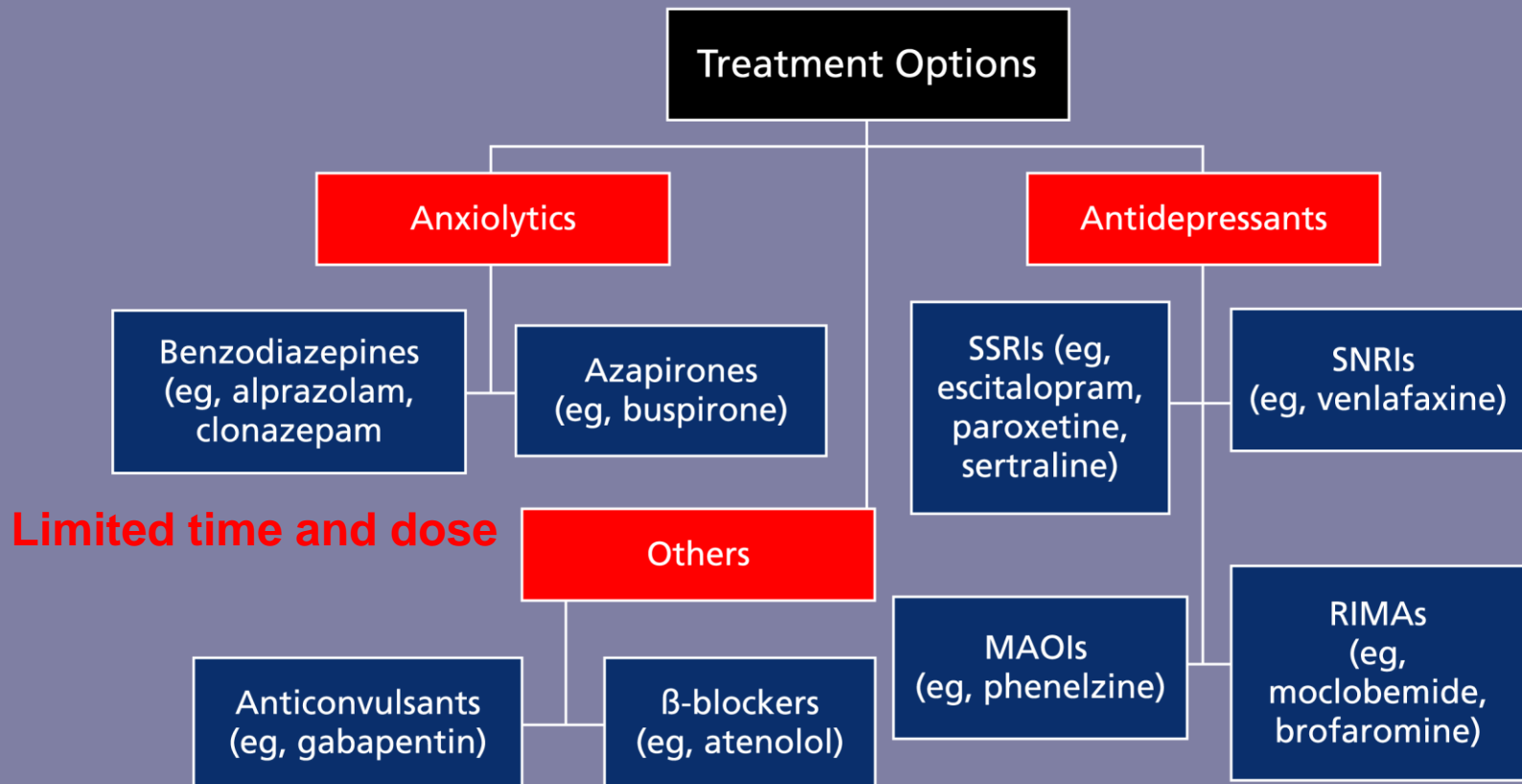
venlafaxin, reboxetine, duloxetine

Table 2. Summary of SSRIs and SNRIs

Generic Name	Brand Name	Approved Use
SSRIs		
Citalopram	Celexa	Depression
Escitalopram	Lexapro	Depression, GAD
Fluoxetine	Prozac, Sarafem, Selfemra	Bulimia nervosa, depression, OCD, panic disorder, PMDD
Fluvoxamine	Luvox	OCD, social phobia
Paroxetine	Paxil, Pexeva	Depression, GAD, OCD, panic disorder, PTSD, PMDD, social phobia
Sertraline	Zoloft	Depression, OCD, panic disorder, PTSD, PMDD, social phobia
Vilazodone	Viibryd	MDD
SNRIs		
Duloxetine	Cymbalta	Depression, diabetic neuropathy, fibromyalgia, GAD, musculo-skeletal pain, osteoarthritis
Venlafaxine	Effexor	Depression, GAD, panic disorder, social phobia
Desvenlafaxine	Pristiq	Depression
Levomilnacipran	Fetzima	Depression

GAD: generalized anxiety disorder; OCD: obsessive-compulsive disorder; MDD: major depressive disorder; PMDD: premenstrual dysphoric disorder; PTSD: posttraumatic stress disorder; SNRI: serotonin-

Pharmacologic Treatments for Social Anxiety Disorder



Blanco C, Raza MS, Schneir FR, et al. *Int J Neuropsychopharmacol.* 2003;6:427-442.

PSYCHOTHERAPEUTIC THERAPIES

- Supportive psychotherapy
- Cognitive and behavioural therapies - (systematic desensitisation) correcting false schemas and cognitive distortions
- Relaxation-meditation
- Psychodynamic psychotherapies - /working through losses, traumas, repressed aggression/
- Group-therapies, family-therapies

- In general, anxiety illnesses, particularly those that are persistent and recurrent, will require medication
- Psychotherapies that research has shown helpful for some forms of anxiety are interpersonal and cognitive/behavioral therapies.
- Interpersonal/family therapists focus on the patient's disturbed personal relationships that both cause and exacerbate (or increase) the anxiety
- Cognitive/behavioral therapists help patients change the negative styles of thinking and behaving often associated with anxiety
- Psychodynamic therapies which are sometimes used to treat anxious persons, focus on resolving the patient's conflicted feelings

Recommendations for treatment of anxiety disorders

1. Mild cases of panic may respond to behavioral interventions, but many patients will need medication (e.g., tricyclic antidepressants [TCAs], monoamine oxidase inhibitors [MAOIs], alprazolam).
2. The agoraphobic patient should be gently encouraged to get out and explore the world.
 - Progress will not occur unless the phobic patient confronts the feared places or situations.
3. Behavioral techniques (i.e., exposure, flooding, desensitization) will help most persons with specific and social phobias.
 - Some people with social phobias respond well to medication (e.g., TCAs, MAOIs, serotonin reuptake inhibitors, alprazolam).
4. Generalized anxiety may respond to simple behavioral techniques (e.g., relaxation training), but many patients will need medication (e.g., benzodiazepines).
 - Be sure that the benzodiazepine is prescribed for a limited time only (e.g., weeks or months).
5. Posttraumatic stress disorder tends to be chronic, but many patients will benefit from the support available in group therapy.
 - Group therapy has become especially popular with Vietnam veterans, and most veteran organizations can offer help in finding a group.