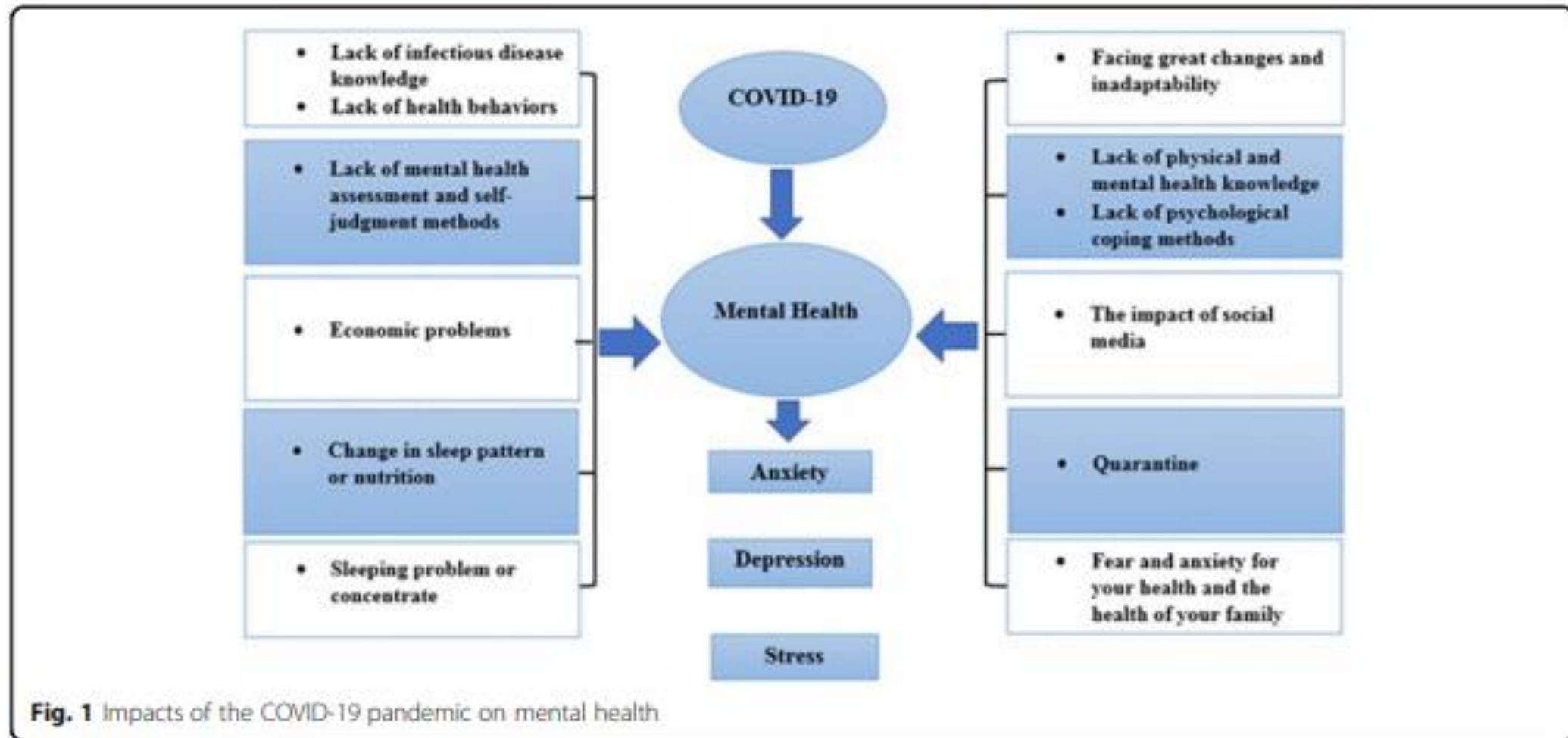


بِناَمِ خُدا

# رویکرد تشخیصی و درمانی به اختلالات افسردگی و اضطرابی و اختلالات خواب در کوید ۱۹

دکتر فریده رنجبران



**Fig. 1** Impacts of the COVID-19 pandemic on mental health

# Adverse psychological outcomes of individuals in quarantine


- anger, anxiety, boredom, confusion, fear, depression, emotional exhaustion, frustration, irritability, and stress.
- PTSD
- Other adverse outcomes included avoidance behaviors (eg, avoiding crowded or public places), detachment from others
- subthreshold symptoms of alcohol use disorder
- excessive preoccupation with distressing somatic symptoms
- Stigma
- domestic violence
- suicidal ideation and behavior

RESEARCH

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# Prevalence of stress, anxiety, depression among the general population during the COVID-19 pandemic: a systematic review and meta-analysis



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The prevalence of stress in 5 of the studies with a sample size of 9074 was 29.6%

The prevalence of anxiety in 17 studies with a sample size of 63,439 was obtained as 31.9%

the prevalence of depression in 14 studies with a sample size of 44,531 was 33.7%



**prevalence of stress, anxiety, depression among the general population  
during the COVID-19 pandemic in different continents**

The highest prevalence of anxiety in Asia is 32.9 (95% CI: 28.2–37.9),  
the highest prevalence of stress in Europe is 31.9 (95% CI: 23.1–42.2),  
and the highest prevalence of depression in Asia is 35.3

- Anxiety, when above normal, weakens body's immune system
- Research shows that people who follow COVID-19 news the most, experience more anxiety
- Misinformation and fabricated reports about COVID-19 can exacerbate depressive symptoms in the general population
- People who have improved and the progress of medications and vaccines, can reduce anxiety levels
- The results of epidemiological studies show that women are at a higher risk of depression
- In recent studies, the prevalence of anxiety and depression and stress during COVID-19 pandemic is shown to be higher in women than in men

# Symptoms of depression

- Sadness
- Tearfulness
- Loss of interest
- pleasureGuilt
- Trouble sleeping
- Decrease or increase appetite
- Low energy
- Diminished sex drive
- Poor concentration



# Probable etiology

- Living an urban area
- Reduce exercise
- Dissatisfaction with sleep
- Hypoxia
- Smoking
- Fear of covid 19 reinfection
- Hospitalization
- Past medical disease
- Self medication
- Persistent symptoms of covid 19
- Stigma

# May affect several groups

- Patients with COVID 19
  - Disease related stresses
  - Neuropsychiatric sequels of infection
- Medical staff who care for patients (outpatient or admitted)
- Patients with prior psychiatric diagnosis:
  - Exacerbation of Psychiatric illnesses (panic disorder, GAD, OCD)
  - Increased risk of infection for major psychiatric disorders (schizophrenia, Bipolar)
- General Population

# Health care workers

- Anxiety – 12 to 20 percent
- Depression – 15 to 25 percent
- Insomnia – 8 percent
- Traumatic distress – 35 to 49 percent
- Need for psychological support – 47 percent

# Risk & Protective Factors

- **Risk factors:**
  - increased contact with affected patients
  - a prior history of psychiatric or medical diseases
  - perceived lack of organizational support or social stigma
- **Protective factors**
  - access to personal protective equipment & trust in the institution's infection control
  - access to psychiatric interventions
  - having supportive peers
  - receiving clear communication from supervisors

# Psychiatric illnesses in infected patients

- during acute infection, approximately 20 to 40 percent of patients manifested neuropsychiatric symptoms
  - insomnia – 42 percent
  - impaired attention or concentration – 38 percent
  - Anxiety – 36 percent
  - Memory impairment – 34 percent
  - Depressed mood – 33 percent
  - Confusion – 28 percent
  - Altered consciousness – 21 percent
- Depression, anxiety, fatigue, and post-traumatic stress disorder in the post-illness stage of previous coronavirus epidemics (few data yet on COVID-19).

# What does stress mean?

- Stress is a normal physiological response to an abnormal situation. As such, it is part and parcel of our lives. It enables our body to adapt to the multiplicity of positive and negative events that we experience, like a birth, marriage, loss of employment, etc. Stress comes and goes on its own, depending of what factors are involved. For example, if you feel stressed on the job but less so at home in the evening or on the weekend, we could deduce that the stressors are work-related.

# What does anxiety mean?

- Contrary to fear, which is a response to a well-defined and very real threat, anxiety is a response to a vague or unknown threat. Anxiety manifests itself when we believe that a dangerous or unfortunate event may take place and are expecting it. Everyone experiences anxiety at their own individual degree and intensity. How the anticipated event is perceived will greatly influence the intensity of the anxiety experience.

### **Physical symptoms:**

- Headaches, neck tension, gastrointestinal problems, etc.
- Sleep problems
- Lower appetite
- Lower energy, fatigue
- Etc.

### **Psychological and emotional symptoms:**

- Virus-related worries and insecurity
- Feelings of being overwhelmed by events, powerlessness
- Self-verbalization that does not always reflect reality
- Negative vision of things or daily events
- Feelings of discouragement, insecurity, sadness, anger, etc.

### **Behavioural symptoms:**

- Difficulty in concentrating
- Irritability, aggression
- Crying
- Withdrawal, insularity
- Difficulty in taking decisions
- Increased use of alcohol, drugs and/or medication
- Etc.



# Mechanism of depression and anxiety in patients with covid 19

- Psychological mechanism
  - quarantine
  - stress

Central mechanism Covid 19 inflammation Hypocamp  
disorderMalnutrition

# Factors Associated with Sleep Disturbance in COVID-19 Patients

- Physiologic Factors
  - Neuronal System Injury
- Psychological Factors
- Environmental Factors

# Sleep disorder

- The collective term sleep disorder refers to conditions that affect sleep quality, timing, or duration and impact a person's ability to properly function while they are awake. These disorders can contribute to other medical problems, and some may also be symptoms for underlying mental health issues



# Who's at Risk for Coronasomnia?

- Patients with COVID-19
- Frontline workers
- Unpaid caregivers
- Essential workers
- Women
- Young adults
- People of color

# Causes Coronasomnia?

- Increased Stress
- Loss of Daily Routines
- Increased Media Consumption

# Cause of sleep disorder

- Neuroinflammation
- Blood - Brain Barrier Disruption

# Insomnia drug therapy in covid 19 patients

- Important factors :
  - interaction of hypnotic medication with common medication in the treatment of covid 19
  - respiratory depression by hypnotic medication
  - liver damage by covid 19 or drug therapy



# Medication for improve sleep quality

- **Benzodiazepines**
- **Z drug**

# Drug interactions

- Drug interactions are major challenge in comorbidity of psychiatric disorders and COVID-19 infection
- QTc prolongation is a major concern while using antiviral medications and/or Chloroquine/hydroxychloroquine in combination with many psychotropics. Cardiac monitoring especially in high risk patients is highly recommended.
- Concomitant use of SSRIs with antiviral medications and/or Chloroquine/hydroxychloroquine increase the risk of hypoglycemia.
- Concomitant use of pimozide or midazolam with antiviral medications is contraindicated.

## Table 1. Summary of drug-drug interaction between psychotropics and pharmacological treatments used for COVID-19

Psychotropic Medication	Lopinavir/Ritonavir	Chloroquine/ hydroxychloroquine	Implication	
	↑ risk for serotonin syndrome and hypoglycemia		Caution in patients with diabetes	
	↑ Ritonavir via CYP2D6 and CYP3A4		Risk of hypoglycemia	
	↑ Sertraline via CYP3A4		Risk of hypoglycemia	
SSRIs				
			↑ QTc	Cardiac monitoring especially in high-risk patients
			↑ QTc	Cardiac monitoring especially in high-risk patients
		↓ Paroxetine via CYP2D6 and protein binding displacement		Risk of hypoglycemia
	↑ Ritonavir via CYP3A4		Risk of hypoglycemia	

Antidepressants	SNRIs		↑ QTc	Cardiac monitoring especially in high-risk patients
	TCA	↑ TCAs via CYP2D6	↑ QTc	dose reduction is not necessary, monitor TCA side effects
	MAOIs		Risk of hypoglycemia	Caution in patients with diabetes
	Bupropion	↓ Bupropion via CYP2B6		Monitor clinical effectiveness of bupropion
	Trazodone	↑ Trazodone via CYP3A4	↑ QTc	Cardiac monitoring especially in high-risk patients Sedation may also occur
	Mirtazapine	↑ Mirtazapine via CYP3A4		Use the lowest efficient dose
	Vortioxetine	↑ Vortioxetine via CYP2D6		Not clinically significant
	Vilazodone	↑ Vilazodone via CYP3A4		Dose reduction by 50% or using alternative is suggested
Mood stabilizers	Valproic acid	↓ Valproate ↑ Lopinavir		Monitor for lopinavir toxicity and virologic response
	Lamotrigine	↓ Lam via UGTs		A dose increase is recommended
	Carbamazepine	↓ Lopinavir	↑ Chloroquine	
	Lithium			

Psychotropic Medication		Lopinavir/Ritonavir	Chloroquine/ hydroxychloroquine	Implication	
Antipsychotics	Second-generation antipsychotics	Risperidone	↑ Risperidone via CYP3A4 and 2D6	Dose reduction	
		Aripiprazole	↑ Aripiprazole via CYP3A4 and 2D6		
		Olanzapine	↓ Olanzapine via CYP1A2	↑QTc	
		Quetiapine	↑ Quetiapine via CYP3A4	↑QTc	Cardiac monitoring
		Ziprasidone	↑ Que via CYP3A4		
		Clozapine	Additive metabolic toxicities	↑QTc	Consider alternative agents
	First-generation antipsychotics	Chlorpromazine		↑QTc	
		Pimozide	↑ Pimozide via CYP3A4	↑QTc	Contraindicated
Anxiolytics	Benzodiazepines	Midazolam	↑ Midazolam via CYP3A4	Contraindicated	
		Diazepam	↑ Diazepam via 3A4	Dose reduction	
		Alprazolam	↑ Alprazolam via 3A4	Monitor sedation and dose reduction	
		Triazolam	↓ Triazolam via 3A4		
	Zolpidem	↑ Zolpidem via 3A4		Not clinically significant	
	Buspirone	↑ Buspirone via 3A4		Dose adjustment is not usually necessary	

## Summary of drug-drug interaction between psychotropics and pharmacological treatments used for COVID-19

Psychotropic Medication		Lopinavir/Ritonavir	Chloroquine/hydroxychloroquine	Implication
		increased risk for serotonin syndrome and hypoglycemia		Caution in patients with diabetes
	Fluoxetine	Increased Ritonavir via CYP2D6 and CYP3A4	Risk of hypoglycemia	
	Sertraline	Increased Sertraline via CYP3A4	Risk of hypoglycemia	
SSRIs	Citalopram		increased QTc	Cardiac monitoring especially in high-risk patients
	Escitalopram		increased QTc	Cardiac monitoring especially in high-risk patients
	Paroxetine	decreased Paroxetine via CYP2D6 and protein binding displacement	Risk of hypoglycemia	
	Fluvoxamine	increased Ritonavir via CYP3A4	Risk of hypoglycemia	

		Lopinavir/Ritonavir	Chloroquine/ hydroxychloroquine	Implication
Antidepressants	SNRIS		Increased TCAs via CYP2D6 QTc	Cardiac monitoring especially in high-risk patients
	TCAS	Increased TCAs via CYP2D6	Increased TCAs via CYP2D6 QTc	dose reduction is not necessary, monitor TCA side effects
	MAOIs		Risk of hypoglycemia	Caution in patients with diabetes
	Bupropion	Decreased Bupropion via CYP2B6		Monitor clinical effectiveness of bupropion
	Trazodone	Increased Trazodone via CYP3A4	Increased TCAs via CYP2D6 QTc	Cardiac monitoring especially in high-risk patients Sedation may also occur
	Mirtazapine	Increased Mirtazapine via CYP3A4		Use the lowest efficient dose
	Vortioxetine	Increased Vortioxetine via CYP2D6		Not clinically significant
	Vilazodone	Increased Vilazodone via CYP3A4		Dose reduction by 50% or using and alternative is suggested

		Lopinavir/Ritonavir	Chloroquine/ hydroxychloroquine	Implication
Mood stabilizers	Valproic acid	Decreased Valproate Increased Lopinavir		Monitor for lopinavir toxicity and virologic response
	Lamotrigine	Decreased Lam via UGTs		A dose increase is recommended
	Carbamazepine	Decreased Lopinavir	Increased Chloroquine	
	Lithium			



			Lopinavir/Ritonavir	Chloroquine/ hydroxychloroquine	Implication
Antipsychotics	Second-generation antipsychotics	Risperidone	Increased Risperidone via CYP3A4 and 2D6		
		Aripiprazole	Increased Aripiprazole via CYP3A4 and 2D6		Dose reduction
		Olanzapine	Decreased Olanzapine via CYP1A2	Increased QTc	
		Quetiapine	Increased Quetiapine via CYP3A4	Increased QTc	Cardiac monitoring
	Ziprasidone	Increased Que via CYP3A4	Increased QTc		
	Clozapine	Additive metabolic toxicities	Increased QTc	Consider alternative agents	
	First-generation antipsychotic	Chlorpromazine		Increased QTc	
		Pimozide	Increased Pimozide via CYP3A4	Increased QTc	Contraindicated

			Lopinavir/Ritonavir	Chloroquine/ hydroxychloroquine	Implication
Anxiolytics	Benzodiazepines	Midazolam	Increased Midazolam via CYP <sub>3A4</sub>		Contraindicated
		Diazepam	Increased Diazepam via 3A <sub>4</sub>		Dose reduction
		Alprazolam	Increased Alprazolam via 3A <sub>4</sub>		Monitor sedation and dose reduction
		Triazolam	Decreased Triazolam via 3A <sub>4</sub>		
	Zolpidem		Increased Zolpidem via 3A <sub>4</sub>		Not clinically significant
	Buspirone		Increased Buspirone via 3A <sub>4</sub>		Dose adjustment is not usually necessary

با تشکر از توجه شما

