Panic disorder in general medical practice- A narrative review

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ABSTRACT

The under-or misdiagnosis, and symptomatic treatment of the panic disorder (PD), despite high prevalent medical illness, is common among non-psychiatric physicians. The non-psychiatrist physician's role is vital in the care of PD as most patients initially approach general medical settings for medical help, including primary care. However, a significant proportion is undiagnosed and undergoes either unnecessary investigation, misdiagnosed, or mismanaged even among post-Coronary Artery Bypass Grafting patients, which profoundly affects the patients functioning and quality of life. This article aims to provide overviews of relevant epidemiological aspects, presenting features across medical specialties with respective diagnostic dilemmas, assessment, and management of the PD in their general medical settings, including emergency visits. Apart from psychiatrists, this will also assist non-psychiatrist physicians across all medical specialties, including general practitioners, to understand, identify, and provide the first line evidence-based pharmacotherapy and address the unmet need of patients with PD in their day-to-day busy clinical practice. This paper also provides a referral guide for non-psychiatrist physicians to refer to psychiatrists for further management after their first-line management.

Keywords: Clinical pathways, general medical practice, panic disorder, treatment

Introduction

Panic disorder (PD) is distinguished by unpredictable recurring intense episodes of extreme anxiety called 'panic attack,' leaving worries in anticipation of a further episode. About 70-80% of panic patients approach primary care physician for help; thus, their role is very crucial as they are the backbone of any healthcare.[1] There is a need to sensitise them as prevalence of panic attack is very high (13.2%)[2] so does their encounter with general physician.^[3] In scarcity of psychiatrists, the major issues that needs attention are high rate of misdiagnosis (85.8%),[4] unnecessary expensive investigation, [5] high rate of benzodiazepine

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dependence due to symptomatic management (60%), 6 disabling chronicity of the illness, [7,8] and consequences of PD such as hindrance in lifestyle, education dropout, unemployment, depression, substance use, and suicidality.

Methodology Search

There is insufficient published literature from India to arrive for a clear guideline in Indian context. The author performed a narrative review to identify the types of available data about PD in general medical practice suitable for an Indian context. This review was chosen as there is no literature to address this topic for all medical doctors ranging from specialties to general practitioners in India. Since this was a narrative review, there were no restrictions for publication type or date limit and quality appraisal for inclusion.

The literature search is done on the electronic database (Medline, Embase/Excerpta Medica, Index Medicus, Web of Science,

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EBSCO, Search Medica, Google Scholar, and Google.) The primary search term used was "panic disorder" that combined with epidemiology, diagnosis, treatment, management, assessment, risk factors, the clinical pathway of care, investigation, presentation, associated symptoms, comorbidity, differential diagnosis, the difference in symptomatology, pathophysiology, pathogenesis, rating scale, assessment, pharmacotherapy, psychotherapy, non-pharmacotherapy, healthcare setting, prognosis, prevention.

Epidemiology

Lifetime prevalence varies from 0.5 to 4%, with peak age of onset being 25 years. It is more common in females, unemployed, divorced/separated/widowed, lower education, and low household income. [2,3,9] A 5.6-19% of primary care patients may have panic attacks, [10] and about 85.8% may subjected to misdiagnosis^[4] (known as 'functional treatment gap).^[11] Apart from primary care, the patients of PD approaches different specialists. About 8% seen by cardiologists presents with PD, [5] with 5-8% may have coronary artery disease. [12] Interestingly 10% of patients with coronary artery bypass surgery, [13] 28% of those who undergo angiography may have PD, but significant proportion remain undetected. [14] Sixteen to 17.5% PD present in emergency department present as chest pain, [13] who may remain undiagnosed or subject to investigation.^[5] Fourteen percent of dizziness in ENT clinics may have PD.[15,16] Up to 30% of PD may present with gastrointestinal symptoms and with chronic obstructive pulmonary disease^[17] and syncope.^[18] Under-recognition of PD by endocrinologists are also reported in patients with thyroiditis, diabetes, polycystic ovarian disease, and Cushing disease.^[19]

Clinical pathways and functional treatment gap in care

The most patients seek treatment in primary care setting, the emergency room and cardiology depending upon nature of the presentation and availability of clinical setting. [20] Duration of untreated illness remain shorter as symptoms mimic of life-threatening condition.^[21] It often remains undiagnosed in up to 90% if present with chest pain, [22] and often undergo invasive cardiac investigation.^[5,22] Even after diagnosis, many would keep believing of having cardiac disease, and do not seek help from psychiatrists^[7] despite having disabling symptoms^[7,23] and continue utilizing emergency room service. [24] In western country only 1/3rd would initially contact mental health professionals out of which 1/5th would be seen by psychiatrists. [25,26] The median rate of untreated PD across the world is 55.9%.[27] Thus, there appears to be urgent need of primary care physician and other medical specialists to be familiar with PD and identify early, to reduce the associated burden and duration of untreated illness. [26] Some of the common investigations carried out in patient with PD are related to cardiopulmonary, neurological, and gastrointestinal [Table 1].

Clinical features of panic disorder

As per the Diagnostic and Statistical Manual of Mental Disorders, Fifth Edition, [28] the diagnostic guidelines are given below.

Recurrent unexpected panic attacks

A panic attack is an abrupt surge of intense fear or intense discomfort that reaches a peak within minutes, and during which time four (or more) of the following symptoms occur (table -2):

B. At least one of the attacks has been followed by one month (or more) of one or both of the following:

- Persistent concern or worry about additional panic attacks or their consequences (e.g., losing control, having a heart attack, "going crazy").
- 2. A significant maladaptive change in behaviour related to the attacks (e.g., behaviors designed to avoid having panic attacks, such as avoidance of exercise or unfamiliar situations).

C. The disturbance is not attributable to the physiological effects of a substance (e.g., a drug of abuse, a medication) or another medical condition (e.g., hyperthyroidism, cardiopulmonary disorders).

D. The disturbance is not better explained by another mental disorder. (e.g., social anxiety disorder; specific phobia; obsessive-compulsive disorder; post-traumatic stress disorder).

Generally, an initial panic attack is entirely spontaneous or provoked by physical or emotional excitement or trauma. Subsequently attacks are precipitated by substance use (Nicotine, Caffeine, Alcohol, etc.), unusual eating or sleep pattern, harsh lighting etc.^[24,28]

The onset of an attack is usually abrupt with rapid progress in the severity of symptoms that peaks within 10 minutes, lasts for 20 to 30 minutes (rarely more than an hour), and may disappear quickly or gradually. Generally, the symptoms remain so severe that patients would run to seek help. Psychiatric examination often shows rumination, inability to talk correctly (e.g., stammering), memory impairment, depersonalization, derealisation, and depression. Between the attack, patients usually have anticipatory anxiety having another attack. Patients usually have anticipatory anxiety having another attack is not a psychiatric disorder. The presence of unexpected, repeated attacks is almost a pathognomonic of panic disorder especially nocturnal panic attacks.

Diagnostic markers

Clinically physiological recording of abrupt surges of arousal of heart rate that reaches a peak within minutes and subsides during the attack is a diagnostic marker. Other proposed markers are aberrant respiratory regulation, heart rate variability, Serotonin 5-TH, and noradrenergic systems activation, structural changes in the amygdala, hippocampus, etc.^[30]

Neurobiology

Critical neurotransmitters implicated in the pathophysiology of PD are Serotonin, Norepinephrine, Gama Aminobutyric Acid, Dopamine, Cholecystokinin. Different parts of the brain have been implicated in the manifestation of other components of symptoms.^[31] [Table 3]

According to neurocircuitry hypothesis of PD, thalamus transmit stimulus information to the lateral amygdala (short loop) directly or through the medial prefrontal cortex and cingulate (long loop), which in turn transmit information to the amygdala. The amygdala then transmits the processed information to the locus ceruleus, parabrachial nucleus, periaqueductal grey region, and hypothalamus, resulting in panic signs and symptoms.^[32]

Heritability of PD is estimated to be about 43%. The common genes implicated are monoamine degradative enzyme

catechol-O-methyltransferase, 5-HT transporter promoter gene, orexin-2 receptor gene, amiloride-sensitive cation channel 2 gene, etc.^[33]

Treatment gap

Treatment gap is inability to avail any intervention for illness. The proportion of eligible people not being able to access treatment at the community level is 'apparent treatment gap'; while sufferers of psychiatric disorders reaching primary care

Symptom	Symptom domain	Presenting to	Common investigations ordered (apart from routine)	Common treatment is given (apart from drugs)
Palpitations, pounding heart, tachycardia Chest pain or discomfort syncope	Cardiovascular	Cardiologist	Electrocardiogram Troponin levels, Oxygen saturation, Doppler ultrasound, Echocardiogram	Cardiac Catheterization
Sweating Chills or hot flushes, Trembling or shaking	Autonomic	Endocrinologists	Hormone levels (T3, T4, TSH) HPA axis evaluations	Hormone supplements, antihormone drugs
Shortness of breath, Smothering sensation, Hyperventilation, Stammering	Pulmonary	Chest physician/ Pulmonologist, ENT specialist	Chest X-Ray Pulmonary function test, Blood gases Laryngoscopy	Nebulization, Laryngoscopy
Paraesthesia, Dizziness or lightheadedness, unsteadiness, faintness, Impaired memory	Neurological	Neurologist	MRI Brain CT scan EEG	-
Choking Nausea Abdominal distress	Gastrointestinal	Gastroenterologists	Endoscopy	-
Depersonalization Derealization The dread of no control or 'going crazy'	Psychological	Faith healers	-	-

Table 2: The symptoms of a panic attack 1. Palpitations, pounding heart or accelerated heart rate. 2. Sweating. 3. Trembling or shaking. 4. Sensations of shortness of breath or smothering. 5. Feelings of choking. 6. Chest pain or discomfort. 7. Nausea or abdominal distress. 9. Chills or heat sensations. 10. Paraesthesias (numbness or tingling sensations). 11. Derealization (feelings of unreality) or depersonalization (being detached from oneself). 12. Fear of losing control or "going crazy." 13. Fear of dying.

Table 3: Neuroanatomy of Panic Disorder		
Structure	The implicated symptoms	
Amygdala	involved in fear and anxiety processing, regulate the behavioral and autonomic responses.	
Parabrachial Nucleus	producing a rapid breathing	
hypothalamus (lateral nucleus)	activating the sympathetic nervous system- perspiration, epigastric discomfort, dryness of mouth, palpitation, tremulousness.	
Locus Ceruleus	Norepinephrine release leading to tachycardia, elevated blood pressure, flight fight response	
Hypothalamus (paraventricular)	Hypothalamo-Pitutory-Axis activation and release of adrenocorticoids	
Periaqueductal Grey	phobic avoidance, defensive behaviors, postural freezing	
Prefrontal cortex	Processing of phobic avoidance, learned extinction processes	

8. Feeling dizzy, unsteady, light-headed, or faint.

with one or the other physical symptoms and do not receive appropriate treatment is 'functional treatment gap'. [27]

So far apparent treatment gap of PD is concerned, World health organization in 2004 reports that the median treatment gap of PD is 55.9%, while a large study in 2018 reported to be up to 91%.^[34] In India, the gap may increase to 98%.^[35] The detection rate in primary care is less than 50%.^[36] Functional treatment gap may reach to 85% due to physician's unawareness and may improve with awareness training.^[4]

Assessment of panic disorder

Panic disorder is clinically diagnosed, based on diagnostic guidelines. However, multiple psychological tools have been used to assess the different dimensions of PD [Table 4].

In the face of uncertainty, a patient may undergo a minimum investigation to exclude relevant probable physical illness. The common investigations considered at baseline include blood count, blood glucose level, thyroid function tests, renal function test, electrocardiography.^[37]

Course, outcome, and prognosis

After several years of initial attack, it recurs in 17-70%. About 1/3rd achieve stable remission, 1/4th achieve relapse-remission and less than ½ would have continuous course, particularly those with agoraphobia and long duration of illness at baseline, and presence mental or physical disorder, being poor and single. [38,39]

It is to be noted that about 56% of patients with PD remain untreated. [26] For those who receive treatment, 17–64% would not achieve remission [40] and 50% of treatment responder would relapse after discontinuation of medication. [8]

Comorbidity and associated symptoms

Most psychiatric comorbidities are agoraphobia (avoiding going out of home in the apprehension of having an attack) in about up to 84%, depression^[41] (10-15%), and other anxiety disorder (up to 30%). [42,43] These comorbidities result in functional impairment. [44] Reported attempted suicide is 25%, while complete suicide is thousand-fold more than the general population. [29,45] Most common medical comorbidities are Asthma [46] (6.5 to 24%), Hyperventilation [47] (35%) and cardiovascular disorder (e.g. Mitral valve prolapse).

Differential diagnosis

The PD must be differentiated from physical illnesses (See the list in table 5). The main difference that would raise a suspicion of underlying aetiology of physical illness are symptoms started with neurological symptoms (such as headache, unusual perception, motor or sensory dysfunction, ataxia), presence of atypical symptoms, extreme age at onset, altered consciousness or orientation, sphincter dyscontrol, and presence of sign/symptoms of a suspected physical illness and history of psychoactive or psychotropic drug use^[48] (Table 5; Table 6).

Clinical guidelines for the management of PD

In the management of PD, intervention shall be aimed at preventing further episodes of attacks (severity and frequency), anticipatory anxiety, agoraphobic avoidance, and achieving premorbid functioning by achieving full remission.^[49] Currently available evidence-based treatment options include pharmacotherapy and psychotherapy.^[50]

General consideration

The management begin with comprehensive information to patients in their understandable languages about PD followed by shared/collaborative decision. Majority can be treated on outpatient basis, while those with safety issues (e.g., suicidality) need inpatient care.

Clinical management during panic attack is to abort panic attack: Most of the panic attacks subsides spontaneously within 20 min, and patients reach medical centres with subsiding attacks

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Table 4: Tools to assess Panic Disorder		
Screening tools	Diagnostic instruments	
Panic Disorder Self-Report ^[46]	Anxiety Disorder Interview Schedule for DSM-5 ^[53]	
Anxiety Disorder Diagnostic Questionnaire ^[47]	Mini International Neuropsychiatric Interview ^[54]	
Generalized Anxiety Disorder -7 ^[48]	Structured Clinical Interview for DSM-IV ^[55]	
Quick Psycho Diagnostics ^[49]	Symptom Driven Diagnostic System for Primary Care ^[56]	
Mental Health Index 5 ^[50]		
Psychiatric Diagnostic Screening Questionnaire ^[51]		
Clinical Schedules for Primary Care Psychiatry ^[52]		
Severity Rating scale	Other purposes	
Beck Anxiety Inventory ^[57]	Panic attack questionnaire ^[63]	
Hamilton Anxiety Rating Scale ^[58]	Panic attack symptoms and cognition questionnaire ^[64]	
Panic Disorder Severity Scale ^[59]	Panic belief questionnaire ^[8]	
Panic and Agoraphobia Scale ^[60]	Self-efficacy to control Panic Attacks Questionnaire [65]	
Panic associated symptoms scale ^[61]	Panic Attack Frequency Calendar ^[66]	
NIMH Panic Questionnaire ^[62]	Panic appraisal questionnaire ^[67]	
	Nocturnal Panic Screen ^[68]	

Table 5: Physical illnesses as a differential diagnosis of Panic Disorder^[55]

	ranic Disorde	<u>L</u>	
System	rstem Diseases		
Cardiovascular	Anaemia	Hypertension	
Diseases	Angina	Mitral valve prolapses	
	Congestive heart failure	Myocardial infarction	
	Hyperactive adrenergic state	Paradoxicalatrial tachycardia	
Pulmonary	Asthma	Pulmonary embolus	
Diseases	Hyperventilation		
Neurological	Cerebrovascular disease	Migraine	
Diseases	Epilepsy	Multiple sclerosis	
	Huntington's disease	Transient ischemic attack	
	Infection	Tumour	
	Meniere's disease	Wilson's disease	
Endocrine	Addison's disease	Hypoglycemia	
Diseases	Carcinoid syndrome	Hypoparathyroidism	
	Cushing's syndrome	Menopausal disorders	
	Diabetes	Pheochromocytoma	
	Hyperthyroidism	Premenstrual syndrome	
Drug	Amphetamine	Hallucinogens	
Intoxications	Amyl nitrite	Marijuana	
	Anticholinergics	Nicotine	
	Cocaine	Theophylline	
	Alcohol	Opiates and opioids	
Drug Withdrawal	Antihypertensives	Sedative-hypnotics	
Other Conditions	Anaphylaxis	Systemic infections	
	B ₁₂ deficiency	Systemic lupus erythematosus	
	Electrolyte disturbances	Temporal arteritis	
	Heavy metal poisoning	Uraemia	

or feelings of anticipatory anxiety. Thus, explaining the very brief nature of illness and reassurance by the treating doctor during such an attack is helpful. A benzodiazepine with rapid onset of action (clonazepam, lorazepam and alprazolam) is the choice (except for those with benzodiazepine dependence) of pharmacotherapy to abort a panic attack on as-needed basis (up to 4- weeks), with instruction not to drive or operate dangerous equipment and abstain from alcohol or other sedative medication. Deep breathing and relaxation technique during attack or post-attack are also useful.

Pharmacotherapy

In general, the first-line drug of choice is Selective Serotonin Reuptake Inhibitors (SSRI), though if patients opt for another antidepressant group (SNRI and tricyclic) they may be tried. In patients with severe symptoms, short duration (4 weeks) of regular benzodiazepine with gradual tapering to be done over weeks along with antidepressants.^[51]

Though all SSRI are equally effective, [52] paroxetine is commonly used due to calm down effect (sedation). It should be started at 5–10 mg (CR preparation 12.5 to 25 mg) for 1 week then increase by 10 mg (CR preparation 12.5 mg) every week and consider the maximum dose of 60 mg (CR preparation 62.5 mg) if warranted for symptom control. Sertraline, Escitalopram, and fluvoxamine are the next best tolerated. Tricyclic (Clomipramine, Imipramine, and Desipramine) may be considered when patients prefer it or when SSRI or SNRI is not effective or cannot be used or not available. Disadvantages are higher propensity of

Features	Panic Attack	Partial Seizure	Syncope	Heart Attack
Immediate precipitating factors	May be situationally predisposed usually none	Usually none	Emotional stress, Valsalva, orthostatic hypotension, cardiac aetiologies	Physical or emotional Stress
Premonitory symptoms	Generalized anxiety in few cases	None or aura (e.g., odd odour) epigastric aura	Tiredness, nausea, diaphoresis, tunnelling of vision	Chest pain, anxiety, abnormal sensation in left arm
Posture at onset	variable	Variable	Usually erect	Variable
unconsciousness	If present gradual over seconds to minutes	Often immediate	Gradual over seconds	If present gradual over seconds to minutes
onset	Peak within minutes	sudden	gradual	usually gradual, over several minutes, and rarely instantaneous
Disorientation and sleepiness after event	Usually absent	Many minutes to hours	<5 min	Usually absent
Aching of muscles after event	Usually absent	Often	Sometimes	Usually absent
Biting of tongue	Usually absent	Sometimes	Rarely	Absent
Incontinence	Usually absent	Sometimes	Sometimes	Usually absent
Headache	Sometimes	Sometimes	Rarely	sometimes
Consciousness	Alert	Alert but may progress to impairment	impaired	Usually alert may be impaired
Dejavu hallucinations	Very rare	More than 5%	Very rare	Very rare
automatisms	Very infrequent	Common, progressing to CPS	Very infrequent	Very infrequent
Depressive symptoms	Commonly associated	uncommon	uncommon	uncommon
Anticipatory anxiety	Very common	uncommon	uncommon	uncommon
Interictal EEG	Normal	Often abnormal	Normal	Normal
ECG	normal	normal	normal	abnormal
MRI of temporal structures	Usually normal	Often abnormal	Usually normal	Usually Normal

severe side effects, higher dose requirement, very slow titration of dose (8–12 weeks), higher discontinuation rate.^[53] Table 7 provides a glimpse of antidepressant drugs.

As antipanic effect of antidepressants are slow, and outcome assessment should be done after 4-6 weeks of trial, after attaining an adequate dose. The effective medication (except benzodiazepines) should be continued for a minimum of 8-12 months, and discontinuation of medication should not be abrupt but gradual over many weeks. If symptoms reappear (30-90%) the medication should be resumed.^[52] Many patients may require long-term treatment for years to an indefinite period.^[54]

Psychotherapy

The most treatment guidelines recommend the cognitive behaviour therapy (CBT). The other psychotherapy used are psychodynamic psychotherapy, interpersonal psychotherapy, and acceptance-based approach. The cognitive-behavioral theory of PD assumes that attacks arise from distorted thoughts and catastrophization of the body symptoms^[55] and can be done online^[56] [Table 8]. This treatment option shall be offered for patients who refuses medication and request for

Table 7: Antipanic Drugs			
Drug	Starting (mg)	Maintenance (mg)	
SSRIs			
Paroxetine	5-10	20-60	
Paroxetine CR	12.5-25	62.5	
Fluoxetine	20	20-60	
Sertraline	12.5-25	50-200	
Fluvoxamine	12.5	100-150	
Citalopram	10	20-40	
Escitalopram	10	20	
Tricyclic Antidepressants			
Clomipramine	5-12.5	50-125	
Imipramine	10-25	150-500	
Desipramine	10-25	150-200	
Benzodiazepines			
Alprazolam	0.25-0.5 tid	0.5-2 tid	
Clonazepam	0.25-0.5 bid	0.5-2 bid	
Diazepam	2-5 bid	5-30 bid	
Lorazepam	0.25-0.5 bid	0.5-2 bid	
Atypical Antidepressants			
Venlafaxine	6.25-25	50-150	
Venlafaxine XR	37.5	150-225	

SSRIs - Selective Serotonin Reuptake Inhibitors; bid - twice a day; tid - three times a day.

non-pharmacological management. In this treating physicians are advised to refer to psychiatrists.

The CBT aims to address these issues in individual or group setting with informational intervention, cognitive restructuring, and exposure (interoceptive and *in vivo*).^[57] Applied relaxation^[58] and Breathing training,^[59,60] and *in-vivo* exposure^[61] are also helpful.

Treatment-resistant panic disorder

Failure to respond to an adequate dose of two or more of the standard pharmacological treatment for 4-6 week constitute a treatment-resistant state. [40] It may be associated with inadequate treatment, comorbidity and environmental even, and stress. [40] The initial approach is to optimise the treatment to ensure adherence, prescribe adequate dose, or increase to the maximum dose. On failure, switch within or between the class of antidepressants. Thus, initial switching between SSRI or SNRI and later to TCA, MAO, BZD, and CBT may be considered. The third approach is combination and augmentation. While considering for a combination, an antidepressant that initially had some response is combined with either another antidepressant of the same or different group or with CBT. The commonly used augmentation strategy is adding benzodiazepine, antipsychotics, and D-cycloserine. [62]

Role of primary care physicians

As mentioned earlier that PD is highly prevalent in the primary care setting, that may increase to 10 times in the patient's population with cardiac, gastrointestinal, ENT or neurological symptoms. They can be treated on an outpatient basis, and required hospitalization only for severe symptoms, comorbidities, and patient safety is a concern. [49] Immediate referral should be considered for severe anxiety, marked functional impairment, self-neglect, risk of self-harm or suicide, presence of psychiatric and medical comorbidities, and inadequate response to primary care intervention. Early referral decreases overall cost in assessment and management of PD.[63] In the Indian scenario, given that psychiatrists are scarce, the role of the non-psychiatric physicians are to identify the PD, and a trial of first-line pharmacotherapy should be provided. If patient does not respond in a month, they may be referred to psychiatrists for further management.^[64]

Prevention

Preventive intervention can be considered for those with high risks (e.g., increased anxiety sensitivity, history of panic

Table 8: CBT model of Panic disorder		
Core patterns in panic disorder	Common Catastrophic Thoughts in Panic Disorder	
Fears of symptoms of	Fears of death or disability	
anxiety (anxiety sensitivity)	Do I have a heart attack? I am having a stroke!; I am going to suffocate!	
Risk for onset of panic attacks	Fears of losing control/insanity	
Risk for biological provocation	I am going to lose control and scream; I am having a nervous breakdown; If I don't escape, I will go crazy	
of panic	Fears of humiliation or embarrassment	
Risk for panic disorder relapse	People will think something is wrong with me; They will think I am a lunatic	

attack). Those with diagnosed PD should be considered for relapse prevention in form of continuing pharmacological and non-pharmacological intervention with which patient responded.^[8]

Indian research

There is a paucity of study from India on PD. An epidemiological study reveals the prevalence of PD to be 0.52% and 1.6% of PD with agoraphobia, while apparent treatment gap was 98%.[9] The common psychiatric comorbidities observed were major depression and generalized anxiety disorder, [38,65] and these comorbidities were associated with more severe illness, increased severity of panic, and other anxiety symptoms. [66] The presence of agoraphobia; comorbid psychiatric disorders and stress were associated with more severe illness, increased severity of panic, and other anxiety symptoms^[67] Haematological feature of increased platelet distribution width and red cell distribution width has been observed. [68] Regarding treatment, CBT was found to be effective. At the same time, Yoga and breathing exercise was useful adjunctive treatment in PD.[11,69] "Clinical Schedule for Primary Care Psychiatry" (CSP) is a recently designed and validated point of care manual to identify psychiatric disorders including panic disorder by primary care doctors[11] including non-psychiatric specialists.

Summary and Conclusions

Panic disorder is highly prevalent in primary care setting. As the symptoms mimic of serious disease, most patient initially seek help of primary care physician or other non-psychiatrist specialist. Due to less familiarity about illness, they are under-diagnosed frequently and undergo unnecessary investigation. Antidepressant and psychotherapies remain the main stay of treatment and prevention. Despite advancement in management, most patients remain untreated or undertreated. Despite treatment, two-third patients experience remission with varying severity, and treatment is required to be continued for long term. There is an urgent need that non-psychiatrist physician to play a vital role in the care of panic disorder.

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Conflicts of interest

There are no conflicts of interest.

References

- Leon AC, Olfson M, Portera L. Service utilization and expenditures for the treatment of panic disorder. Gen Hosp Psychiatry 1997;19:82-8.
- de Jonge P, Roest AM, Lim CC, Florescu SE, Bromet EJ, Stein DJ, et al. Cross-national epidemiology of panic disorder and panic attacks in the world mental health surveys. Depress Anxiety 2016;33:1155-77.
- 3. Grant BF, Hasin DS, Stinson FS, Dawson DA, Goldstein RB,

- Smith S, *et al.* The epidemiology of DSM-IV panic disorder and agoraphobia in the United States: Results from the National Epidemiologic Survey on Alcohol and Related Conditions. J Clin Psychiatry 2006;67:363-74.
- Vermani M, Marcus M, Katzman MA. Rates of detection of mood and anxiety disorders in primary care: A descriptive, cross-sectional study. Prim Care Companion CNS Disord 2011;13:PCC.10m01013.
- Dammen T, Arnesen H, Ekeberg Ø, Husebye T, Friis S. Panic disorder in chest pain patients referred for cardiological outpatient investigation. J Intern Med 1999;245:497-507.
- Fujii K, Uchida H, Suzuki T, Mimura M. Dependence on benzodiazepines in patients with panic disorder: A crosssectional study. Psychiatry Clin Neurosci 2015;69:93-9.
- Beitman BD, Kushner MG, Basha I, Lamberti J, Mukerji V, Bartels K. Follow-up status of patients with angiographically normal coronary arteries and panic disorder. JAMA 1991;265:1545-9.
- Caldirola D, Alciati A, Daccò S, Micieli W, Perna G. Relapse prevention in panic disorder with pharmacotherapy: Where are we now? Expert Opin Pharmacother 2020;21:1699-711.
- 9. Gururaj G, Varghese M, Benegal V, Rao GN, Pathak K, Singh LK, *et al.* National Mental Health Survey of India, 2015-16: Prevalence, patterns and outcomes. Bengaluru, National Institute of Mental Health and Neurosciences, NIMHANS Publication No. 129, 2016. p. 92.
- King M, Nazareth I, Levy G, Walker C, Morris R, Weich S, et al. Prevalence of common mental disorders in general practice attendees across Europe. Br J Psychiatry 2008;192:362-7.
- 11. Pahuja E, Kumar ST, Uzzafar F, Manjunatha N, Kumar CN, Gupta R, *et al.* An impact of a digitally driven primary care psychiatry program on the integration of psychiatric care in the general practice of primary care doctors. Indian J Psychiatry 2020;62:690-6.
- 12. Todaro JF, Shen BJ, Raffa SD, Tilkemeier PL, Niaura R. Prevalence of anxiety disorders in men and women with established coronary heart disease. J Cardiopulm Rehabil Prev 2007;27:86-91.
- 13. Tully PJ, Baker RA. Depression, anxiety, and cardiac morbidity outcomes after coronary artery bypass surgery: A contemporary and practical review. J Geriatr Cardiol 2012;9:197-208.
- 14. Shakeri J, Tatari F, Vaezi N, Golshani S, Farnia V, Alikhani M, *et al.* The prevalence of panic disorder and its related factor in hospitalized patients with chest pain and normal angiography. J Educ Health Promot 2019;8:61.
- 15. Stein MB, Asmundson GJ, Ireland D, Walker JR. Panic disorder in patients attending a clinic for vestibular disorders. Am J Psychiatry 1994;151:1697-700.
- 16. Shipko S. Panic disorder in otolaryngologic practice: A brief review. Ear Nose Throat J 2001;80:867-8.
- 17. Livermore N, Sharpe L, McKenzie D. Panic attacks and panic disorder in chronic obstructive pulmonary disease: A cognitive behavioral perspective. Respir Med 2010;104:1246-53.
- 18. Smitherman TA, Kolivas ED, Bailey JR. Panic disorder and migraine: Comorbidity, mechanisms, and clinical implications. Headache 2013;53:23-45.
- 19. Hall RC, Hall RC. Anxiety and endocrine disease. Semin Clin Neuropsychiatry 1999;4:72-83.
- 20. Chang HM, Pan CH, Chen PH, Chen YL, Su SS, Tsai SY, et al. Identification and medical utilization of newly

- diagnosed panic disorder: A nationwide case-control study. J Psychosom Res 2019;125:109815.
- 21. Vigne P, Fortes P, Dias RV, Laurito LD, Loureiro CP, de Menezes GB, *et al.* Duration of untreated illness in a cross-diagnostic sample of obsessive-compulsive disorder, panic disorder, and social anxiety disorder. CNS Spectr 2019;24:526-32.
- 22. Fleet R, Lavoie K, Beitman BD. Is panic disorder associated with coronary artery disease? A critical review of the literature. J Psychosom Res 2000;48:347-56.
- 23. Lynch P, Galbraith K. Panic in the emergency room. Can J Psychiatry 2003;48:361-6.
- 24. Lavey EB, Winkle RA. Continuing disability of patients with chest pain and normal coronary arteriograms. J Chronic Dis 1979;32:191-6.
- 25. Kaplan DS, Masand PS, Gupta S. The relationship of irritable bowel syndrome (IBS) and panic disorder. Ann Clin Psychiatry 1996;8:81-8.
- 26. Ballenger JC. Panic disorder in the medical setting. J Clin Psychiatry 1997;58(Suppl 2):13-7.
- 27. Manjunatha N, Kumar CN, Math SB, Thirthalli J. Designing and implementing an innovative digitally driven primary care psychiatry program in India. Indian J Psychiatry 2018;60:236-44.
- DSM-5. The Diagnostic and Statistical Manual of Mental Disorders. 5th ed. DSM-5; American Psychiatric Association; 2013.
- 29. Roy-Byrne PP, Craske MG, Stein MB. Panic disorder. Lancet 2006;368:1023-32.
- 30. Cosci F, Mansueto G. Biological and clinical markers in panic disorder. Psychiatry Investig 2019;16:27-36.
- 31. Gorman JM, Kent JM, Sullivan GM, Coplan JD. Neuroanatomical hypothesis of panic disorder, revised. Am J Psychiatry 2000;157:493-505.
- 32. de Carvalho MR, Rozenthal M, Nardi AE. The fear circuitry in panic disorder and its modulation by cognitive-behaviour therapy interventions. World J Biol Psychiatry 2010;11 (2 Pt 2):188-98.
- 33. Tretiakov A, Malakhova A, Naumova E, Rudko O, Klimov E. Genetic biomarkers of panic disorder: A systematic review. Genes 2020;11:1310.
- 34. Alonso J, Liu Z, Evans-Lacko S, Sadikova E, Sampson N, Chatterji S, *et al.* Treatment gap for anxiety disorders is global: Results of the World Mental Health Surveys in 21 countries. Depress Anxiety 2018;35:195-208.
- Sagar R, Pattanayak RD, Chandrasekaran R, Chaudhury PK, Deswal BS, Lenin Singh RK, et al. Twelve-month prevalence and treatment gap for common mental disorders: Findings from a large-scale epidemiological survey in India. Indian J Psychiatry 2017;59:46-55.
- 36. Culpepper L. Use of algorithms to treat anxiety in primary care. J Clin Psychiatry 2003;64(Suppl 2):30-3.
- 37. Andrews G, Bell C, Boyce P, Gale C, Lampe L, Marwat O, *et al.* Royal Australian and New Zealand College of Psychiatrists clinical practice guidelines for the treatment of panic disorder, social anxiety disorder and generalised anxiety disorder. Aust N Z J Psychiatry 2018;52:1109-72.
- 38. Bruce SE, Yonkers KA, Otto MW, Eisen JL, Weisberg RB, Pagano M, *et al.* Influence of psychiatric comorbidity on recovery and recurrence in generalized anxiety disorder, social phobia, and panic disorder: A 12-year prospective study. Am J Psychiatry 2005;162:1179-87.

- 39. Teismann T, Lukaschek K, Hiller TS, Breitbart J, Brettschneider C, Schumacher U, *et al.* Suicidal ideation in primary care patients suffering from panic disorder with or without agoraphobia. BMC Psychiatry 2018;18:1-5.
- 40. Freire RC, Zugliani MM, Garcia RF, Nardi AE. Treatment-resistant panic disorder: A systematic review. Expert Opin Pharmacother 2016;17:159-68.
- 41. Baldwin D. Easing the burden of panic disorder: Comorbidity, diagnosis, and implications for treatment. CNS Spectrums 2005;10(S12):5-11.
- 42. Roy-Byrne PP, Stein MB, Russo J, Mercier E, Thomas R, McQuaid J, *et al.* Panic disorder in the primary care setting: Comorbidity, disability, service utilization, and treatment. J Clin Psychiatry 1999;60:492-9.
- 43. Barlow DH. Comorbidity among anxiety disorders: Implications for treatment and DSM-IV. J Consult Clin Psychol 1992;60:835-44.
- 44. Kim HJ, Kim JE, Lee SH. Functional impairment in patients with panic disorder. Psychiatry Investig 2021;18:434-42.
- *45.* Tietbohl-Santos B, Chiamenti P, Librenza-Garcia D, Cassidy R, Zimerman A, Manfro GG, *et al.* Risk factors for suicidality in patients with panic disorder: A systematic review and meta-analysis. Neurosci Biobehav Rev 2019;105:34-8.
- Shavitt RG, Gentil V, Mandetta R. The association of panic/ agoraphobia and asthma. Contributing factors and clinical implications. Gen Hosp Psychiatry 1992;14:420-3.
- 47. Hoes MJ, Colla P, van Doorn P, Folgering H, de Swart J. Hyperventilation and panic attacks. J Clin Psychiatry 1987;48:435-7.
- 48. Cameron OG. The differential diagnosis of anxiety: Psychiatric and medical disorders. Psychiatr Clin North Am 1985;8:3-23.
- 49. Bandelow B, Baldwin DS. Pharmacotherapy for panic disorder. The American Psychiatric Association Publishing Textbook of Anxiety, Trauma, and OCD-Related Disorders. 3rd ed. Washington DC: American Psychiatric Pub; 2020. p. 385-407.
- 50. Zulfarina MS, Syarifah-Noratiqah SB, Nazrun SA, Sharif R, Naina-Mohamed I. Pharmacological therapy in panic disorder: Current guidelines and novel drugs discovery for treatment-resistant patient. Clin Psychopharmacol Neurosci 2019;17:145-54.
- 51. Quagliato LA, Cosci F, Shader RI, Silberman EK, Starcevic V, Balon R, *et al.* Selective serotonin reuptake inhibitors and benzodiazepines in panic disorder: A meta-analysis of common side effects in acute treatment. J Psychopharmacol 2019;33:1340-51.
- 52. Du Y, Du B, Diao Y, Yin Z, Li J, Shu Y, *et al.* Comparative efficacy and acceptability of antidepressants and benzodiazepines for the treatment of panic disorder: A systematic review and network meta-analysis. Asian J Psychiatry 2021;60:102664.
- 53. Bakker A, Van Balkom AJ, Spinhoven P. SSRIs vs. TCAs in the treatment of panic disorder: A meta-analysis. Acta Psychiatr Scand 2002;106:163-7.
- 54. Vanelli M. Improving treatment response in panic disorder. Primary Psychiatry 2005;12:68-73.
- 55. Maisto D, Barca L, Van den Bergh O, Pezzulo G. Perception and misperception of bodily symptoms from an active inference perspective: Modelling the case of panic disorder. Psychol Rev 2021;128:610-710.
- Efron G, Wootton BM. Remote cognitive behavioral therapy for panic disorder: A meta-analysis. J Anxiety Disord 2021;79:102385.

- 57. Curtiss JE, Levine DS, Rosenbaum JF, Baker A. Cognitive-behavioral strategies to manage panic disorder. Psychiatr Ann 2021;51:216-20.
- 58. Ost LG, Westling BE. Applied relaxation vs cognitive behavior therapy in the treatment of panic disorder. Behav Res Ther 1995;33:145-58.
- 59. Taylor S. Breathing retraining in the treatment of panic disorder: Efficacy, caveats and indications. Scan J Beha Ther 2001;30:49-56.
- 60. Roberson-Nay R, Kendler KS. Panic disorder and its subtypes: A comprehensive analysis of panic symptom heterogeneity using epidemiological and treatment seeking samples. Psychol Med 2011;41:2411-21.
- 61. Öst LG, Westling BE, Hellström K. Applied relaxation, exposure *in vivo* and cognitive methods in the treatment of panic disorder with agoraphobia. Behav Res Ther 1993;31:383-94.
- 62. Holt RL, Lydiard RB. Management of treatment-resistant panic disorder. Psychiatry (Edgmont) 2007;4:48-59.
- 63. Chen YH, Chen SF, Lin HC, Lee HC. Healthcare utilization patterns before and after contact with psychiatrist care for panic disorder. J Affect Disord 2009;119:172-6.
- 64. Manjunatha N, Naveen Kumar C, Suresh B, Jagadisha T.

- Clinical schedules for primary care psychiatry-version 2.2. In: Manjunatha N, Kumar CN, Math SB, Basvaraju V, Shashidhara HN, Parthasarathy P, eds. Karnataka Telemedicine, Mentoring and Monitoring Program: An Implementation Manual of Primary Care Psychiatry, Published by Department of Psychiatry, National Institute of Mental Health and Neurosciences, Bengaluru, NIMHANS Publication No. 157, 2019 (ISBN: 978-81-86506-00-4).
- 65. Srinivasa K, Neerakal I. A study of panic patients with and without depression. Indian J Psychiatry 2002;44:246.
- 66. Salhotra N, Bhattacharyya D, Wadhawan JM, Yadav P. Sociodemographic and Clinical Variables related to panic disorder with and without Agoraphobia. J Clin Diag Res 2018;12:VC01-4.
- 67. Amatya M, Paudel BH. Effects of Yoga on anxiety and autonomic function in panic disorder patients. Inter J Pharm Biol Arch 2012;3:1082-5.
- 68. Ransing RS, Patil B, Grigo O. Mean platelet volume and platelet distribution width level in patients with panic disorder. J Neurosci Rural Pract 2017;8:174-8.
- 69. Bhagat V, Haque M, Jaalam K. Breathing exercise-A commanding tool for self-help management during panic attacks. Res J Pharm Tech 2017;10:4471-3.