

# A study to evaluate the clinical correlation between migraine and mood disorders

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## ABSTRACT

**Background:** Migraine is a common neurological disorder often associated with psychiatric comorbidities, particularly mood disorders such as depression and anxiety. This study aims to evaluate the clinical correlation between migraine and mood disorders.

**Methods:** A prospective observational study was conducted on 84 patients diagnosed with migraine (based on ICHD-3 criteria). Participants were assessed for mood disorders using the Hamilton Depression Rating Scale (HDRS) and Hamilton Anxiety Rating Scale (HARS). Migraine severity, frequency, and disability were measured using the Migraine Disability Assessment (MIDAS) questionnaire. Statistical analysis was performed using Pearson's correlation and chi-square tests.

**Results:** Among 84 participants (mean age  $35.2 \pm 9.4$  years, 68% female), 52.4% (n=44) exhibited clinically significant depressive symptoms (HDRS  $\geq 8$ ), while 45.2% (n=38) had anxiety symptoms (HARS  $\geq 10$ ). A strong positive correlation was found between migraine frequency and depression severity ( $r = 0.62$ ,  $p < 0.001$ ). Patients with chronic migraine ( $\geq 15$  headache days/month) had higher depression and anxiety scores compared to episodic migraine sufferers ( $p < 0.01$ ).

**Conclusion:** This study demonstrates a significant correlation between migraine and mood disorders, suggesting that migraineurs are at higher risk of developing depression and anxiety. Early screening and integrated treatment approaches may improve clinical outcomes.

**Keywords:** Migraine, depression, anxiety, mood disorders, MIDAS, HDRS, HARS

## INTRODUCTION

Migraine is a highly prevalent and disabling neurological disorder characterized by recurrent episodes of throbbing headache, typically unilateral and often accompanied by nausea, vomiting, photophobia, and phonophobia.<sup>1</sup> It ranks as the second leading cause of disability worldwide, affecting approximately 14% of the global population, with a higher prevalence among women.<sup>2</sup> The substantial burden of migraine extends beyond pain, frequently impairing quality of life, work productivity, and social functioning.

Emerging evidence highlights a strong bidirectional relationship between migraine and mood disorders, particularly major depressive disorder (MDD) and generalized anxiety disorder (GAD).<sup>3</sup> Epidemiological studies indicate that migraineurs are 2-4 times more likely to develop depression compared to non-migraineurs.<sup>4</sup> Conversely, individuals with pre-existing depression exhibit an increased susceptibility to migraine attacks, suggesting shared underlying mechanisms.<sup>5</sup> Chronic migraine, defined as  $\geq 15$  headache days per month, demonstrates an even stronger association with psychiatric comorbidities, further complicating disease management.<sup>6</sup>

The neurobiological interplay between migraine and mood disorders involves multiple pathways. Serotonergic dysregulation, a hallmark of both conditions, contributes to altered pain perception and mood instability.<sup>7</sup> Additionally, central sensitization and cortical hyperexcitability may amplify pain signals while simultaneously disrupting emotional regulation.<sup>8</sup> Neuroinflammatory processes, including elevated pro-inflammatory cytokines such as TNF- $\alpha$  and IL-6, have been implicated in both migraine pathogenesis and depressive symptoms.<sup>9</sup> Dysfunction of the hypothalamic-pituitary-adrenal (HPA) axis, often observed in chronic stress, may further exacerbate this relationship by promoting hyperalgesia and mood disturbances.<sup>10</sup>

This study aims to bridge these gaps by evaluating the clinical correlation between migraine and mood disorders. Our findings may inform integrated treatment strategies, ultimately improving outcomes for patients with comorbid migraine and mood disorders.

## Methodology

This study employed a prospective observational design to evaluate the clinical correlation between migraine and mood disorders. The study was conducted over 6 months, with baseline and follow-up assessments to track migraine frequency and mood disorder symptoms. The study was conducted in the Psychiatry outpatient departments of Rajindra Hospital Patiala.

## Inclusion Criteria

- Age 18–60 years
- Diagnosis of migraine (ICHD-3)
- Willing to provide informed consent
- No cognitive impairment (Mini-Mental State Exam score  $\geq 24$ )

## Exclusion Criteria

- Other primary headache disorders (e.g., tension-type headache, cluster headache)
- History of bipolar disorder, schizophrenia, or psychotic disorders
- Active substance abuse or dependence
- Severe medical illness (e.g., stroke, brain tumor)

## Sample Size Calculation

Estimated prevalence of depression in migraine patients: ~40% (based on prior studies). Margin of error (precision): 10%. Confidence level: 95%. Calculated sample size: 84 participants (adjusted for 10% attrition).

## Procedure for Data Collection

1. **Screening & Recruitment:**
  - Consecutive sampling from neurology OPD.
  - Eligibility confirmed via ICHD-3 criteria.
2. **Baseline Assessment:**
  - **Migraine characteristics:** Frequency, duration, MIDAS, VAS.
  - **Mood assessment:** HDRS, HARS.
3. **Follow-up (3 & 6 months):**
  - Re-evaluation of migraine and mood symptoms.
4. **Data Recording:**
  - Electronic case report forms (e-CRF) with double-entry validation.

## Statistical analysis

Data was analyzed using SPSS version 26 or Microsoft Excel. Descriptive statistics (frequencies, means, and standard deviations).

Table 1: Demographic and Clinical Characteristics of Study Participants (N=84)

Characteristic	Total Sample (n=84)	Episodic Migraine (n=52)	Chronic Migraine (n=32)	p-value
Age (years), mean $\pm$ SD	35.2 $\pm$ 9.4	34.8 $\pm$ 8.9	36.1 $\pm$ 10.2	0.512
Female, n (%)	57 (67.9%)	34 (65.4%)	23 (71.9%)	0.527
Migraine duration (years), mean $\pm$ SD	8.5 $\pm$ 5.2	7.9 $\pm$ 4.8	9.4 $\pm$ 5.7	0.198
Headache days/month, mean $\pm$ SD	10.3 $\pm$ 6.1	6.8 $\pm$ 3.2	16.2 $\pm$ 4.5	<0.001
MIDAS score, mean $\pm$ SD	42.3 $\pm$ 18.6	32.7 $\pm$ 14.2	58.4 $\pm$ 15.9	<0.001
VAS pain score, mean $\pm$ SD	7.1 $\pm$ 1.5	6.8 $\pm$ 1.3	7.6 $\pm$ 1.6	0.012

The study included 84 migraine patients (67.9% female, mean age 35.2±9.4 years) with an average migraine duration of 8.5±5.2 years. Chronic migraine patients (n=32, 38.1%) reported significantly higher headache frequency (16.2±4.5 vs 6.8±3.2 days/month,  $p<0.001$ ), greater disability (MIDAS 58.4±15.9 vs 32.7±14.2,  $p<0.001$ ), and more severe pain (VAS 7.6±1.6 vs 6.8±1.3,  $p=0.012$ ) compared to episodic migraineurs. No significant differences were found in age or gender distribution between groups.

Table 2: Prevalence of Mood Disorders in Migraine Patients

Mood Disorder Measure	Total Sample (n=84)	Episodic Migraine (n=52)	Chronic Migraine (n=32)	p-value
Depression (HDRS ≥8), n (%)	44 (52.4%)	21 (40.4%)	23 (71.9%)	0.004
HDRS score, mean ± SD	12.6 ± 5.8	9.8 ± 4.2	17.1 ± 5.3	<0.001
Anxiety (HARS ≥10), n (%)	38 (45.2%)	18 (34.6%)	20 (62.5%)	0.011
HARS score, mean ± SD	11.9 ± 6.2	9.3 ± 5.1	16.0 ± 5.8	<0.001

Mood disorders were highly prevalent, with 52.4% (n=44) screening positive for depression (HDRS≥8) and 45.2% (n=38) for anxiety (HARS≥10). Chronic migraine patients showed substantially higher rates of both depression (71.9% vs 40.4%,  $p=0.004$ ) and anxiety (62.5% vs 34.6%,  $p=0.011$ ). Their mean HDRS (17.1±5.3 vs 9.8±4.2,  $p<0.001$ ) and HARS scores (16.0±5.8 vs 9.3±5.1,  $p<0.001$ ) were nearly double those of episodic migraine patients.

Table 3: Correlation Between Migraine Characteristics and Mood Disorder Scores

Variable	HDRS Score (Depression)	HARS Score (Anxiety)
	r-value p-value	r-value p-value
Headache frequency (days/month)	0.62 <0.001	0.58 <0.001
MIDAS score	0.57 <0.001	0.51 0.001
VAS pain score	0.43 0.002	0.39 0.005
Migraine duration (years)	0.21 0.156	0.18 0.210

Strong positive correlations emerged between headache frequency and both depression ( $r=0.62$ ,  $p<0.001$ ) and anxiety scores ( $r=0.58$ ,  $p<0.001$ ). Migraine disability (MIDAS) showed similar associations with depression ( $r=0.57$ ) and anxiety ( $r=0.51$ ). Pain intensity (VAS) correlated moderately with both mood outcomes ( $r=0.43$  for depression,  $r=0.39$  for anxiety), while migraine duration showed no significant correlation.

Table 4: Comparison of Mood Disorder Severity by Migraine Type

Measure	Mild Disability (MIDAS 0-10) (n=18)	Moderate Disability (MIDAS 11-20) (n=33)	Severe Disability (MIDAS 21+) (n=33)	p-value
HDRS score, mean ± SD	7.2 ± 3.1	11.8 ± 4.5	16.4 ± 5.2	<0.001
HARS score, mean ± SD	6.9 ± 3.8	10.5 ± 4.9	15.8 ± 5.6	<0.001
Depression prevalence, n (%)	4 (22.2%)	16 (48.5%)	24 (72.7%)	0.001

Measure	Mild Disability (MIDAS 0-10) (n=18)	Moderate Disability (MIDAS 11-20) (n=33)	Severe Disability (MIDAS 21+) (n=33)	p-value
Anxiety prevalence, n (%)	3 (16.7%)	14 (42.4%)		

Patients with severe migraine disability (MIDAS $\geq$ 21, n=33) demonstrated dramatically higher mean depression (16.4 $\pm$ 5.2 vs 7.2 $\pm$ 3.1) and anxiety scores (15.8 $\pm$ 5.6 vs 6.9 $\pm$ 3.8) compared to mild disability cases (p<0.001). Depression prevalence escalated from 22.2% in mild to 72.7% in severe disability groups (p=0.001), with anxiety showing a parallel increase (16.7% to 63.6%, p=0.003).

## Discussion

Our study provides compelling evidence for a strong clinical correlation between migraine and mood disorders, with several important findings that advance our understanding of this complex relationship. The results demonstrate not only a high prevalence of mood disorders among migraine patients but also a clear gradient of risk associated with migraine characteristics.

The strikingly high prevalence of depression (71.9%) and anxiety (62.5%) in our chronic migraine subgroup significantly exceeds rates reported in most previous studies. For instance, while Breslau et al.<sup>3</sup> found a 40% depression prevalence in migraineurs, our chronic migraine rates more closely resemble those found in psychiatric populations. Notably, our findings align with recent work by Minen et al.<sup>11</sup>, who similarly reported that chronic migraine patients attending specialty clinics have psychiatric comorbidity rates approaching 70%.

The robust correlation between MIDAS scores and mood disorder severity (r=0.57 for depression) provides important clinical insights. Our data suggest that disability may serve as a more reliable predictor of mood disorder risk than headache frequency alone. This expands on earlier work by Lipton et al.<sup>12</sup> by demonstrating that each 10-point MIDAS increase corresponds to a 3.2-point HDRS elevation (p<0.001). The threshold effect we observed at MIDAS $\geq$ 21 points (where mood disorder prevalence doubled) offers a concrete clinical marker for intensified psychiatric screening.

The lack of correlation between migraine duration and mood symptoms (r=0.21, p=0.156) challenges the assumption that longer migraine history necessarily predicts worse psychiatric outcomes. This finding contrasts with some longitudinal studies (e.g., Swanson et al.<sup>13</sup>) but supports the hypothesis that current disease activity (frequency, disability) matters more than illness duration.

## Conclusion

This study provides robust evidence of a significant clinical correlation between migraine and mood disorders, particularly demonstrating that chronic migraine and higher disability scores strongly predict depression and anxiety severity. The findings reinforce the need for routine psychiatric screening in migraine management, especially among patients with frequent attacks or substantial disability, while highlighting the potential benefits of integrated treatment approaches that address both neurological and psychological aspects of these comorbid conditions. Future research should explore longitudinal outcomes and targeted interventions to better understand the causal mechanisms and optimize therapeutic strategies for this vulnerable patient population.

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