### 1. Bipolar Disorder:

**Zarate et al. (2012)** This article reviews the effects of ketamine in treating depressive episodes in individuals with bipolar disorder. The findings indicate that ketamine produces rapid and significant reductions in depressive symptoms, even in patients who have not responded to traditional treatments. The study highlights ketamine's unique mechanism of action, involving NMDA receptor antagonism and enhanced neuroplasticity, which addresses the underlying neurobiological factors of bipolar depression. The authors suggest that ketamine is a promising intervention for managing treatment-resistant bipolar depression, offering a rapid-acting alternative to conventional therapies. (International Journal of Neuropsychopharmacology)

**Rodolico et al. (2021)** reviewed the efficacy and safety of ketamine and esketamine in bipolar depression, highlighting their potential as treatment options. The analysis revealed that both ketamine and esketamine are effective in reducing depressive symptoms across various forms of depression, including bipolar depression. Frontiers **Jawad et al. (2023)** This scoping review analyzed 10 clinical studies, including 5 randomized controlled trials and 5 open-label studies, to evaluate the efficacy and safety of ketamine in treating bipolar depression. The findings suggest that ketamine provides rapid antidepressant effects, particularly for individuals with treatment-resistant bipolar depression. However, the authors emphasize the need for further research to confirm these preliminary findings and establish safety profiles. (MDPI - Brain Sciences)

### 2. Post-Traumatic Stress Disorder (PTSD):

**Philipp-Muller et al. (2023):** This systematic review and meta-analysis evaluate the combined use of ketamine and psychotherapy in treating posttraumatic stress disorder (PTSD). The findings suggest that integrating ketamine with trauma-focused psychotherapies may enhance treatment outcomes, leading to significant reductions in PTSD symptoms. <u>Psychiatrist</u>

**Reis et al. (2021):** In anxiety disorders, KAP shows potential for reducing symptoms of generalized anxiety disorder (GAD) and social anxiety by promoting emotional flexibility and reducing hypervigilance. While evidence is preliminary, it suggests ketamine's anxiolytic effects can enhance psychotherapy outcomes. (<u>Cambridge University Press</u>) **Dore et al. (2019):** For post-traumatic stress disorder (PTSD), ketamine reduces hyperarousal and intrusive thoughts, creating an optimal window for trauma-focused psychotherapy. It also promotes emotional processing and symptom reduction. (<u>Taylor & Francis Online</u>)

**Feder et al. (2014)** This randomized, double-blind, placebo-controlled crossover study evaluated the effects of a single ketamine infusion on individuals with chronic posttraumatic stress disorder (PTSD). The study found that ketamine produced rapid and significant reductions in PTSD symptoms, including intrusive thoughts, hyperarousal, and overall symptom severity, within 24 hours of administration. These effects were sustained for up to two weeks in some participants. The authors concluded that ketamine's ability to enhance neuroplasticity and modulate glutamate pathways makes it a promising treatment for PTSD, particularly for patients with treatment-resistant symptoms JAMA Network

### 3. Anxiety Disorders:

**Reis et al. (2021):** In anxiety disorders, KAP shows potential for reducing symptoms of generalized anxiety disorder (GAD) and social anxiety by promoting emotional flexibility and reducing hypervigilance. While evidence is preliminary, it suggests ketamine's anxiolytic effects can enhance psychotherapy outcomes. (Cambridge University Press) **Kolp et al. (2007):** This article explores the use of ketamine-enhanced psychotherapy (KEP) in addressing death anxiety among terminally ill patients. Through a review of literature and two detailed case studies, the findings suggest that KEP facilitates profound transpersonal experiences, reducing death-related fears and enhancing inner peace. The authors advocate for further research into KEP as a therapeutic approach for managing death anxiety in end-of-life care. (CIIS - International Journal of Transpersonal Studies)

**Dore et al. (2019):** For anxiety disorders, including generalized anxiety disorder (GAD) and social anxiety disorder, KAP appears to reduce symptoms by modulating the brain's stress response systems. The article suggests that ketamine enhances the effectiveness of psychotherapeutic interventions for persistent anxiety. (<u>Taylor & Francis Online</u>)

# 4. Obsessive-Compulsive Disorder (OCD):

**Lee et al. (2021):** This review highlights ketamine's potential to provide rapid symptom relief for individuals with Obsessive–Compulsive Disorder (OCD), especially those who have not responded to traditional treatments. However, the effects are often short-lived, underscoring the need for further research to determine optimal dosing and long-term efficacy. (MDPI - Brain Sciences)

**Grözinger et al. (2023):** This case report highlights significant improvements in a 28-year-old patient with severe obsessive-compulsive disorder (OCD) and comorbid depression following treatment with esketamine combined with psychotherapy. The findings suggest that ketamine-assisted psychotherapy may offer a promising option for individuals with treatment-resistant OCD and depression. (Frontiers in Psychiatry) **Dore et al. (2019):** For obsessive-compulsive disorder (OCD), ketamine-assisted psychotherapy helps reduce obsessive thoughts and compulsive behaviors by targeting maladaptive neural pathways and enhancing therapeutic engagement. (Taylor & Francis Online)

# 5. Chronic Pain Conditions:

**Batievsky et al. (2023):** This pilot study explores two approaches to ketamine-assisted psychotherapy (KAP) for individuals suffering from chronic pain and comorbid depression. The findings suggest that KAP may effectively alleviate symptoms of both chronic pain and depression, as well as associated anxiety and PTSD. The study indicates that a psychedelic approach to KAP could be more beneficial, highlighting the need for further research to optimize treatment protocols for this population. Frontiers

**Niesters et al. (2014):** This review evaluates the risks and benefits of ketamine for chronic pain management. It highlights ketamine's efficacy in reducing pain intensity and opioid consumption, particularly for conditions like complex regional pain syndrome

(CRPS) and neuropathic pain. The article also addresses potential neuropsychiatric side effects, emphasizing the need for careful patient selection and monitoring. While ketamine shows promise as a treatment for chronic pain, the authors call for further research to clarify its long-term efficacy and safety. (British Journal of Clinical Pharmacology)

**Schwartzman et al. (2009)** This study explored the long-term effects of ketamine infusions for patients with intractable complex regional pain syndrome (CRPS). The findings demonstrated that ketamine, administered as a continuous infusion over several days, resulted in significant reductions in pain intensity and improvements in function. The study highlights ketamine's effectiveness in targeting central sensitization and reducing neuropathic pain symptoms. Importantly, the authors emphasize ketamine's potential as a therapeutic option for CRPS patients who have not responded to conventional treatments, though they advocate for careful patient selection and monitoring. <u>Oxford Academic</u>

**Sigtermans et al. (2009)** Results showed that ketamine significantly reduced pain intensity compared to placebo, with improvements persisting beyond the infusion period. The authors concluded that ketamine's NMDA receptor antagonism plays a critical role in alleviating central sensitization and chronic pain associated with CRPS. This study highlights ketamine's potential as a valuable treatment for severe, treatment-resistant chronic pain conditions.. <u>ScienceDirect</u>

### 6. Suicidal Ideation and Self-Harm Behaviors:

**Wolfson and Vaid. (2024):** This article highlights the potential benefits of ketamine-assisted psychotherapy for addressing self-harm and suicidal ideation. The authors emphasize that ketamine's rapid antidepressant effects, when combined with psychotherapy, can help patients process underlying psychological issues contributing to these behaviors. By facilitating emotional insight and resolution, this integrated approach shows promise in reducing self-harm and suicidal thoughts. The article also underscores the importance of the therapeutic setting and subjective experiences in achieving optimal outcomes. (Frontiers in Psychiatry)

**Witt et al. (2020):** This systematic review and meta-analysis assess the efficacy of ketamine in reducing suicidal ideation among adults with psychiatric disorders. The findings indicate that ketamine administration leads to a significant and rapid decrease in suicidal thoughts, with effects observed as early as 4 hours post-administration and lasting up to 7 days. The study highlights ketamine's potential as a valuable intervention for acute suicidal ideation, especially in cases where traditional treatments may not provide immediate relief.

#### **SAGE Journals**

**Grunebaum et al. (2017)** This randomized, double-blind, placebo-controlled trial investigated the effects of ketamine on suicidal ideation in patients with major depressive disorder. The study found that a single infusion of ketamine led to a rapid and significant reduction in suicidal thoughts within 24 hours compared to the placebo group. These improvements were maintained for several days following the infusion. The authors highlight ketamine's potential as a critical intervention for patients experiencing acute

suicidal ideation, offering a rapid-acting treatment option where conventional therapies may take longer to produce effects. <u>PsychiatryOnline</u>

# 7. Alcohol Use Disorder and Substance Use Disorders:

**Goldfine et al. (2023):** This scoping review evaluates the therapeutic use and efficacy of ketamine in treating alcohol use disorder (AUD) and alcohol withdrawal syndrome (AWS). The findings suggest that ketamine may reduce cravings and alcohol consumption in AUD patients and serve as a safe adjunct in managing AWS, potentially leading to longer abstinence rates and improved treatment outcomes. Frontiers

**Dakwar et al. (2020):** This randomized, midazolam-controlled pilot trial investigated the effects of a single ketamine infusion combined with motivational enhancement therapy (MET) on individuals with alcohol use disorder (AUD). The study found that participants receiving ketamine plus MET had a higher likelihood of abstinence and a longer time to relapse compared to the control group, suggesting that ketamine, when integrated with MET, may enhance treatment outcomes for AUD. (<u>American Journal of Psychiatry</u>) **Lee et al. (2021):** The article discusses ketamine's role in reducing cravings and preventing relapse for individuals with Substance Use Disorders (SUD), including alcohol and cocaine dependence. It suggests that ketamine, particularly when combined with psychotherapy, may improve motivation for abstinence and enhance treatment outcomes. (<u>MDPI - Brain Sciences</u>)

**Dore et al. (2019):** For substance use disorders, ketamine has a role in decreasing cravings and withdrawal symptoms, while enhancing motivation and emotional readiness for sobriety. Combined with psychotherapy, ketamine supports long-term recovery efforts. (Taylor & Francis Online)

### 8. Eating Disorders:

**Lee et al. (2021):** Preliminary findings suggest that ketamine may benefit individuals with Eating Disorders (ED), such as anorexia nervosa and binge eating disorder, by improving mood regulation and cognitive flexibility. These mechanisms address core challenges in eating disorder treatment. (<u>MDPI - Brain Sciences</u>)

**Dunlop et al. (2021):** This article reviews the potential use of ketamine in treating eating disorders, emphasizing its effects on neural circuits involved in mood regulation and reward processing. The study highlights ketamine's rapid action and its potential to address core symptoms such as rigidity and negative affect in conditions like anorexia nervosa and binge eating disorder. <u>Frontiers in Psychiatry</u>

**Calabrese et al. (2022):** This pilot study explored the combined use of a therapeutic ketogenic diet and ketamine infusions in adults with chronic anorexia nervosa. The results indicated significant improvements in eating disorder psychopathology, suggesting that this novel treatment approach may be safe and effective for addressing persistent symptoms in anorexia nervosa.

Springer Link

**Huber et al. (2022):** This study reviews the emerging evidence on ketamine's potential for treating eating disorders, such as anorexia nervosa and binge eating disorder. The findings suggest that ketamine may positively influence cognitive flexibility, mood regulation, and reward processing, which are critical factors in managing eating disorders. (MDPI - Brain Sciences)

**Tomasik et al. (2021):** This study explores the role of ketamine in treating eating disorders, particularly anorexia nervosa, highlighting its potential to address symptoms such as cognitive rigidity and mood dysregulation. The findings suggest that ketamine's impact on glutamate modulation and neural plasticity could be beneficial in improving treatment outcomes for eating disorders. (MDPI - Nutrients)