

**EMOTIONALLY UNSTABLE PERSONALITY DISORDER
(EUPD) GOOD PRESCRIBING PRACTICE**
MAY 2020

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1. Introduction

This guideline makes recommendations for prescribing in people with emotionally unstable personality disorder (EUPD). Psychotropic drugs have been shown to have only modest and inconsistent positive effects and there is no evidence that they change the course of EUPD. No drug has UK marketing authorisation for the treatment of EUPD. This guideline contains recommendations about prescribing to manage crises, comorbid conditions and insomnia.

2. Aims and objectives

To provide guidance to prescribers on the safe, effective and appropriate use of drug treatment in EUPD.

3. Scope of guideline

This guidance relates to prescribing in adult patients with a diagnosis of EUPD in primary and secondary care.

Borderline personality disorder (BPD) is also known as EUPD and is used interchangeably in this guidance.

4. Background

- NICE Guideline on the treatment of EUPD states that drug treatment should not be used specifically for borderline personality disorder (BPD) or for the individual symptoms or behaviour associated with the disorder (for example, repeated self-harm, marked emotional instability, risk-taking behaviour and transient psychotic symptoms). Antipsychotic drugs should not be used for the medium- and long-term treatment of BPD.
- Short-term use of sedative medication may be considered cautiously as part of the overall treatment plan for people with BPD in a crisis. The duration of treatment should be agreed with them, but should be no longer than 1 week
- Prescribing for EUPD is common and often patients are maintained on psychotropic medications for long periods with limited benefit and a potential burden of side effects.
- The majority of studies of drug treatment in EUPD last for only 6 weeks and a large number of different outcome measures have been used, making it difficult to evaluate and compare studies. The placebo response rate in RCTs of BPD is uniformly high. In the context of there being a very limited and inconsistent evidence base for the use of medication in EUPD, prescribing practice may be disproportionately influenced by even small RCT's, case series, and single case reports published in widely read journals.
- With patients' and carers' anxiety and frustration that something needs to be done to help them there may be an experience of pressure to prescribe for a clinician and sometimes there is a need for something to be done concretely for these patients to feel that they are being cared for.

- In those individuals with severe presentations of EUPD who experience disturbed and distressing states of mind and challenging behaviors and are unable for various reasons to engage with evidence-based interventions and psychological therapies there can be pressure on clinicians to prescribe psychotropic medication. Often it is in these cases that polypharmacy is found. In these instances as in all cases there should be clarity around the rationale for and potential efficacy when prescribing and there should be a discussion where patients are given clear information regarding this.
- There is an understanding that symptoms that are sub-threshold for the diagnosis of a co-morbid psychiatric disorder show a poor response to medication.
- Treatment concordance is often inconsistent and can lead to accumulation of unsafe stockpiles of medication given the risk of overdose in some patients with EUPD.
- Consideration should be given to the psychological meaning of prescribing for the patient and prescriber in addition to the potential impact changes in prescribing may have on the therapeutic relationships and treatment strategies longer-term.
- Prior to prescribing, patients should be given clear, detailed and understandable information about the rationale for, potential efficacy and possible side effects of the medication and treatment alternatives such as psychotherapy, so that they can make an informed decision and to enable them to manage their expectations appropriately.

5. Main prescribing principles

This guidance has been compiled with reference to POMH-UK 2012, relevant NICE guidance on Borderline Personality Disorder, Maudsley Guideline and updated research evidence.

1. Psychotropic drugs have been shown to have only modest and inconsistent positive effects and there is no evidence that they change the course of EUPD. Overall the evidence base for prescribing in EUPD is limited due to small samples with a wide range of drugs, short treatment duration, diverse outcome measures and infrequent replication of findings.
2. In the UK there are no psychotropic drugs licensed for the treatment of EUPD. Psychotropic drugs are most appropriately prescribed for the treatment of co-morbid psychiatric disorders in accordance with specific relevant guidance and should not be used routinely to treat symptoms intrinsic to EUPD. Note however that a diagnosis of BPD predicts a poorer outcome from treatment of depression with antidepressants and ECT. Symptoms of OCD in BPD may be less responsive to clomipramine (Maudsley Guidelines)
3. If psychotropic drugs are prescribed in the absence of a clear co-morbid psychiatric disorder, it should be considered an adjunct to appropriate evidence-based psychological interventions to manage specific symptoms and not instead of. The prescription should be time-limited with a clear plan to review its efficacy and discontinue if there is no response.
4. In prescribing in EUPD Polypharmacy should be avoided.
5. Due consideration should be given to the medication's potential lethality in overdose, risk of dependency and any interactions with alcohol and other psychoactive substances before a decision to prescribe is made.

6. In making decisions about commencing a psychotropic medication where possible collaboration with other professionals involved in the ongoing or longer-term care of the patient should be undertaken and consensus reached about the prescription with the main prescriber identified before prescription commences.
7. Prescribing decisions and plans should be clearly documented in a readily accessible area of the patient's health record and should be consistent with the patient's agreed care plan. Required physical health monitoring should be considered and included in this documentation.
8. Psychotropic drugs may be considered in acute crises situations where psychological interventions are not deemed sufficient. However these should be withdrawn as soon as the crisis has resolved.
9. A collaborative crisis plan should be agreed with the patient and carer (if appropriate) which indicates clearly if and which medication prescription should be considered during a crisis.
10. If prescribing is considered appropriate in the longer-term for a co-morbid condition, then it is reasonable to ask the GP to continue this, (unless a secondary care only drug is being used).

6. The management of crises

Short-term use of drug treatments may be helpful for people with EUPD during a crisis. Use a single drug and avoid polypharmacy whenever possible. Choose a drug (such as a sedative antihistamine, promethazine) that has a low side-effect profile, low addictive properties, minimum potential for misuse and relative safety in overdose. Anticipated side-effect profile and potential toxicity in overdose should guide choice. For example, benzodiazepines (particularly short-acting drugs) can cause disinhibition in this group of patients, potentially compounding problems; sedative antipsychotics can cause EPS and/or considerable weight gain, and tricyclic antidepressants are particularly toxic in overdose.

A thoughtful, joint crisis plan should help people with EUPD, as well as carers and the clinical team, manage during difficult times. This crisis plan should be easily accessible and make specific mention of whether or not medication is indicated at such times, and if so, what medication should be considered.

Discontinue a drug after a trial period if the target symptoms do not improve. Consider alternative treatments, such as psychological treatments, if target symptoms do not improve or the level of risk does not diminish.

Review after the crisis has subsided. Plan to stop drug treatment begun during a crisis, usually within 1 week. If drug treatment started during a crisis cannot be stopped within 1 week, there should be a regular review of the drug to monitor effectiveness, side effects, misuse and dependency.

When patients with EUPD present in crisis it is not an advisable time to review response to and make changes to regular prescribed medication such as anti-depressants or mood stabilisers. This should be reviewed after the period of crisis has subsided with a view to exploring the efficacy of the treatment over the longer term including periods of greater

stability and crisis and then considering making changes or discontinuing medications that aren't effective.

7. Specific medication guidance

Current practice and evidence for prescribing in EUPD, in the absence of co-morbid psychiatric disorder, tends to be organised around treatment of prominent symptom domains including affective symptoms or affect dysregulation, cognitive perceptual symptoms, impulsivity and aggression.

Antidepressants

The limited evidence base indicates that SSRI's do not have a major role in treating any of the symptom domains of EUPD. Amitriptyline has been shown not to be effective, and both may cause behavioural disinhibition. Reboxetine has also been reported to worsen symptoms. Other antidepressants require further research to inform prescribing guidance. In addition, the risk of discontinuation symptoms in the context of inconsistent compliance may be a significant factor for some patients and the discontinuation reactions and side effects may make them feel worse. Side-effects including sexual dysfunction should be considered.

Mood Stabilisers

Although NICE does not recommend routine prescription of mood stabilisers in EUPD, there is limited evidence of some effect for the prescription of sodium valproate and topiramate in reducing affective symptoms or affect dysregulation and impulsive aggression respectively. Significant risks of teratogenicity and potentially serious side-effects limit prescription however. Available evidence indicates that carbamazepine is not effective in treating EUPD, it can cause behavioral disinhibition and that its propensity to interact with other drugs is of particular importance. While lithium is licensed for aggressive and self-harming behavior its clinical utility in treating EUPD is limited due to inconsistent compliance, lethality in overdose and monitoring requirements. The Maudsley Guidelines suggests that the use of Lithium may reduce mood variation, anger, and suicidal ideation. The evidence of reduction in mood variation is based on a study from 1972 looking at 21 patients and that on anger and suicidal ideation is based on a study with a sample size of 10 in which Lithium was compared to desimipramine. Although the 2010 Cochrane review and NICE Guideline (2009) suggest that there is evidence from small scale studies to indicate the use of Lamotrigine in EUPD the recent LABILE study in 2018 has confirmed that there is no evidence of clinical or cost effectiveness in treating EUPD with Lamotrigine.

Antipsychotics

There is some evidence of positive effect for haloperidol in reducing anger when prescribed in lower doses than for psychotic disorders. However, this is based on a small number of participants. Haloperidol is known to be associated with extrapyramidal symptoms and can prolong the cardiac QTc interval. There is evidence from a small RCT that Flupenthixol Decanoate is effective in reducing suicidal behavior, but this has not been replicated. Second generation antipsychotics Aripiprazole, Olanzapine and Quetiapine have shown some effect in treating affective dysregulation and cognitive-perceptual symptoms. One small RCT of Aripiprazole showed a reduction in depression, anxiety and anger. The evidence for Clozapine is at present limited to small sample studies and case study data. However they do indicate that in patients with a personality disorder there is a reduction in self harm behavior when treated with Clozapine (Rhode et al. 2018; Zarzar et al, 2019) but there are yet to be large scale randomized controlled trials evaluating the effectiveness of clozapine in BPD. Side-effects including potential long-term metabolic effects are important considerations. In the absence of a co-morbid psychiatric disorder, the prescription should be time-limited and it should be made clear to the patient that this will be for short-term use.

Benzodiazepines and related drugs

While these drugs can be useful in improving sleep short-term and in reducing anxiety in crises situations (up to 14 days), the intrinsic risk of dependence, potential for causing disinhibition or paradoxical symptoms (e.g. aggression) and impact on memory pose a significant challenge and should not be prescribed long-term. Sedating antihistamines such as promethazine may be a suitable alternative in the first instance.

8. References

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Appendix 1

Equality impact assessment tool

	Yes/No	Comments
1. Does the policy/guidance affect one group less or more favourably than another on the basis of:		
Race	No	

	Yes/No	Comments
Ethnic origins (including gypsies and travellers)	No	
Nationality	No	
Gender	No	
Culture	No	
Religion or belief	No	
Sexual orientation including lesbian, gay and bisexual people	No	
Age	No	
Disability - learning disabilities, physical disability, sensory impairment and mental health problems	No	
2. Is there any evidence that some groups are affected differently?	No	
3. If you have identified potential discrimination, are any exceptions valid, legal and/or justifiable?	N/A	
4. Is the impact of the policy/guidance likely to be negative?	No	
5. If so can the impact be avoided?	N/A	
6. What alternatives are there to achieving the policy/guidance without the impact?	N/A	
7. Can we reduce the impact by taking different action?	N/A	