Electroconvulsive therapy, catatonia, deep vein thrombosis and anticoagulant treatment: a case report

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ABSTRACT
Electroconvulsive therapy (ECT) is considered an effective treatment for pharmacotherapy-resistant severe mental disorders. Catatonia is a complex syndrome characterised by important psychomotor disturbances. Deep vein thrombosis (DVT) and pulmonary embolism (PE) are frequent side effects of prolonged immobility in catatonic patients. Therefore, it is important to resolve the catatonia as soon as possible. ECT is the most effective therapy available and is generally considered a safe procedure. Nevertheless, its use in patients with DVT or PE and anticoagulant treatment remains controversial. We describe a case of a woman in her 40s with a previous diagnosis of bipolar disorder and dysfunctional personality traits. She was hospitalised with persecutory and reference delusions, high emotional liability, anxiety, somatisation and regressive conduct. She later developed catatonic symptoms. No progress was achieved after a month of hospitalisation, despite several pharmacological treatments. She suffered multiple complications of prolonged bed rest, such as an extensive DVT of the left common femoral, the external iliac and the common iliac veins. ECT was conducted under treatment with bemiparin. After the third administration, she showed improvement. No major bleeding or PE was developed. The safety of ECT while receiving anticoagulant therapy has been documented, though dosage and type of anticoagulant must be considered. Location of DVT (proximal or distal) may be an important topic to take into account. This report provides further evidence about the efficacy and safety of undergoing ECT in the context of concomitant serious medical conditions, such as DVT and anticoagulant therapy administration.

INTRODUCTION
Electroconvulsive therapy (ECT) might improve pharmacotherapy-resistant, severe mental disorders such as depression, mania, schizophrenia or catatonia, regardless of the underlying diagnosis. Catatonia is a complex syndrome characterised by a considerable psychomotor disturbance, which may range from marked unresponsiveness to marked agitation. Clinical presentation may include stupor, catalepsy, waxy flexibility, mutism, negativism, posturing, stereotypies, echolalia or echopraxia. It can occur in the context of various mental disorders and other medical conditions. Catatonic patients often experience prolonged periods of immobility, increasing the risk of blood stasis and, as a result, deep vein thrombosis (DVT) or a pulmonary embolism (PE). Other complications, such as decubitus ulcer, rhabdomyolysis, infections, urinary retention and flexion contractures, might also occur. Therefore, it is important to resolve the catatonia as soon as possible.

Although ECT is generally considered a safe and effective procedure for catatonic symptoms, its use in patients with DVT and anticoagulant treatment is still controversial. To provide further evidence on this topic, we report the case of a patient who presented with catatonic symptoms and extensive DVT. ECT was conducted under anticoagulation treatment, ultimately resulting in the patient showing improvement and without developing any complications.

CASE REPORT
The patient was a woman in her 40s, with a previous diagnosis of bipolar disorder and dysfunctional personality traits. She also suffered from fibromyalgia, chronic fatigue syndrome and generalised spondylarthrosis. She had two previous admissions and an outpatient follow-up since 2001 and a history of multiple psychopharmacological treatments. She was hospitalised with persecutory and reference delusions, emotional liability, anxiety, somatisations and regressive conduct. She referred to autolytic ideas, owing to her illnesses, ailments and physical limitations. At admission, she was taking quetiapine 500 mg, topiramate 50 mg, lorazepam 30 mg, clorazepate 100 mg and venlafaxine 300 mg daily; morphine and tramadol on demand. The admission blood test showed no coagulation abnormalities: prothrombin time (PT) 13.40 (reference value (RV): 11.0–15.3), activated partial thromboplastin time (aPTT) 50.40 (RV: 24.1–34.0) and antithrombin activity (AT) 94% (RV: 80–120). Catatonic symptoms were noted, including catalepsy, waxy flexibility, mutism, negativism, posturing, stereotypies, echolalia, echopraxia and echolalia. She was hospitalised with persecutory and reference delusions, high emotional liability, anxiety, somatisations and regressive conduct. She referred to autolytic ideas, owing to her illnesses, ailments and physical limitations. At admission, she was taking quetiapine 500 mg, topiramate 50 mg, lorazepam 30 mg, clorazepate 100 mg and venlafaxine 300 mg daily; morphine and tramadol on demand. The admission blood test showed no coagulation abnormalities: prothrombin time (PT) 13.40 (reference value (RV): 11.0–15.3), activated partial thromboplastin time (aPTT) 50.40 (RV: 24.1–34.0) and antithrombin activity (AT) 94% (RV: 80–120).

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The patient was discharged after 3 months without psychotic or mood symptoms. She maintained certain anxiety and chronic somatisations. She was diagnosed with histrionic personality disorder, unspecified dissociative disorder, and benzodiazepine and opioid dependence according to the International Statistical Classification of Diseases and Related Health Problems, 10th Edition. Treatment at discharge was venlafaxine 225 mg, mirtazapine 30 mg, trazadone 100 mg, pregabalin 50 mg, olanzapine 7.5 mg and bemiparin 7500 UI daily. The coagulation blood test at discharge showed PT 12.90 (RV: 11.0–15.3) s, PA 100.00% (RV: 70.0%–120.0%), INR 1.41 (RV: 0.8–1.2), APTT 39.80 (RV: 29.2–39.0) s and D-dimer (immunoturbidimetry) 3.98 (RV: 0.1–0.5) µg/mL. The Doppler of lower limbs showed DVT at the left common femoral, the left external iliac and the left common iliac veins. A computed tomography (CT) of the pulmonary artery ruled out PE. Enoxaparin was raised to 70 UI/12 hours.

No progress was achieved after a month, with the patient developing catatonic symptoms, such as immobility, mutism, negativism, posturing and stereotypies. She suffered complications owing to prolonged bedding, such as infections, autonomic dysfunction, urinary retention and constipation. Nasogastric tube placement was required to ensure nutrition and medication. In the absence of improvement, and owing to recurrent urinary retention, paliperidone was suspended. Gabapentin was also withdrawn; venlafaxine and mirtazapine were maintained and lorazepam was started up to 9 mg per day. Since benzodiazepines were not effective, ECT was considered. Owing to the theoretical risk of bleeding and PE, the case was consulted with internal medicine and neurology specialists. The patient underwent both a cranial CT scan and an MRI. No significant abnormalities were found (see online supplemental material 1). Enoxaparin was replaced by bemiparin 7500 UI once daily. After obtaining valid informed consent from the patient’s husband, bilateral ECT was started. General anaesthesia was induced by intravenous propofol (100–200 mg) and myorelaxation by succinylcholine (50–75 mg). Bemiparin was administered not less than 18 hours before each ECT session and at least 6 hours after, to prevent the risk of bleeding. The stimulus settings were based on the age and the length of seizures, measured by electroencephalogram, up to a maximum charge of 425.6 mC (85%).

After the third administration, the patient showed improvement. She was awake, communicative and cooperative. She started oral intake and walking. She showed self-limited temporary disorientation and amnesia of the episode. A total of seven ECT sessions were carried out without major bleeding or PE. Before discharge, the patient was oriented in all areas. The episodic amnesia was partially maintained but possibly related to the catatonia itself, rather than to the ECT. No other cognitive side effects were observed.

**DISCUSSION**

The use of ECT in patients with venous thromboembolism (VTE) and anticoagulant treatment is still a debated topic. Although cases of PE in patients with previous DVT who have undergone ECT sessions have been reported, the risk remains unclear. Some authors suggest that muscle contractions during a seizure can dislodge a clot from residual DVT, despite the use of succinylcholine, therefore increasing the risk of PE. Nonetheless, several reports show that ECT can be performed safely without major complications in patients with prior DVT.

It has been suggested that there may be an association between the location of the residual DVT and the onset of PE. In a recent case series, various patients with distal DVT and anticoagulant treatment received ECT without complications. Meanwhile, a patient suffering proximal DVT developed PE one day after the second ECT session. On the other hand, several cases of proximal DVT have safely undergone ECT treatment as happened with our patient. Given the inconsistency, it has been recommended that anticoagulation should be continued until the resolution of proximal DVT, before going through the ECT sessions safely. ECT is also safe in patients with PE, after proper stabilisation of the medical conditions and initiation of anticoagulant treatment.

Concerns about intracerebral haemorrhage have been raised regarding the use of ECT in patients undergoing anticoagulant therapy since ECT increases momentarily systemic and intracerebral blood pressure. In this context, the safety of ECT during the administration of anticoagulant therapy has been reported. However, two matters should be taken into consideration: the dosage and the type of anticoagulant. While

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an overdose may induce an intracerebral haemorrhage during ECT, suboptimal anticoagulation can contribute to the recurrence of thrombosis.12 Regarding the type of anticoagulant, some data demonstrate the safety of treatment with heparin,2 4 12 14 as well as with warfarin,11 while undergoing ECT. A case series of 35 patients, who had undergone a total of 300 ECT sessions while receiving long-term warfarin treatment, showed no intracerebral haemorrhage despite increases in blood pressure and pulse rate.11 However, some authors recommended the use of heparin instead of warfarin, owing to its shorter half-life, which allows its transitory discontinuation before each ECT session and its restitution afterward.12 14 For instance, Suzuki et al interrupted a patient’s heparin infusion 2 hours before each ECT session to prevent brain haemorrhage, with good results.12 Regarding our patient, the administration time of bemiparin was strictly controlled before and after ECT sessions, to minimise the bleeding risk. In any case, when warfarin or heparin is used, adequate values of INR or APTT, respectively, should be achieved to reduce the risk of bleeding.12 Recently, dabigatran, rivaroxaban, apixaban and edoxaban have emerged as direct oral anticoagulants, constituting the optimal first-line treatment for VTE in uncomplicated patients.15 They are associated with a slightly lower risk of haemorrhage compared with warfarin; hence they are a convenient alternative of treatment.9 15 Several reports have demonstrated the safety of treatment with these types of anticoagulants.2 5 10 13 16 17 For instance, a recent case series described eight patients with severe psychiatric conditions who were taking either rivaroxaban, apixaban or edoxaban for DVT. They were successfully treated with ECT, without major bleeding complications.9 It is worth noting that no evidence to date reports the optimal duration of anticoagulant treatment before ECT, which should be a focus of attention in further studies.

In conclusion, it is important to address catatonic symptoms as soon as possible to avoid medical complications. This report provides further evidence about the efficacy and safety of ECT in a patient with proximal DVT and anticoagulant treatment with bemiparin. However, the available evidence on this matter is still scarce and mostly consists of case reports or small case series. Controlled studies are necessary to support the use of ECT in patients with VTE and anticoagulation treatment.

Contributors Both authors made a significant contribution to this study and have read and approved its final version. Both authors were involved in the follow-up and treatment of the patient during her admission to the inpatient unit. AP-B obtained informed consent from the patient for the publication of the article. She also carried out the bibliographic search related to the topic and wrote the main part of the article. IS-R completed the bibliographic search and carried out the revision and correction of the text, contributing to the drafting of the article.

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