

Hematological toxicity of psychotropic medications: A Case Report

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1 **Hematological toxicity of psychotropic medications: A Case Report**

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3 **Abstract**

4 Hematological toxicity can be clinically important during psychotropic treatment.

5 We present the case of a 40-year-old Guinean man with schizoaffective disorder who
6 developed neutropenia and leukopenia after starting chlorpromazine (300 mg/day), and a
7 second neutropenic episode under carbamazepine (800 mg/day). he had never been treated
8 with antipsychotics before. Then a first hematological abnormality appeared ten days after
9 initiating chlorpromazine, and the second episode occurred three weeks after introducing
10 carbamazepine. In these situations, cessation of the suspected drug led to rapid normalization
11 of hematological values. These events were reported to the pharmacovigilance center, and
12 these medications were formally contraindicated for our patient.

13 Chlorpromazine and mood stabilizers like carbamazepine are known causes of leukopenia and
14 agranulocytosis, in this case , ethnic neutropenia was initially considered but the clear
15 temporal association between drug exposure and hematological abnormalities supported a
16 diagnosis of drug-induced neutropenia.

17 This case illustrates the importance of systematic blood count monitoring when initiating
18 psychotropic drugs known to cause hematologic adverse effects.

19 Keywords: Psychotropicmedication, Hematologicalreaction, Neutropenia,Leukopenia,
20 Chlorpromazine.

21

22 **Introduction**

23 Several psychotropic medications can cause hematological adverse effects, especially due to
24 bone marrow suppression or immune-mediated mechanisms. Clozapine is the best-known
25 example of these hematological effects , but there is other antipsychotics, particularly
26 phenothiazines, may also cause neutropenia or agranulocytosis (1).

27 Data from the AMSP European pharmacovigilance system (Arzneimittelsicherheit in der
28 Psychiatrie) program revealed various cytopenia among more than 120,000 psychiatric
29 inpatients for showing the need for routine blood monitoring (1).

30 Carbamazepine is generally used as a mood stabilizer, she has also been associated with
31 several blood adverse effects like leukopenia, anemia, thrombocytopenia, and agranulocytosis
32 (2,3).

33 the objective of this report is to illustrate the clinical and therapeutic challenges encountered
34 in managing hematological effects induced by chlorpromazine and carbamazepine, and to
35 document the importance of biological monitoring to ensure optimal patient care.

36 **Case Report**

37 A 40-year-old Guinean man with schizoaffective disorder was admitted in our hospital due a
38 symptomatic reactivation of his mental disorder. He had never treated with antipsychotics
39 before. On admission, the initial psychiatric evaluation revealed a manic syndrome
40 accompanied by delusional features and insomnia. His only medical history was chronic
41 tobacco use.

42 We started for him a treatment combining risperidone (2 mg with gradual dose escalation),
43 chlorpromazine (300 mg/day), and sodium valproate (500 mg/day) and then baseline blood
44 tests demonstrated mild leukopenia and neutropenia which were confirmed on repeat testing.
45 Chlorpromazine was therefore stopped and replaced with lorazepam (5 mg/day) then four
46 days later, the complete hematological evaluation normalized.

47 sodium valproate was replaced with carbamazepine, titrated up to 800 mg/day, because of
48 limited clinical improvement, while quetiapine (300 mg/day) and risperidone (8 mg/day)
49 were continued, and three weeks later, hematological evaluation revealed a new neutropenic
50 episode, which led to immediate discontinuation of carbamazepine, reduction of Quetiapine to
51 150 mg/day, and the rest of treatment was continued. After one week, the patient's blood
52 values had returned to normal.

53 These hematological reactions were reported to the pharmacovigilance center, and
54 chlorpromazine and carbamazepine were officially contraindicated for our patient, then due
55 to the limited clinical improvement, we started a low-dose of olanzapine (5 mg/day) with a
56 strict hematological monitoring.

57

58 **Discussion**

59 Agranulocytosis is a rare but potentially serious complication, with an incidence of roughly
60 6–8 cases per million individuals every year, and medications account for most cases (2). In
61 this case, the initial leukopenia could have been related to benign ethnic neutropenia, which is
62 more frequent in people of African origin (3), but the clear normalization of blood values after
63 discontinuation of the suspected drugs confirmed this hypothesis (3). Chlorpromazine-induced
64 neutropenia is consistent with the historical association between phenothiazines and bone
65 marrow suppression. Leukopenia occurs in approximately 0.8% of patients, and
66 agranulocytosis has been reported in 0.05%, especially in the first three months of treatment
67 (3). Carbamazepine has been linked to hematological abnormalities, a pharmaco-
68 epidemiological study from McLean Hospital involving 977 patients reported leukopenia in
69 2.1% of treated individuals, with most cases occurring in the early weeks of treatment. And
70 recovery is usually rapid approximately 6 days after the drug is stopped (3).

71 A review from Sedky and Lippmann reported hematological toxicities across multiple
72 psychotropic classes, including neutropenia, leukopenia, and agranulocytosis (4);
73 phenothiazines may cause leukopenia and agranulocytosis (4) also, atypical agents especially
74 clozapine remains the most frequently associated with severe neutropenia, while olanzapine,
75 quetiapine, and risperidone have also been implicated. Mood stabilizers cause different
76 hematological effects; lithium tends to produce leukocytosis, whereas carbamazepine is
77 implicated for inducing leukopenia, agranulocytosis, or thrombocytopenia. Valproate is
78 mainly associated with thrombocytopenia, with leukopenia representing a less common (4).

79 Within antidepressant agents, including sertraline, have rarely been associated to
80 agranulocytosis. Tricyclics have also been related with this risk, also trazodone has been
81 reported to cause leukopenia or anemia in some cases (4).

82 **Conclusion**

83 The treatment approach of patients treated with antipsychotics or mood stabilizers must
84 include individualized strategies and systematic hematological monitoring to identify the
85 adverse effects and prevent a severe complication. Every clinician should be vigilant,
86 especially during the first weeks of treatment with agents known for their potential of
87 hematologic toxicity.

88 **Conflicts of Interest**

89 The authors declare no conflicts of interest.

90 **Authors' Contributions**

91 All authors approved the final version.

92 **Patient Consent**

93 ¹ informed consent was obtained from the patient for publication.

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