

# Atrial fibrillation and unresponsiveness with muscadol - A case report

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## Introduction

Muscadol is a combination medication containing paracetamol 450mg and orphenadrine citrate 35mg used in the treatment of muscle spasms. The Paracetamol component is intended to provide analgesia, and the orphenadrine is an anticholinergic muscle relaxant. The adverse effects of paracetamol as well as orphenadrine individually are well described. Here we describe an uncommon adverse event-atrial fibrillation with unresponsiveness occurring in association with initiation of Muscadol treatment for osteoporosis.

## Case description

An 89-year old woman with no previous comorbidities other than osteoporosis was brought to our ED by ambulance after she was found unresponsive for about 30 minutes at home. There was no witnessed seizures or abnormal movements. Her physical examination included: GCS 15, T 37°C, HR 130-140/minute, BP 120/ 80mmHg, RR 16/minute, and pO<sub>2</sub> 99% on room air. Her cranial nerves were normal, pupils 6mm and reactive, mucous membranes moist, bowel sounds present, and dry skin. Her 12-lead ECG demonstrated atrial fibrillation (which was a new finding compared to her old ECG). The patient was subsequently evaluated as a case of TIA secondary to atrial fibrillation. Her lab assays including complete blood count, electrolytes, renal function test were normal. Noncontrast brain CT revealed nothing indicating ischemia or hemorrhage. Upon reviewing her medication profile it was found that the patient was recently started on Muscadol (paracetamol 450mg/orphenadrine 35mg) for osteoporosis 7 days prior to the event. Consultations to cardiology and neurology service were obtained, and these included additional investigations of echocardiogram, the result of which was normal. At 4 hours after presentation, the patient was treated with Metoprolol 25mg orally and no further doses of muscadol were given. In 72 hrs the patient reverted back to sinus rhythm, and she was discharged from the hospital.

## Discussion

The physiologic mechanism for orphenadrine to cause atrial fibrillation is uncertain, but action as an anticholinergic agent is a possible explanation. A recent study of 484 people taking orphenadrine found atrial fibrillation in 12 of them (2.48%). Of these 100% of them were female and 85.71% were >60 years. Although many conditions and medications can cause atrial fibrillation, Muscadol or any drug containing orphenadrine should be used with the understanding that atrial fibrillation is a recognized adverse event. In General drugs that increase or decrease adrenergic or vagal activity, such as sympathicomimetics, parasympathomimetics, and their inhibitors, may be able to cause AF, especially in susceptible patients with a history of cardiovascular disease (disease is the substrate, drug is the trigger), but also in "healthy" patients. These drugs represent a substantial part of cardiovascular, respiratory, and central nervous system medications.<sup>1-5</sup> Check Table for some of the examples

| Drug Class              | Representative Drugs |
|-------------------------|----------------------|
| Anti-inflammatory drugs | NSAIDs               |
| High dose               | Methylprednisolone   |
| Anti-convulsions drugs  | Lacosamide           |
| Dopamine agonist        | Apomorphine          |
| Cholinergic             | Donepezil            |
| Antipsychotics          | Olanzapine           |
| Antidepressants         | Fluoxetine           |
| Xanthine                | Aminophylline        |
| Miscellaneous           | Nicotine             |

\*\* Note: This list is not conclusive just examples.

## Conclusion

The risk of orphenadrine-associated atrial fibrillation is greatest in elderly females above 60 years of age. In our case, the atrial fibrillation was easily reversed by cessation of the drug and use of a beta blocker.

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## Conflicts of interest

Author declares there are no conflicts of interest.

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