

Date of report 28 Jul 2020

# Reported case interaction between Ritonavir and Paclitaxel

# **Drugs suspected to be involved in the DDI**

Perpetrator	Daily Dose
Ritonavir	100 (mg)
Dose adjustment performed	Administration Route
No	Oral
Start date	End date
<b>Unknown</b>	<b>Unknown</b>
Victim	Daily Dose
<b>Paclitaxel</b>	<b>Unknown</b>
Dose adjustment performed	Administration Route
No	Intravenous
Start date	End date
<b>Unknown</b>	<b>Unknown</b>

## **Complete list of drugs taken by the patient**

#### Antiretroviral treatment Abacavir/Lamivudine Darunavir (with Ritonavir or Cobicistat)

Complete list of all comedications taken by the patient, included that involved in the DDI

#### Paclitaxel, Carboplatin

#### **Clinical case description**

Gender	Age
Male	60
eGFR (mL/min) 60-30	Liver function impairment <b>No</b>

#### Description

This is about an HIV-infected patient successfully treated with ABC/3TC/DRV/r who was diagnosed with metastatic gastric cancer for which palliative treatment with paclitaxel/ carboplatin was indicated. The HIV physician and oncologist were aware of a potential interaction between DRV/r and paclitaxel (see Liverpool site) but a switch to an integrase inhibitor was not chosen due to concern about a psychiatric history in this patient. Rilpivirine was considered but not attractive due to expected use of PPIs around the chemotherapy cycles. As the Liverpool site recommended close monitonring of toxicity and uncertainty whether paclitaxel levels would go up or down, the DRV/r regimen was continued and the patient was monitored according to usual toxicity evaluation by the oncologist. The 1st cycle of chemotherapy was administered in normal dose and no

toxicity occurred. The 2nd cycle was also given in normal dose. Three days later the patient became ill, got fever, mucositis, positive blood culture for E coli, and two days later died. Obduction showed extensive mucositis, no perforation.

# Clinical Outcome Toxicity Drug Interaction Probability Scale (DIPS)

Score

2 - Possible

#### **Editorial Comment**

Paclitaxel is primarily metabolized by CYP2C8 and to a lesser extent 3A4. Darunavir/cobicistat could potentially increase paclitaxel exposure (inhibition of CYP3A4). Monitor paclitaxel induced toxicity. This case report suggests that there was a DDI leading to extensive mucositis (increased paclitaxel toxicity) that resulted in death. This is indeed possible but not substantiated (there were no levels done), and complicated by E coli bacteraemia.

### **University of Liverpool Recommendation**

• These drugs should not be coadministered

For more information <u>click here</u>