

Date of report 28 Jul 2020

Reported case interaction between Ritonavir and Paclitaxel

Drugs suspected to be involved in the DDI

Perpetrator

Ritonavir

Daily Dose

100 (mg)

Dose adjustment performed

No

Administration Route

Oral

Start date

Unknown

End date

Unknown

Victim

Paclitaxel

Daily Dose

Unknown

Dose adjustment performed

No

Administration Route

Intravenous

Start date

Unknown

End date

Unknown

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) Liver function impairment

60-30 No

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

