

Date of report 21 Dec 2021

# Reported case interaction between Ritonavir and Daclatasvir

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

Perpetrator	Daily Dose
Ritonavir	100 (mg)
Dose adjustment performed No	Administration Route Oral
Start date	End date
June 13, 2016	Ongoing
Victim	Daily Dose
<b>Daclatasvir</b>	60 (mg)
Dose adjustment performed	Administration Route
No	Oral
Start date	End date
Sept. 8, 2020	Ongoing

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

#### Antiretroviral treatment Atazanavir (with Ritonavir or Cobicistat) Emtricitabine/Tenofovir-DF

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

Metformin 850 mg OD, Sofosbuvir 300mg/Daclatasvir 60mg

# **Clinical case description**

Gender	Age
Male	32
eGFR (mL/min)	Liver function impairment
>60	Yes

Child-Pugh Child-Pugh B

Description

#### **Social Hx:**

- Injecting drug user
- Alcoholic
- Heavy smoker

#### Medical Hx:

- Living with HIV five years ago and on HAART
- Diagnosed with DM type II 2 years ago and on Metformin Upon follow up after 4 weeks starting Hep C (Tab Sofosbuvir 300mg/Daclatasvir 60mg) :
- Patient complaint easy feel tired and lose of appetite
- Physical examination jaundice
- Elevated ALT (71 U/L), anaemia (HB = 9 d/L)

- Blood glucose plasma level = 11 mm/mol
  Suspect:
- Toxicity of Daclatasvir (CYP3A4 metabolite) due to inhibition of Ritonavir (CYP3A4 inhibitor) decrease the elimination and remain in the plasma.
- Action is to reduce the Daclatasvir dose from 60mg to 30mg.

# Clinical Outcome Toxicity Drug Interaction Probability Scale (DIPS)

#### Score

### 6 - Probable

## **Editorial Comment**

This interaction is highlighted in the Daklinza Prescribing Information ie a PK study is described showing a decrease in exposure of daclatasvir (non dose normalized data) with the recommendation to 'decrease the daclatasvir dose to 30 mg once daily'. This case highlights the importance of the prescriber/health care professional having ready access to the relevant information before starting treatment. This patient should not have been on daclatasvir 60 mg. Note that hepatic impairment (CP-A, CP-B or CP-C) has no effect on daclatasvir exposure unlike the major impact on HCV protease inhibitors. This patient was CP-B.

# **University of Liverpool Recommendation**

Potential interaction - may require close monitoring, alteration of drug dosage or timing of administration

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