

Guidelines for the use of PF-07321332 plus Ritonavir (Paxlovid) for non-hospitalised patients with COVID-19

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Commissioning position

Antivirals or neutralising monoclonal antibodies (nMABs) are recommended to be available as a treatment option through routine commissioning for non-hospitalised adults with COVID-19 treated in accordance with the criteria set out in this document.

This policy applies to non-hospitalised patients with COVID-19 who are symptomatic and showing no evidence of clinical recovery and covers the following treatment options:

- First-line: PF-07321332 plus ritonavir (Paxlovid, antiviral) OR Sotrovimab (nMAB), as clinically indicated
- Second-line: Molnupiravir (antiviral)

Combination treatment with an nMAB and an antiviral is NOT routinely recommended

Where patients are ineligible for treatment under this policy, recruitment to the [PANORAMIC trial](#), which is building the evidence for novel oral antivirals in a broader cohort of at risk patients, should be supported.

PF-07321332 plus ritonavir

Evidence

Final results from the EPIC HR trial indicate that the dual oral antiviral PF-07321332 (nirmatrelvir) plus ritonavir resulted in a relative risk reduction of hospitalisation or death by 89% (within 3 days of symptom onset) and 88% (within 5 days of symptom onset) compared to placebo in non-hospitalised, high-risk adults with COVID-19.

Marketing authorisation

PF-07321332 plus ritonavir administered orally has conditional marketing authorisation in Great Britain (England, Scotland and Wales) for the treatment of COVID-19 in adults who do not require supplemental oxygen and who are at increased risk for progression to severe COVID-19.

Eligibility criteria

Patients must meet all of the eligibility criteria and none of the exclusion criteria. Non-hospitalised patients are eligible for treatment if:

- SARS-CoV-2 infection is confirmed by either:
 - o Polymerase chain reaction (PCR) testing OR
 - o Lateral flow test (registered via gov.uk or NHS 119)within the last **5 days**
- AND
- Symptomatic with COVID-19 and showing no signs of clinical recovery
- AND
- Onset of symptoms of COVID-19 within the last **7 days**¹
- AND
- A member of a 'highest' risk group (as defined in Appendix 1).

Exclusion criteria

- Pregnancy or possibility of being pregnant
- Requirement for hospitalisation for COVID-19
- The pattern of clinical presentation indicates that there is recovery rather than risk of deterioration from infection
- Swallowing difficulties and difficulty with compliance to a complex dosage 5 day course of tablets
- The patient has a history of advanced decompensated liver cirrhosis or stage 3-5 chronic kidney disease
- < 18years of age
- The patient is taking any of the medications listed in Appendix 2

Dose and administration

The recommended dose of PF-07321332 plus ritonavir is 300mg (two 150mg tablets) PF-07321332 with 100mg (one 100mg tablet) ritonavir taken together orally twice daily for 5 days.

PF-07321332 plus ritonavir should be given as soon as possible after positive results of direct SARS-CoV-2 viral testing and within 7 days of onset of symptoms. Clinicians should assure themselves that patients are able to swallow the oral tablets.

A missed dose should be taken as soon as possible and within 8 hours of the scheduled time, and the normal dosing schedule should be resumed. If more than 8 hours has elapsed, the missed dose should not be taken and the treatment should resume according to the normal dosing schedule.

If a patient requires hospitalisation due to severe or critical COVID-19 after starting treatment with PF-07321332 plus ritonavir, the patient should complete the full 5-day treatment course at the discretion of his/her healthcare provider.

1. **Treatment commencement may be extended up to a maximum of 7 days from symptom onset if clinically indicated (treatment commencement beyond 5 days from symptom onset is off-label)**

Importance of medication history taking

It is of utmost importance that a patient's current medication is considered before prescribing PF-07321332 plus ritonavir as many patients will be ineligible for treatment. See Appendix 2.

Ineligibility could be due to a contraindication or where strong advice to not use concomitantly exists. In these circumstances PF-07321332 plus ritonavir must NOT be prescribed together with the patient's medication. However, if the factors impacting decisions about risk assessment are met and the prescriber is confident of appropriate follow up, management options can be considered e.g. temporary stopping of concomitant medication.

Further management options are available via the **University of Liverpool COVID-19 Drug Interaction checker**- <https://www.covid19-druginteractions.org/>

Drug Interactions and Cautions

Please refer to the Summary of Product Characteristics (SmPC) for PF-07321332 plus ritonavir for special warnings and precautions for use at

<https://www.medicines.org.uk/emc/product/13145/smpc#gref>

PF-07321332 plus ritonavir has a risk of serious adverse reactions due to interactions with other medicinal products (see Appendix 2 for a list of these products).

Initiation of PF-07321332 plus ritonavir, a CYP3A inhibitor, in patients receiving medicinal products metabolised by CYP3A or initiation of medicinal products metabolised by CYP3A in patients already receiving PF-07321332 plus ritonavir, may increase plasma concentrations of medicinal products metabolised by CYP3A. Initiation of medicinal products that inhibit or induce CYP3A may increase or decrease concentrations of PF-07321332 respectively.

These interactions may lead to:

- Clinically significant adverse reactions, potentially leading to severe, life-threatening or fatal events from greater exposures of concomitant medicinal products.
- Clinically significant adverse reactions from greater exposures of PF-07321332 plus ritonavir.
- Loss of therapeutic effect of PF-07321332 plus ritonavir and possible development of viral resistance.

Use the **University of Liverpool COVID-19 Drug Interaction checker**- <https://www.covid19-druginteractions.org/> to confirm that patients current medications DO NOT interact with PF-07321332 plus ritonavir before prescribing.

Also refer to Appendix 2 which contains a list of where PF-07321332 plus ritonavir is not an appropriate option.

Hepatic transaminase elevations, clinical hepatitis and jaundice have occurred in patients receiving ritonavir. Therefore PF-07321332 plus ritonavir should NOT be prescribed to patients with pre-existing liver diseases, liver enzyme abnormalities or hepatitis.

Pregnancy and women of childbearing potential

Paxlovid (PF-07321332 and Ritonavir) is not recommended during pregnancy. There are no human data on the use of PF-07321332 plus ritonavir during pregnancy to inform the drug-associated risk of adverse developmental outcomes, women of childbearing potential should avoid becoming pregnant during treatment with PF-07321332 plus ritonavir. PF-07321332 plus ritonavir is **not recommended** during pregnancy and in women of childbearing potential not using effective contraception.

Use of ritonavir may reduce the efficacy of combined hormonal contraceptives. Patients using combined hormonal contraceptives should be advised to use an effective alternative contraceptive method or an additional barrier method of contraception during treatment and until after one complete menstrual cycle after stopping PF-07321332 plus ritonavir.

Safety reporting

It is vital that any suspected adverse reactions (including congenital malformations and/or neurodevelopmental problems following treatment during pregnancy) are reported directly to the MHRA via the new dedicated COVID-19 Yellow Card reporting site at: <https://coronavirus-yellowcard.mhra.gov.uk/>.

Reference

1. MHRA CAS Alert: Antivirals or neutralising monoclonal antibodies (nMABs) for non-hospitalised patients with COVID-19 - <https://www.cas.mhra.gov.uk/ViewandAcknowledgment/ViewAlert.aspx?AlertID=103191>
2. Pfizer Press Release <https://www.pfizer.com/news/press-release/press-release-detail/pfizer-announces-additional-phase-23-study-results>

Medicines Governance Committee Statement on antivirals and nMABS for COVID-19 - February 10th 2022:

Please note:

- For Paxlovid, conditional marketing authorisation has been given by the MHRA and the Department of Health, through the NHS, has approved its use. COVID-19 Medicines Delivery Units (CMDUs) can therefore prescribe the Paxlovid for patients. Full results of the EPIC HR trial awaited but there is currently enough information to guide safe prescribing decisions.
- The nMABs and antivirals for COVID-19 are expected to work against the omicron variant and further evidence is emerging.
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- As further evidence emerges clinicians will be kept informed.

Appendix 1: Patient cohorts considered at highest risk from COVID-19 and to be prioritised for treatment with nMABs.

The following patient cohorts were determined by an independent advisory group commissioned by the Department of Health and Social Care (DHSC).

Cohort	Description
Down's syndrome	All patients with Down's syndrome
Patients with a solid cancer	<ul style="list-style-type: none"> • Active metastatic cancer and active solid cancers (at any stage) • All patients receiving chemotherapy within the last 3 months • Patients receiving group B or C chemotherapy 3-12 months prior (see Appendix 3) • Patients receiving radiotherapy within the last 6 months
Patients with haematological diseases and stem cell transplant recipients	<ul style="list-style-type: none"> • Allogeneic haematopoietic stem cell transplant (HSCT) recipients in the last 12 months or active graft vs host disease (GVHD) regardless of time from transplant (including HSCT for non-malignant diseases) • Autologous HSCT recipients in the last 12 months (including HSCT for non-malignant diseases) • Individuals with haematological malignancies who have <ul style="list-style-type: none"> ○ received chimaeric antigen receptor (CAR)-T cell therapy in the last 24 months, or ○ radiotherapy in the last 6 months • Individuals with haematological malignancies receiving systemic anti-cancer treatment (SACT) within the last 12 months except patients with chronic phase chronic myeloid leukaemia (CML) in molecular response or first or second line tyrosine kinase inhibitors (TKI). • All patients with myeloma (excluding MGUS) or chronic B-cell lymphoproliferative disorders (e.g. chronic lymphocytic leukaemia, follicular lymphoma) or myelodysplastic syndrome (MDS) who do not fit the criteria above. • All patients with sickle cell disease. • Individuals with non-malignant haematological disorder (e.g. aplastic anaemia or paroxysmal nocturnal haemoglobinuria) receiving B-cell depleting systemic treatment (e.g. anti-CD20, anti-thymocyte globulin [ATG] and alemtzumab) within the last 12 months.

Patients with renal disease	<ul style="list-style-type: none"> • Renal transplant recipients (including those with failed transplants within the past 12 months), particularly those who: <ul style="list-style-type: none"> ◦ Received B cell depleting therapy within the past 12 months (including alemtuzumab, rituximab [anti-CD20], anti-thymocyte globulin) ◦ Have an additional substantial risk factor which would in isolation make them eligible for nMABs or oral antivirals ◦ Not been vaccinated prior to transplantation • Non-transplant patients who have received a comparable level of immunosuppression • Patients with chronic kidney stage (CKD) 4 or 5 (an eGFR less than 30 ml/min/1.73m²) without immunosuppression
Patients with liver disease	<ul style="list-style-type: none"> • Patients with cirrhosis Child's-Pugh class B and C (decompensated liver disease). • Patients with a liver transplant • Liver patients on immune suppressive therapy (including patients with and without liver cirrhosis) • Patients with cirrhosis Child's-Pugh class A who are not on immune suppressive therapy (compensated liver disease)
Patients with immune-mediated inflammatory disorders (IMID)	<ul style="list-style-type: none"> • IMID treated with rituximab or other B cell depleting therapy in the last 12 months • IMID with active/unstable disease on corticosteroids, cyclophosphamide, tacrolimus, cyclosporin or mycophenolate. • IMID with stable disease on either corticosteroids*, cyclophosphamide, tacrolimus, cyclosporin or mycophenolate. • IMID patients with active/unstable disease including those on biological monotherapy and on combination biologicals with thiopurine or methotrexate
Immune deficiencies	<ul style="list-style-type: none"> • Common variable immunodeficiency (CVID) • Undefined primary antibody deficiency on immunoglobulin (or eligible for Ig) • Hyper-IgM syndromes • Good's syndrome (thymoma plus B-cell deficiency) • Severe Combined Immunodeficiency (SCID) • Autoimmune polyglandular syndromes/autoimmune polyendocrinopathy, candidiasis, ectodermal dystrophy (APECED syndrome) • Primary immunodeficiency associated with impaired type I interferon signalling • X-linked agammaglobulinaemia (and other primary agammaglobulinaemias) • Any patient with a secondary immunodeficiency

	receiving, or eligible for, immunoglobulin replacement therapy
HIV/AIDS	<ul style="list-style-type: none"> • Patients with high levels of immune suppression, have uncontrolled/untreated HIV (high viral load) or present acutely with an AIDS defining diagnosis • On treatment for HIV with CD4 <350 cells/mm³ and stable on HIV treatment or CD4>350 cells/mm³ and additional risk factors (e.g. age, diabetes, obesity, cardiovascular, liver or renal disease, homeless, those with alcohol-dependence)
Solid organ transplant recipients	All recipients of solid organ transplants not otherwise specified above
Rare neurological conditions	<ul style="list-style-type: none"> • Multiple sclerosis • Motor neurone disease • Myasthenia gravis • Huntington's disease

Appendix 2: Drug-drug interactions involving PF-07321332 (nirmatrelvir) plus ritonavir¹⁰

Table 1 below lists medicines in alphabetical (by generic name) order indicating that concurrent prescribing of PF-07321132 (nirmatrelvir) plus ritonavir is not an appropriate option. This list is not comprehensive. If a medicine is not listed also check [University of Liverpool COVID-19 Drug Interaction checker](https://covid19-druginteractions.org/checker) (<https://covid19-druginteractions.org/checker>).

The last column gives an indication of when advice to not prescribe together applies or where risks and benefits need careful consideration taking account of the practicalities of managing such patients in a CMDU or non-specialist setting.

Table 1: Alphabetical (by generic name) list of medicines indicating that PF-07321132 (nirmatrelvir) plus ritonavir is not an appropriate option to be prescribed together.

Specific medicines	Medicine used for	Use of PF-07321132 (nirmatrelvir) plus ritonavir
Abemaciclib	Cancer	Consider risks and benefits
Acalabrutinib	Cancer	Consider risks and benefits
Alfuzosin	Prostate gland enlargement	Do not use
Aliskiren	High blood pressure (hypertension)	Do not use*
Amiodarone	Irregular heartbeats	Do not use
Apalutamide	Cancer	Consider risks and benefits
Apixaban	Treating or preventing blood clots	Do not use
Avanafil	Erection problems	Do not use
Bedaquiline	Infections	Consider risks and benefits
Bosentan	Pulmonary arterial hypertension	Do not use
Budesonide (inhaled, nasal spray)	Relieving asthma or COPD, or cold-like symptoms caused by allergic rhinitis	Consider risks and benefits
Carbamazepine	Epilepsy, nerve pain or trigeminal neuralgia	Do not use
Ceritinib	Cancer	Consider risks and benefits
Ciclosporin	Immunosuppressant	Do not use
Cisapride	Gastrointestinal motility problems	Do not use
Clonazepam	Epilepsy or anxiety	Do not use
Clopidogrel	Treating or preventing blood clots	Do not use*
Clozapine	Schizophrenia	Do not use
Colchicine	Gout	Do not use
Contraception, hormonal	Contraception	Consider risks and benefits
Dabigatran	Treating or preventing blood clots	Consider risks and benefits
Delamanid	Infections	Consider risks and benefits
Dexamphetamine	Narcolepsy or attention deficit hyperactivity disorder (ADHD)	Consider risks and benefits
Diazepam	Anxiety, muscle spasms or fits	Do not use
Digoxin	Irregular heartbeats or heart failure	Consider risks and benefits
Dihydroergotamine	Cluster headaches	Do not use
Disopyramide	Irregular heartbeats	Do not use*
Dronedarone	Irregular heartbeats	Do not use
Eletriptan	Migraines	Consider risks and benefits
Encorafenib	Cancer	Consider risks and benefits
Enzalutamide	Cancer	Consider risks and benefits
Eplerenone	Heart failure	Do not use*
Ergotamine	Cluster headaches	Do not use

¹⁰ The information in this appendix is based on SPS guidance and is correct at the time of publication. Please refer to the SPS [guidance](#) for the most up to date information.

Specific medicines	Medicine used for	Use of PF-07321132 (nirmatrelvir) plus ritonavir
Everolimus	Cancer or immunosuppressant	Do not use
Exviera (contains dasabuvir)	Hepatitis C	Consider risks and benefits
Fentanyl	Pain	Consider risks and benefits
Flecainide	Irregular heartbeats	Do not use
Flurazepam	Anxiety or problems sleeping	Do not use
Fluticasone propionate (inhaled or nasal spray)	Relieving asthma or COPD Cold-like symptoms caused by allergic rhinitis	Consider risks and benefits
Fostamatinib	Blood disorder	Consider risks and benefits
Fusidic acid (oral)	Infections	Do not use
Ibrutinib	Cancer	Consider risks and benefits
Illegal drugs	Substance abuse	Check advice in University of Liverpool COVID-19 Drug Interaction checker
Ivabradine	Heart failure or angina	Do not use*
Ketoconazole	Infections	Consider risks and benefits
Lamotrigine	Epilepsy or bipolar disorder	Consider risks and benefits
Lercanidipine	High blood pressure (hypertension)	Do not use*
Letemovir	Transplant	Consider risks and benefits
Levothyroxine	Underactive thyroid (hypothyroidism)	Consider risks and benefits
Lomitapide	Lowering cholesterol	Do not use
Lurasidone	Schizophrenia	Do not use
Maviret (contains glecaprevir and pibrentasvir)	Hepatitis C	Do not use
Methadone	Heroin dependence	Consider risks and benefits
Methylphenidate	Narcolepsy or attention deficit hyperactivity disorder (ADHD)	Consider risks and benefits
Midazolam	Epilepsy	Do not use
Neratinib	Cancer	Do not use
Pethidine	Pain	Do not use
Phenobarbital	Epilepsy	Do not use
Phenytoin	Epilepsy	Do not use
Pimozide	Schizophrenia	Do not use
Piroxicam	Pain	Do not use
Propafenone	Irregular heartbeats	Do not use
Propoxyphene	Analgesics	Do not use
Quetiapine	Bipolar disorder, depression, schizophrenia	Do not use
Quinidine	Antiarrhythmic	Do not use
Ranolazine	Heart failure or angina	Do not use
Rifabutin	Infections	Consider risks and benefits
Rifampicin	Infections	Do not use
Riociguat	Pulmonary arterial hypertension	Consider risks and benefits
Rivaroxaban	Treating or preventing blood clots	Do not use
Rosuvastatin	Lowering cholesterol	Consider risks and benefits
Salmeterol (inhaled)	Relieving asthma or COPD	Do not use
Sildenafil	Erection problems or pulmonary arterial hypertension	Do not use
Simvastatin	Lowering cholesterol	Do not use
Sirolimus	Immunosuppressant	Do not use*
Sodium fusidate (oral)	Infections	Do not use
St. John's Wort (Hypericum perforatum)	Herbal medicine	Do not use
Tacrolimus	Immunosuppressant	Do not use

Specific medicines	Medicine used for	Use of PF-07321132 (nirmatrelvir) plus ritonavir
Tadalafil	Erection problems or pulmonary arterial hypertension	Do not use
Theophylline	Relieving asthma or COPD	Consider risks and benefits
Ticagrelor	Treating or preventing blood clots	Do not use*
Vardenafil	Erection problems	Do not use
Valproic acid	Bipolar disorder, epilepsy or migraine	Consider risks and benefits
Venetoclax	Cancer	Do not use
Viekirax (contains ombitasvir, paritaprevir and ritonavir)	Hepatitis C	Consider risks and benefits
Vinblastine	Cancer	Consider risks and benefits
Vincristine	Cancer	Consider risks and benefits
Voriconazole	Infections	Consider risks and benefits
Warfarin	Treating or preventing blood clots	Consider risks and benefits
Zepatier (contains elbasvir and grazoprevir)	Hepatitis C	Do not use*

*Not listed in PF-07321132 (nirmatrelvir) plus ritonavir SmPC but use NOT advised by [COVID-19 Drug Interaction checker](#)

Table 2 below lists medicines by what they are used for indicating when PF-07321132 (nirmatrelvir) plus ritonavir is not an appropriate option to be prescribed concurrently. This list is not comprehensive. If a medicine is not listed also check [University of Liverpool COVID-19 Drug Interaction checker](#) (<https://covid19-druginteractions.org/checker>).

The last column gives an indication of when advice to not prescribe together applies or where risks and benefits need careful consideration taking account of the practicalities of managing such patients in a CMDU or non-specialist setting.

Table 2: Medications interacting with PF-07321132 (nirmatrelvir) plus ritonavir listed by use

What the medicine is used for	Specific medicines	Use of PF-07321132 (nirmatrelvir) plus ritonavir
Underactive thyroid (hypothyroidism)	Levothyroxine	Consider risks and benefits
Lowering cholesterol	Lomitapide	Do not use
	Rosuvastatin	Consider risks and benefits
	Simvastatin	Do not use
Treating or preventing blood clots	Apixaban	Do not use
	Clopidogrel	Do not use*
	Dabigatran	Consider risks and benefits
	Rivaroxaban	Do not use
	Ticagrelor	Do not use*
	Warfarin	Consider risks and benefits
Relieving asthma or COPD (inhaled or oral)	Budesonide	Consider risks and benefits
	Fluticasone propionate	Consider risks and benefits
	Salmeterol	Do not use
	Theophylline	Consider risks and benefits
Bipolar disorder, schizophrenia, epilepsy, migraine or cluster headaches	Carbamazepine	Do not use
	Clonazepam	Do not use
	Clozapine	Do not use
	Dihydroergotamine	Do not use
	Eletriptan	Consider risks and benefits
	Ergotamine	Do not use
	Lamotrigine	Consider risks and benefits
	Lurasidone	Do not use
	Phenobarbital	Do not use
	Phenytoin	Do not use

	Pimozide Quetiapine Valproic acid Midazolam	Do not use Do not use Consider risks and benefits Do not use
Erection problems	Avanafil Sildenafil Tadalafil Vardenafil	Do not use Do not use Do not use Do not use
Contraception, hormonal	Elicit name of medication and check COVID-19 Drug Interaction checker .	Consider risks and benefits
Irregular heartbeats	Amiodarone Digoxin Disopyramide Dronedarone Flecainide Propafenone Quinidine	Do not use Consider risks and benefits Do not use* Do not use Do not use Do not use Do not use
High blood pressure (hypertension)	Aliskiren Lercanidipine	Do not use* Do not use*
Prostate gland enlargement	Alfuzosin	Do not use
Cold-like symptoms caused by allergic rhinitis (nasal spray)	Budesonide Fluticasone propionate	Consider risks and benefits Consider risks and benefits
Pain	Fentanyl Midazolam Pethidine Propoxyphene Piroxicam	Consider risks and benefits Do not use Do not use Do not use Do not use
Nerve pain or trigeminal neuralgia	Carbamazepine	Do not use
Heart failure or angina	Eplerenone Ivabradine Ranolazine Digoxin	Do not use* Do not use* Do not use Consider risks and benefits
Gout	Colchicine	Do not use
Heroin dependence	Methadone	Consider risks and benefits
Substance abuse	Various illicit drugs	Check COVID-19 Drug Interaction checker
Herbal medicines	St. John's Wort (<i>Hypericum perforatum</i>)	Do not use
Infections	Bedaquiline Delamanid Fusidic acid/ sodium fusidate (oral) Ketoconazole Rifabutin Rifampicin Voriconazole	Consider risks and benefits Consider risks and benefits Do not use Consider risks and benefits Consider risks and benefits Do not use Consider risks and benefits
Pulmonary arterial hypertension (PAH)	Bosentan Riociguat Sildenafil (Revatio) Tadalafil	Do not use* Consider risks and benefits Do not use Do not use
Anxiety, problems sleeping, muscle spasms, fits, attention deficit hyperactivity disorder (ADHD) or narcolepsy	Diazepam Flurazepam Clonazepam St John's Wort Dexamphetamine Methylphenidate	Do not use Do not use Do not use Do not use Consider risks and benefits Consider risks and benefits
Immunosuppressant medicines which	Ciclosporin	Do not use*

can be used for a range of conditions	Everolimus Sirolimus Tacrolimus	Do not use* Do not use* Do not use*
Transplant	Letemovir	Consider risks and benefits
Hepatitis C	Exviera (contains dasabuvir) Maviret (contains glecaprevir and pibrentasvir) Viekirax (contains ombitasvir, paritaprevir and ritonavir) Zepatier (contains elbasvir and grazoprevir)	Consider risks and benefits Do not use Consider risks and benefits Do not use*
Cancer	Abemaciclib Acalabrutinib Apalutamide Ceritinib Encorafenib Enzalutamide Everolimus Ibrutinib Neratinib Venetoclax Vinblastine Vincristine	Consider risks and benefits Consider risks and benefits Consider risks and benefits Consider risks and benefits Consider risks and benefits Consider risks and benefits Do not use Consider risks and benefits Do not use Do not use Consider risks and benefits Consider risks and benefits
Blood disorders	Fostamatinib	Consider risks and benefits

*Not listed in PF-07321132 (nirmatrelvir plus ritonavir SmPC but use NOT advised by [COVID-19 Drug Interaction checker](#)

Appendix 3: Chemotherapy agents (Groups B and C)

Patients currently on or who have received the following chemotherapy regimens (Groups B and C in the table below) in the last 12 months and are considered to be at higher risk of Grade 3/4 febrile neutropenia or lymphopenia.

Group B 10-50% risk of grade 3/4 febrile neutropenia or lymphopenia	Group C >50% risk of grade 3/4 febrile neutropenia or lymphopenia
<ul style="list-style-type: none"> • Etoposide based regimens • CMF • Irinotecan and Oxaliplatin based regimens • Cabazitaxel • Gemcitabine • Chlorambucil • Temozolomide • Daratumumab • Rituximab • Obinutuzumab • Pentostatin • Proteasome inhibitors • IMiDs • PI3Kinase inhibitors • BTK inhibitors • JAK inhibitors • Venetoclax • Trastuzumab-emtansine • Anthracycline-based regimens • Fluorouracil, epirubicin and cyclophosphamide (FEC) • Methotrexate, vinblastine, adriamycin/doxorubicin, cisplatin (MVAC) • Adriamycin/doxorubicin, bleomycin, vinblastine, dacarbazine (ABVD) • Cyclophosphamide, doxorubicin, vincristine, prednisolone (CHOP) • Bleomycin, etoposide, doxorubicin, cyclophosphamide, vincristine, procarbazine and prednisolone (BEACOPP) • Liposomal doxorubicin • Taxane – 3-weekly • Nab-paclitaxel • Carboplatin-based regimens • Ifosfamide-based regimens • Bendamustine • Cladribine • Topotecan • Cyclophosphamide/Fludarabine combinations • Ifosfamide, carboplatin, etoposide (ICE) • Gemcitabine, dexamethasone, cisplatin (GDP) • Isatuximab • Polatuzumab • Acalabrutinib • Dexamethasone, cytarabine, cisplatin (DHAP) • Etoposide, methylprednisolone, cytarabine, 	<ul style="list-style-type: none"> • All acute myeloid leukaemia/acute lymphocytic regimens • Bleomycin, etoposide and platinum • Highly immunosuppressive chemotherapy (e.g. FluDAP, high dose Methotrexate & Cytarabine) • Trifluradine/ Tipiracil • KTE-X19 • Gilteritinib

<p>cisplatin (ESHAP)</p> <ul style="list-style-type: none">• Cyclophosphamide, vincristine, doxorubicin, dexamethasone (CVAD)• Dacarbazine-based regimens• Lomustine• Magalizumab• Brentuximab vedotin• Asparaginase-based regimens	
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