

*Supplementary Materials*
**Table S1.** Characteristics of the reviewed 17 studies.

Publication	Type of Study	Inclusion Criteria	Exclusion Criteria	No. of participants	Geographic region	Study period
[1]	Retrospective study	18+, hospitalized		92	Saudi-Arabia	Prescription between January and December 2022
[2]	Observation, retrospective cohort study	40+, confirmed COVID-19 infection, considered high risk for severe disease	Treatment with molnupiravir, patients residing in long-term care facilities, patients hospitalized during the study period but before the positive SARS-CoV-2 test result	109,254	Israel	Jan 9, 2022 until March 31, 2022
[3]	Retrospective cohort study	18+, one or more risk factors for severe COVID-19 as per CDC guidelines (including 50+ years old), presence of underlying medical conditions associated with a conclusive higher risk of severe COVID-19	<18, patients hospitalized on or before the COVID-19 index date or date of treatment with nirmatrelvir/ritonavir, patients receiving nirmatrelvir/ritonavir before their index date, late beginning of treatment (later than 5 days after the patient's index date)	410,642	USA	Positive SARS-CoV-2 test or COVID-19 diagnosis between December 23, 2021 and December 31, 2022
[4]	Retrospective cohort study	18+, hospitalized and eGFR < 90ml/min (chronic kidney disease)	Hemodialyzed patients, peritoneal patients, pregnancy and breastfeeding, patients taking drugs such as amiodarone, propafenone and warfarin	1,279	China	April 7, 2022 and June 21, 2022
[5]	Retrospective controlled study	High-risk outpatients (patients with comorbidities)		386	Italy	Diagnosis between February 1, 2022 and May 31, 2022

Publication	Type of Study	Inclusion Criteria	Exclusion Criteria	No. of participants	Geographic region	Study period
		and/or immune depression)				
[6]	Population-based cohort study	50+, nonhospitalized, no contraindications for nirmatrelvir plus ritonavir	PCR positive in the 90 days before the diagnosis, patients with an estimated glomerular filtration rate less than 30mL/min	44,551	USA	Diagnosis between Jan 1, 2022 and July 17, 2022
[7]	Retrospective cohort study	18-50 years old, vaccination, development of COVID-19 at least one month after vaccination	Treatment with other approved COVID-19 medications (monoclonal antibodies, convalescent plasma, molnupiravir)	5,094	USA	December 1, 2021 until July 30, 2022
[8]	Comparative retrospective cohort study	18+, vaccinated, development of COVID-19 at least one month after vaccination	Treatment with monoclonal antibodies, convalescent plasma, molnupiravir; receipt of nirmatrelvir/ritonavir more than five days after diagnosis; patients requiring initial hospitalization	2,260		Diagnosis between Dec 1, 2021 and Apr 18, 2022
[9]	phase 2-3, double-blind, randomized, placebo-controlled trial	18+, confirmed SARS-CoV-2 infection, symptom onset of no more than five days before randomization, have at least one sign or symptom of COVID-19 on the day of randomization and at least one characteristic or coexisting condition associated with	Previous vaccination or receipt of convalescent plasma, previous confirmed COVID-19 infection or hospitalization for COVID-19, and anticipated need for hospitalization within 48 hours of randomization	2,246	Participants from the United States, Bulgaria, South Africa, Brazil, India, Mexico, Ukraine, Turkey, Japan, Spain, Russia, Argentina, Colombia, Poland, South Korea, Hungary, Taiwan,	"Delta dominance"

Publication	Type of Study	Inclusion Criteria	Exclusion Criteria	No. of participants	Geographic region	Study period
		high risk of progression to severe COVID-19.			Malaysia, Czech Republic, Thailand and Puerto Rico	
[10]	Retrospective cohort study	18+, at least one risk factor for severe COVID-19	Contraindicated medical conditions, usage of one or more contraindicated medications, hospitalization within 3 days of index	1,029,910	USA	Positive test or diagnosis between December 2021 and February 2023
[11]	Retrospective cohort study	12+, prescription within 5 days of symptom onset, untreated patients who met the same eligibility criteria as in the nirmatrelvir/ritonavir guidelines (60+ OR 12+ (weighing at least 40kg) with underlying disease and/or immunosuppression, no supplemental oxygen)	Treatment with molnupiravir or remdesivir, < 12, missing data in basic epidemiological investigation and patients who did not meet the nirmatrelvir/ritonavir eligibility criteria	1,936,925	South Korea	Diagnosis between July 1 until November 30, 2022
[12]	Randomized controlled trial	18-90 years old, hospitalized with severe comorbidities, admission within 48 hours of positive PCR test	History of active liver disease, patients on dialysis or EGFR < 45 mL/min/1.73m <sup>2</sup> within the past 6 months, HIV infection, history of allergy to nirmatrelvir/ritonavir	264	China	April 10, 2022 until May 19, 2022
[13]	Restrospective cohort study	18+, first-ever positive COVID-19 test, at least one comorbidity or condition	Treatment with molnupiravir, late (> 5 days after positive test) beginning of	180,351	Israel	Diagnosis between Jan 1, 2022 and Feb 28, 2022

Publication	Type of Study	Inclusion Criteria	Exclusion Criteria	No. of participants	Geographic region	Study period
		associated with high risk for severe COVID-19 (including age 60+ years, body mass index (BMI) $\geq 30$ kg/m <sup>2</sup> , diabete, hypertension, cardiovascular disease, chronic liver disease, chronic lung disease, chronic kidney disease, neurological disease, immunosuppression, and malignancy)	treatment, use of medications that were contraindicated for use with Nirmatrelvir/Ritonavir, estimated glomerular filtration rate $< 30$ mL/minute/1.73m <sup>2</sup> , dialysis, weight $< 40$ kg, pregnancy			
[14]	Retrospective cohort study	Patients and workers at long-term care facilities		2,241	South Korea	February 6, 2022 until April 2, 2022
[15]	Retrospective study	18+, outpatient encounter associated with the COVID-19 diagnosis, at least one previous face-to-face encounter in Cosmos during the 3 years preceding the diagnosis	Pregnancy, pharmacologic or medical contraindications to nirmatrelvir/ritonavir usage	699,848	USA	April until August 2022
[16]	Retrospective study	Severely and critically ill hospitalized patients	30 mL/min $\leq$ creatinine clearance $< 60$ mL/min, severe liver dysfunction (Child-Pugh C), known or suspected pregnancy, hypersensitivity to an nirmatrelvir/ritonavir, HIV infection, receiving Salmeterol, Rifampicin,	1,082	China	December 8, 2022 until February 9, 2023

Publication	Type of Study	Inclusion Criteria	Exclusion Criteria	No. of participants	Geographic region	Study period
			Tacrolimus, Sirolimus, Domperidone, Simvastatin, Rivaroxaban, Estazolam, Atorvastatin, Amiodarone, Propafenone, or Carbamazepine as usual medications prior to this hospitalization or any administration of drugs mentioned above within 72 h prior to assessment of eligibility			
[17]	Retrospective cohort study	Hospitalized, non-oxygen-dependent patients, hospital admission date within 3 days before or after COVID-19 diagnosis	< 18, admission to hospital more than 5 days after symptom onset, history of oral antiviral use before admission, requirement of supplemental oxygen on admission, drug-related contraindications to nirmatrelvir/ritonavir use, severe renal or severe liver impairment	1,780	Hong Kong	Diagnosis between Feb 26, 2022 and Apr 26, 2022

This supplementary material contains three tables: The first table contains the preprints and papers, which met the inclusion criteria and were considered in our review. The second table contains the preprints and papers, which were included in our study, but not the review itself and the final table includes all papers and preprints, whose title and abstracts were scanned, but which did not meet the inclusion criteria.

Used

**Table S2.** Publications which were considered in our review.

<b>Author(s)</b>	<b>Title</b>	<b>Type of publication &amp; journal</b>
[1]	Paxlovid for Treating COVID-19 Patients: A Case- Control Study From Two Hospitals in the Eastern Province of Saudi Arabia	Published in <i>Cureus</i>
[2]	Nirmatrelvir Use and Severe Covid-19 Outcomes during the Omicron Surge	Published in <i>The New England Journal of Medicine</i>
[3]	Effect of Nirmatrelvir/Ritonavir (Paxlovid) on Hospitalization among Adults with COVID-19: an EHR-based Target Trial Emulation from N3C	Preprint
[4]	Paxlovid for hospitalized COVID-19 patients with chronic kidney disease	Published in <i>Antiviral Research</i>
[5]	Molnupiravir, Nirmatrelvir/Ritonavir, or Sotrovimab for High-Risk COVID-19 Patients Infected by the Omicron Variant: Hospitalization, Mortality, and Time until Negative Swab Test in Real Life	Published in <i>pharmaceuticals</i>
[6]	Nirmatrelvir Plus Ritonavir for Early COVID-19 in a Large U.S. Health System	Published in <i>Annals of Internal Medicine</i>
[7]	Oral Nirmatrelvir and Ritonavir for Coronavirus Disease 2019 in Vaccinated, Nonhospitalized Adults Aged 18–50 Years	Published in <i>Clinical Infectious Diseases</i>
[8]	Oral Nirmatrelvir and Ritonavir in Non-hospitalized Vaccinated Patients with Covid-19	Published in <i>Clinical Infectious Diseases</i>

[9]	Oral Nirmatrelvir for High-Risk, Nonhospitalized Adults with Covid-19	Published in <i>New England Journal of Medicine</i>
[10]	Paxlovid (nirmatrelvir/ritonavir) effectiveness against hospitalization and death in N3C: A target trial emulation study	Preprint
[11]	Effectiveness of Paxlovid, an Oral Antiviral Drug, Against the Omicron BA.5 Variant in Korea: Severe Progression and Death Between July and November 2022	Published in <i>Journal of Korean Medical Science</i>
[12]	Efficacy and safety of Paxlovid in severe adult patients with SARS-Cov-2 infection: a multicenter randomized controlled study	Published in <i>The Lancet Regional Health - Western Pacific</i>
[13]	Effectiveness of Nirmatrelvir/Ritonavir in Reducing Severe Coronavirus Disease 2019 and Mortality in High-Risk Patients	Published in <i>Clinical Infectious Diseases</i>
[14]	The effectiveness of Paxlovid treatment in long-term care facilities in South Korea during the outbreak of the Omicron variant of SARS-CoV-2	Published in <i>Osong Public Health and Research Perspectives</i>
[15]	Paxlovid associated with decreased hospitalization rate among adults with COVID-19 – United States, April–September 2022	Published in <i>American Journal of Transplantation</i>
[16]	Composite Interventions on Outcomes of Severely and Critically Ill Patients with COVID-19 in Shanghai, China	Published in <i>Microorganisms</i>
[17]	Real-world effectiveness of early molnupiravir and nirmatrelvir/ritonavir among hospitalized, non-oxygen-	Preprint

	dependent COVID-19 patients on admission during Hong Kong's Omicron BA.2 wave: an observational study	
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**Used, but not in review**

**Table S3.** Publications which were considered in our paper, but not in the review itself.

Author(s)	Summary	Used in section
[18]	Examination (clinically, virologically and immunologically) of seven patients who experienced rebound symptoms. Six experienced rebound symptoms after taking Nirmatrelvir/Ritonavir, the final patient did not receive the treatment.	Discussion
[19]	Short sections on who is eligible for treatment with Nirmatrelvir/Ritonavir, how does Nirmatrelvir/Ritonavir work, what peer reviewed evidence is there for Nirmatrelvir/Ritonavir, which countries are using Nirmatrelvir/Ritonavir, how much does Nirmatrelvir/Ritonavir cost, what is the potential for this drug.	Nirmatrelvir/Ritonavir
[20]	Argue that Nirmatrelvir/Ritonavir might harm transplant patients as there is a chance of medication interaction and that the ritonavir component of Nirmatrelvir/Ritonavir might pose a safety risk for these patients	Discussion
[21]	Enrollment of around 260 patients of which	Methods

Author(s)	Summary	Used in section
	approximately half were treated with molnupiravir, while the other half was treated with Nirmatrelvir/Ritonavir. Recorded hospitalization and deaths during a 14-day follow-up period. As no (untreated) control group is provided, this work is not included in the review.	
[22]	Discussion of if and how rebound symptoms after taking Nirmatrelvir/Ritonavir differ between Omicron BA.5 and Omicron BA.2.12.1 infected patients	Discussion
[23]	Review which summarized the effects of ritonavir on drug disposition and debates what influences the probability of drug-drug interaction when taking Nirmatrelvir/Ritonavir.	Discussion
[24]	Discusses side-effects of Nirmatrelvir/Ritonavir in a single kidney transplant patient, whos treatment with Nirmatrelvir/Ritonavir had to be interrupted and who suffered acute kidney injury from taking the drug	Discussion
[25]	No abstract available on Pubmed, published under "Medical News & Perspectives", discussing rebound effects potentially associated with Nirmatrelvir/Ritonavir treatment.	Discussion
[26]	Assessment of the in vitro antiviral effect of GS-441524, remdesivir, EIDD-1931,	Discussion

Author(s)	Summary	Used in section
	molnupiravir and nirmatrelvir against the various SARS-CoV-2 VOCs, including Omicron.	
[27]	Review of efficacy and safety of the three antiviral treatments molnupiravir, fluvoxamine and Nirmatrelvir/Ritonavir. Published, when only Nirmatrelvir/Ritonavir's clinical results were available.	Introduction

**Unused Table**

**Table S4.** Publications, which were taken into consideration, but which did not meet our inclusion criteria and were hence not included in the review.

Author(s)	Summary	Reason for Exclusion
[28]	Discussing Thymoquinone and other natural products as possible treatments for COVID-19.	Does not meet the inclusion criteria.
[29]	Case discussion of two patients who were prescribed Nirmatrelvir/Ritonavir and experiences a rebound of COVID-19	Does not meet the inclusion criteria.
[30]	Nirmatrelvir/Ritonavir binds non-covalently at regions other than the catalytic sites with energies stronger than reported, namely to mMpro.	Does not meet the inclusion criteria.
[31]	Review which presents the efficacy of repurposed drugs for COVID-19. Discussion of clinical trials, combination therapies and novel methods followed for treatment. Written when Nirmatrelvir/Ritonavir was undergoing Phase III studies.	Does not meet the inclusion criteria.
[32]	Debating evidence from pivotal	Does not meet the inclusion

Author(s)	Summary	Reason for Exclusion
	trials that led to the approval of effective COVID-19 therapeutics and categorizing them as effective outpatient and inpatient management strategies.	criteria.
[33]	Description of drugs that contraindicated and/or should (not) be used carefully when also taking Nirmatrelvir/Ritonavir based on fact sheets by the Spanish Agency for Medicines and Health Products.	Does not meet the inclusion criteria.
[34]	Proposition of a hybrid multiscale mathematical approach to assess Nirmatrelvir/Ritonavir.	Does not meet the inclusion criteria.
[35]	Usage of techniques of structure modeling, in silico docking and pharmacokinetics prediction to test compounds from algae for their ability to inhibit SARS-CoV-2's protease Mpro.	Does not meet the inclusion criteria.
[36]	No abstract available on pubmed.	Abstract unavailable.
[37]	Discussion of the case of a male veteran reporting rebound symptoms after Nirmatrelvir/Ritonavir treatment. The patient was found to be hypoxic with pulmonary emboli.	Does not meet the inclusion criteria.
[38]	No abstract available on pubmed.	Abstract unavailable.
[39]	No abstract available on pubmed.	Abstract unavailable.
[40]	No abstract available on pubmed.	Abstract unavailable.

Author(s)	Summary	Reason for Exclusion
[41]	The English abstract solely states that Nirmatrelvir/Ritonavir is an available treatment for COVID-19 infections, administered orally and to people at high risk for severe disease.	Only the abstract was available in English.
[42]	No abstract available on pubmed.	Abstract unavailable.
[43]	Written when Nirmatrelvir/Ritonavir was still under development, embryofetal development studies in rats and rabbits to assess clinically relevant risks when prescribing Nirmatrelvir/Ritonavir to males and females in reproductive age.	Does not meet the inclusion criteria.
[44]	Aim to critically assess benefits and shortcomings of using molecular models for drug repurposing when trying to develop effective COVID-19 treatments..	Does not meet the inclusion criteria.
[45]	No abstract available on pubmed.	Abstract unavailable.
[46]	Discussing panax ginseng's (a medical plant) ability to control cytokine storm in COVID-19	Does not meet the inclusion criteria.
[47]	Identification of patients potentially eligible for treatment with Nirmatrelvir/Ritonavir in the UK. In consequence, assessment of the coverage of new treatments among these patients with the conclusion that there were variants in coverage between key clinical,	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	geographic and demographic groups.	
[48]	No abstract available on pubmed.	Abstract unavailable.
[49]	No abstract available on pubmed.	Abstract unavailable.
[50]	For 36 mRNA-vaccinated and Omicron-infected individuals viral kinetics were measured. 11 of the 36 were treated with Nirmatrelvir/Ritonavir and treatment was associated with larger incidence of viral rebound.	Does not meet the inclusion criteria.
[51]	Report of simulation of H172Y mutation on Mpro's structure leading to decreased structural stability and binding affinity.	Does not meet the inclusion criteria.
[52]	Evaluation of the incidence of viral and symptom rebound in untreated mild to moderate COVID-19 outpatients.	Does not meet the inclusion criteria.
[53]	Review presenting the progress in clinical trials concerning the effectiveness of treatments for COVID-19. Written when solely the results from Nirmatrelvir/Ritonavir's clinical trials were available.	Does not meet the inclusion criteria.
[54]	No abstract available on pubmed.	Abstract unavailable.
[55]	Presenting the preclinical disposition, metabolism and potential drug-drug interaction of nirmatrelvir.	Does not meet the inclusion criteria.
[56]	Discussion of levels of total cholesterol, LDL-C, HDL-C, and apolipoprotein B and A-I levels in patients with COVID-19 infections.	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
[57]	Discussing the potential of Yindan Jiedu granules as a treatment for COVID-19. Comparison of Yindan Jiedu granules and Nirmatrelvir/Ritonavir when treating COVID-19 patients.	Does not meet the inclusion criteria.
[58]	Present how Pfizer has successfully improved its research and development (R&D) productivity between 2010 and 2020.	Does not meet the inclusion criteria.
[59]	Presentation of literature on monotherapy use of Nirmatrelvir/Ritonavir and monotherapy use of remdesivir and discussion of the hypothesis of using nirmatrelvir and remdesivir to increase efficacy of COVID-19 treatment.	Does not meet the inclusion criteria.
[60]	No abstract available on pubmed.	Abstract unavailable.
[61]	Provides a review of currently available COVID-19 treatments. As this was published in December 2021, it only include the clinical trials for Nirmatrelvir/Ritonavir.	Does not meet the inclusion criteria.
[62]	Data from December 2021 until May 2022 were analyzed to describe oral antiviral treatment prescription dispensing overall and by week, stratified by zip code and social vulnerability.	Does not meet the inclusion criteria.
[63]	Evaluation of in vitro potency of normatrelvir against the Mpro of currently circulating (and previous) variants of	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	concern. Their in vitro data suggests that Nirmatrelvir/Ritonavir has the ability to inhibit SARS-CoV-2 replication, even vor VOCs like Omicron.	
[64]	No abstract available on pubmed.	Abstract unavailable.
[65]	Retrospective analysis of benefits and limitations of (previous) treatments for COVID-19. Calls Nirmatrelvir/Ritonavir a promising treatment, but only presents clinical results.	Does not meet the inclusion criteria.
[66]	People with cystic fibrosis (CF) are at risk for drug-drug interaction when being treated with Nirmatrelvir/Ritonavir. Simulation of coadministration of elexacaftor-tezacaftor-ivacaftor and Nirmatrelvir/Ritonavir to examine these potential interactions.	Does not meet the inclusion criteria.
[67]	Case report of a severe aplastic anemia child who was successfully treated with Nirmatrelvir/Ritonavir.	Does not meet the inclusion criteria.
[68]	Written before Nirmatrelvir/Ritonavir was authorized and before clinical results were available. Hence, discussing the promising antiviral effect of nirmateelvir, while also naming (then) unresolved concerns.	Does not meet the inclusion criteria.
[69]	Evaluation of drugs authorized in the US for treating COVID-19. For Nirmatrelvir/Ritonavir's	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	effectiveness, solely the clinical results are discussed.	
[70]	Discussing medical management (but excluding intensive care management) of COVID-19.	Article only available in French.
[71]	Review to present the potential of nirmatrelvir when treating COVID-19, discussing its history of rational design, its target selectivity, synthesis and drug resistance.	Does not meet the inclusion criteria.
[72]	News article introducing the most important Nirmatrelvir/Ritonavir-related information for the general public and patients.	Does not meet the inclusion criteria.
[73]	No abstract available on pubmed.	Abstract unavailable.
[74]	No abstract available on pubmed.	Abstract unavailable.
[75]	Examination of rates and relative risks of COVID-19 rebound in patients treated with Molnupiravir or Nirmatrelvir/Ritonavir during January-June 2022	Does not meet the inclusion criteria.
[76]	Summary of the milestones during the development of normatrelvir plus ritonavir leading to its first authorizations and approval for the treatment of COVID-19	
[77]	No abstract available on pubmed.	Abstract unavailable.
[78]	No abstract available on pubmed.	Abstract unavailable.
[79]	Discussion of potential nirmatrelvir escape mutations from emerging variants of	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	SARS-CoV-2 and exploration of the mutational landscape of Mpro.	
[80]	Discussion of drug interactions of Nirmatrelvir/Ritonavir and immunosuppressant drugs and provision of general recommendations for therapeutic drug monitoring when co-administering the two.	Does not meet the inclusion criteria.
[81]	Providing recommendations on behalf of the national French society of pharmacology for possible drug-drug interactions between Nirmatrelvir/Ritonavir and other commonly used drugs.	Does not meet the inclusion criteria.
[82]	No abstract available on pubmed.	Abstract unavailable.
[83]	Introduction of a potential future treatment, the orally available ribonucleoside analog 4'-fluorouridine (4'-FIU). Furthermore, reviewing currently approved and emerging medicines against COVID-19.	Does not meet the inclusion criteria.
[84]	Discussing statin use in the context of COVID-19, coming to no evidence suggesting interference between statins and COVID-19 vaccines. But, simultaneous statins and Nirmatrelvir/Ritonavir administration may increase statin exposure and the risk of adverse effects.	Does not meet the inclusion criteria.
[85]	Development of an efficient LC-MS/MS method for simultaneously determining	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	nirmatrelvir and ritonavir in human plasma.	
[86]	Show that Apilimod and other PIKfyve inhibitors worsen disease in a COVID-19 murine model when given prophylactically or therapeutically. Abstract solely mentions Nirmatrelvir/Ritonavir as an available treatment.	Does not meet the inclusion criteria.
[87]	Multicentre cohort study describing clinical characteristics, and assessing risk and protective factors for geriatric Omicron severe infections.	
[88]	Discussing herbal extracts as potential treatments for COVID-19. Solely mentions that molnupiravir and Nirmatrelvir/Ritonavir are not widely available.	Does not meet the inclusion criteria.
[89]	News article in BMJ, published shortly after the analysis of the phase II-III data was published. Hence, discussing and reporting the results of this analysis.	Does not meet the inclusion criteria.
[90]	Consideration of electronic health record from a large integrated health care system in California to analyze and quantify hospital admissions and emergency department encounters related to COVID-19 during the 5-15 days after receiving Nirmatrelvir/Ritonavir	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	treatment	
[91]	No abstract available, article in French.	
[92]	Report the discovery and description of nirmatrelvir. Comparison of effectiveness of molnupiravir and normatrelvir). Written while Nirmatrelvir/Ritonavir was under study in phase III of the clinical trial.	Does not meet the inclusion criteria.
[93]	Praxis-relevant recommendations for the usage and prescription of Paxlovid.	Does not meet the inclusion criteria.
[94]	No abstract available on pubmed.	
[95]	News article published in EMBO Molecular Medicine, discussing [96]	Does not meet the inclusion criteria.
[97]	Compilation of a list of drugs and their potentially relevant interactions when being administered simultaneously with Nirmatrelvir/Ritonavir as well as a list of commonly prescribed drugs for which no such interactions exist.	Does not meet the inclusion criteria.
[98]	Discussion of how ongoing virus evolution has the potential to yield variants with resistance to clinical protease inhibitors (like Nirmatrelvir/Ritonavir).	Does not meet the inclusion criteria.
[99]	Published when solely Nirmatrelvir/Ritonavir's clinical results were available. Hence, only these are discussed.	Does not meet the inclusion criteria.
[100]	Review of normatrelvir's binding to SARS-CoV-2's Mpro	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	and its potential inefficacy when novel mutations arise.	
[101]	Presentation of current antiviral drugs for COVID-19 as well as potential future directions.	Does not meet the inclusion criteria.
[102]	Review of literature on the drug design landscape of SARS-CoV-2 Mpro inhibitors. Mentions nirmatrelvir as one such inhibitor.	Does not meet the inclusion criteria.
[103]	Development of a non-pathogenic system in which yeast growth is a proxy for Mpro activity. Consequently, mutants which exhibit drug sensitivity and altered enzymatic activity can quickly be identified.	Does not meet the inclusion criteria
[104]	Discussion of review of risk factors for severe covid and Pre-exposure prophylaxis (PrEP) strategies against SARS-CoV-2. Additionally, presentation of potential SARS-CoV-2 PrEP drugs	Does not meet the inclusion criteria.
[105]	Published in late 2021, reporting the current status of oral treatments for COVID-19	Does not meet the inclusion criteria.
[106]	Analysis of newly reported bat coronaviruses with regard to the similarities and differences between their 3CL protease and SARS-CoV-2.	Does not meet the inclusion criteria
[107]	Analysis of the structural features of the Spike protein and the Mpro of the SARS-CoV-2 variant XE and the closely related variants XD and XF.	Does not meet the inclusion criteria.
[108]	Comparison and changes in the epidemiology, clinical profile,	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	therapeutics and public health measures for the COVID-19 pandemic in the Asia Pacific region. Abstract mentions recent introduction of Nirmatrelvir/Ritonavir.	
[109]	Report of 4 cases from a post-COVID cohort study who received nirmatrelvir as part of clinical care and who experiences different outcomes.	Does not meet the inclusion criteria.
[110]	Published in late 2021, when the federal government rationed Nirmatrelvir/Ritonavir doses. Hence, this paper identifies relevant ethical principles and priority groups for access to Nirmatrelvir/Ritonavir.	Does not meet the inclusion criteria.
[111]	No abstract available on pubmed.	
[112]	No abstract available on pubmed.	
[113]	Examine modifications of a promising peptide-based inhibitor of the spike protein, LCB3, against common mutations in the target protein. In consequence, LCB3 retains its efficacy against the spike protein.	Does not meet the inclusion criteria.
[114]	Discussing Nirmatrelvir/Ritonavir's efficacy after publication of clinical results. Article is in Spanish.	Does not meet the inclusion criteria.
[115]	Discussion of a new class of sartan-like arterial antihypertensive drugs (referred to as "bisartans") as treatment for COVID-19.	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	Abstract notes that bisartans do not inhibit SARS-CoV-2 infection in bioassays as effectively as Nirmatrelvir/Ritonavir.	
[116]	No abstract available on pubmed.	
[117]	Discussion of risk of drug interactions between treatments for COVID-19 and drugs used in treating comorbid conditions like diabetes or cardiovascular illness.	Does not meet the inclusion criteria.
[118]	Summarizes diagnostic and therapeutic management of COVID-19 for outpatients. Abstract names Nirmatrelvir/Ritonavir was the preferred treatment for mild cases with high risk of disease progression.	Does not meet the inclusion criteria.
[119]	Assessing currently available COVID-19 drugs for their potential toxicological effects and mechanisms.	Does not meet the inclusion criteria.
[120]	Describes clinical experience with 25 organ transplant recipients who were prescribed Nirmatrelvir/Ritonavir. Their results suggest that clinically significant interaction between Nirmatrelvir/Ritonavir and immunosuppressive agents can be reasonably managed with a standardized dosing protocol	Does not meet the inclusion criteria.
[121]	No abstract available on pubmed.	
[122]	Review of molnupiravir and Nirmatrelvir/Ritonavir with	Does not meet the inclusion criteria

Author(s)	Summary	Reason for Exclusion
	regard to their mechanisms of action, their antiviral activity, pharmacokinetics, drug interactions and clinical experience including trials, adverse events, recommended indications and formulary considerations.	
[123]	Providing vivo safety assessments of Nirmatrelvir/Ritonavir by studies in rats and monkeys.	Does not meet the inclusion criteria.
[124]	Simulation of molnupiravir treatment to judge effectiveness of antiviral therapy in highly-transmissible variants.	Does not meet the inclusion criteria.
[125]	No abstract available on pubmed.	
[126]	Discussing the intrinsic stability of Nirmatrelvir and the degrading products formed under forced conditions.	Does not meet the inclusion criteria.
[127]	No abstract available on pubmed.	
[128]	Introduce the ability of the combination of pamapimod and pioglitazone to inhibit SARS-CoV-2 replication in vitro. Hence, this combination is a potential treatment of COVID-19 and is evaluated (at the time of writing) in a phase II clinical study.	Does not meet the inclusion criteria.
[129]	Provides practical clinical guidelines for using molnupiravir in COVID-19 patients.	Does not meet the inclusion criteria.
[130]	Discussing contraceptive care in times of the Pandemic. Abstract mentions that	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	combined hormonal contraceptive users who take Nirmatrelvir/Ritonavir should consider an additional contraceptive method for the duration of Nirmatrelvir/Ritonavir treatment.	
[131]	No abstract available on pubmed.	
[132]	Development and validation of a method to quantify almonertinib in rat plasma to study the effects of Nirmatrelvir/Ritonavir on the pharmacokinetics of almonertinib in rats.	Does not meet the inclusion criteria.
[133]	No abstract available on pubmed.	
[134]	Presenting the role of endothelial and vascular components as major targets for COVID-19-induced tissue injury, spreading to various organs and injury healing as well as current COVID-19 treatments.	Does not meet the inclusion criteria.
[135]	Israeli study to determine the number of patients who could be included in a prospective Real World Evidence Study to study Nirmatrelvir/Ritonavir's effect on patients' outcomes as well as assessment of comparability between patients who received the treatment and patients who did not.	
[136]	Discussing the important role of the signal transducer and	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	activator of transcription (STAT) 1 protein in antiviral immune response and how viruses like Ebola and SARS-CoV-2 have developed the ability to inhibit this transcription factor.	
[137]	No abstract available on pubmed.	
[138]	No abstract available on pubmed.	
[139]	No abstract available on pubmed.	
[140]	Review of the indications of immunosuppressants and immunomodulators and guidance on which mild to moderate COVID-19 patients might benefit from their use in dermatology.	Does not meet the inclusion criteria.
[141]	Review which provides an overview regarding the molecular profile of the Omicron variant, as well as its transmissibility and the remaining vaccine effectiveness. Mentioned molnupiravir and Nirmatrelvir/Ritonavir as oral treatments of COVID-19.	Does not meet the inclusion criteria.
[142]	Written when the Omicron variant had just emerged and molnupiravir and Nirmatrelvir/Ritonavir had only undergone clinical study. Briefly reviewing the impact of these two new oral antivirals.	Does not meet the inclusion criteria.
[143]	Discussion of drug interactions when simultaneously taking	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	Nirmatrelvir/Ritonavir and cholesterol-lowering drugs.	
[144]	Summary of the current progress in the structural biology of SARS-CoV-2 as well as presentation of structure-based design of Nirmatrelvir/Ritonavir, molnupiravir and VV116 to emphasize the importance of structure in drug development for COVID-19	Does not meet the inclusion criteria.
[145]	No abstract available on pubmed.	
[146]	Recruitment of 5 pediatric cases with underlying disease who were treated with Nirmatrelvir/Ritonavir as well as 30 age-matched patients with underlying disease who were not treated with Nirmatrelvir/Ritonavir as controls. Assessment of efficacy and safety of Nirmatrelvir/Ritonavir as well as inter-group comparisons.	Does not meet the inclusion criteria.
[147]	From evolutionary and structural standpoints possible mutations in Mpro leading to evasion of nirmatrelvir are discussed.	Does not meet the inclusion criteria.
[148]	Case report of a 14 year old female kidney transplant, whose COVID-19 infection was treated with Nirmatrelvir/Ritonavir.	Does not meet the inclusion criteria.
[149]	Cystic fibrosis patients are at increased risk for drug-drug interaction when taking Nirmatrelvir/Ritonavir. Here,	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	these interactions are explored using a physiologically- based pharmacokinetic modeling approach.	
[150]	Letter to the editor discussing the potential of Nirmatrelvir/Ritonavir to treat COVID-19	Does not meet the inclusion criteria.
[151]	Review presenting an update on fluorinated COVID-19 drugs. Here, current knowledge of these drugs' molecular design, metabolism and pharmacokinetics as well as mechanism of action, is provided.	Does not meet the inclusion criteria.
[152]	Review which aims to provide a compact and updated summary of pediatric COVID-19 diagnosis and management. Abstract mentions that in patients at increased risk of progression, Nirmatrelvir/Ritonavir (among other treatments) should be considered.	Does not meet the inclusion criteria.
[153]	No abstract available on pubmed.	Abstract unavailable.
[154]	Study of some of the key human transporters interacting with Molnupiravir and Nirmatrelvir.	Does not meet the inclusion criteria.
[155]	No abstract available on pubmed.	Abstract unavailable.
[156]	Exploration of several reactive warheads in the design of Mpro inhibitors, as SARS-CoV-2 main protease (Mpro) is a validated antiviral drug target of nirmatrelvir.	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
[157]	Presentation of the case of a small bowel transplant patient who had a COVID-19 infection by discussing the management experience and reviewing the relevant literature.	Does not meet the inclusion criteria.
[158]	Listing of etiopathogenic theories of long coronavirus disease and discussion of currently existing main lines of treatment.	Does not meet the inclusion criteria.
[159]	Discussion of possible means to prevent Long COVID, including vaccination, orally effective antiviral agents, and presentation of ongoing clinical trials on dietary supplements or drugs with different mechanisms of action.	Does not meet the inclusion criteria.
[160]	Exploration of the cause for prolonged viral shedding time to establish increased age, comorbidities, incomplete vaccination, severe or critical infections, and delayed Paxlovid treatment as the risk factors.	Does not meet the inclusion criteria.
[161]	Suggestion of structural modification of nirmatrelvir to fight the drug resistance resulting from the appearance of many novel COVID-19 variants.	Does not meet the inclusion criteria.
[162]	No abstract available on pubmed.	Abstract unavailable.
[163]	Report a case of simultaneous mixed-type autoimmune hemolytic anemia with immune thrombocytopenia which followed after a severe infection	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	with COVID-19, treatment with Paxlovid and a subsequent Rhinovirus infection.	
[164]	Discuss the speed of the development of Nirmatrelvir.	Does not meet the inclusion criteria.
[165]	Application of a parsimonious mathematical COVID-19 immunity model to explore the effect of nirmatrelvir/ritonavir in (un)vaccinated patients and potentially occurring viral rebound.	Does not meet the inclusion criteria.
[166]	Case report of a heart-transplant patient to explore the drug-drug interaction between nirmatrelvir/ritonavir and Tacrolimus.	Does not meet the inclusion criteria.
[167]	Review the development of small molecule anti-SARS-CoV-2 drugs in China.	Does not meet the inclusion criteria.
[168]	Prediction of the length of stay (LoS) for hospitalized COVID-19 patients and analysis of cohorts of risk factors reducing or prolonging LoS.	Does not meet the inclusion criteria.
[169]	No abstract available on pubmed.	Abstract unavailable.
[170]	<b>Explore clinical efficacy of Paxlovid in children with hematological diseases.</b>	Does not meet the inclusion criteria.
[171]	Modeling study exploring the public health impact of bivalent vaccines and nirmatrelvir-ritonavir against COVID-19 and coming to the inclusion and coming to the conclusion that the most efficient strategy for both is to target the group of	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	people 75 and older.	
[172]	Exploration of risk factors for Omicron-associated pneumonia.	Does not meet the inclusion criteria.
[173]	No abstract available on pubmed.	Abstract unavailable.
[174]	Investigation of potential herb-drug interactions between traditional Chinese medicine and COVID-19 treatments.	Does not meet the inclusion criteria.
[175]	Investigation of the outcomes of Covid-19 infection in patients with Hairy cell leukemia.	Does not meet the inclusion criteria.
[176]	Assessment of the cost-effectiveness of Paxlovid in reducing severe COVID-19 and mortality in China to come to the conclusion that Paxlovid's only cost-effective in patients 80 and older (regardless of vaccination status).	Does not meet the inclusion criteria.
[177]	Modeling study to explain COVID-19 rebound after nirmatrelvir/ritonavir treatment.	Does not meet the inclusion criteria.
[178]	Discovery of novel bicyclic[3.3.0]proline peptidyl $\alpha$ -ketoamides as potent 3CL-protease inhibitors for SARS-CoV-2.	Does not meet the inclusion criteria.
[179]	Case study of a patient exhibiting petechial rash found to be seronegative IgA vasculitis presenting concurrently with dyspnea subsequently to being infected with COVID-19.	Does not meet the inclusion criteria.
[180]	Screening of a library of microbial metabolites to	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	discover new SARS-CoV-2 inhibitors to complement and extend existing COVID-19 drugs.	
[181]	Investigation of potential herb–drug interactions in low- and middle-income countries, in which access to Paxlovid has recently increased.	Does not meet the inclusion criteria.
[182]	No abstract available on pubmed.	Abstract unavailable.
[183]	Case study of a patient, who was treated with Paxlovid and continued to take Ranolazine, resulting in a drug-drug interaction and an emergency room visit.	Does not meet the inclusion criteria.
[184]	Introduction of novel Peptidomimetics for the treatment of COVID-19.	Does not meet the inclusion criteria.
[185]	Investigate the immunity acquired through BA.2 and BA.2.12.1 infections and come to the conclusion that these infections fail to protect against the BA.4/BA.5 variant.	Does not meet the inclusion criteria.
[186]	Present the case of a patient who suffered from acute pancreatitis after taking Paxlovid.	Does not meet the inclusion criteria.
[187]	Present the case of an elderly patient who, after taking Paxlovid, experienced medication interactions with benzodiazepines and narcotics leading to altered mental status.	Does not meet the inclusion criteria.
[188]	Present four cases who suffered from a rebound after taking Paxlovid and progressed to severe COVID-19.	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
[189]	Literature review to examine if nirmatrelvir/ritonavir can be used for solid organ transplant recipients suffering from COVID-19.	Does not meet the inclusion criteria.
[190]	Perform a national survey on the nirmatrelvir/ritonavir prescribing processes and access barriers.	Does not meet the inclusion criteria.
[191]	Explore possible adverse cutaneous reactions resulting from nirmatrelvir/ritonavir intake.	Does not meet the inclusion criteria.
[192]	No abstract available on pubmed.	Abstract unavailable.
[193]	No abstract available on pubmed.	Abstract unavailable.
[194]	Present VV116, a chemically-modified version of the antiviral remdesivir, as a potential treatment for COVID-19.	Does not meet the inclusion criteria.
[195]	No abstract available on pubmed.	Abstract unavailable.
[196]	Discuss the potential of Paxlovid for treating mild COVID-19 cases of pregnant and/or lactating patients.	Does not meet the inclusion criteria.
[197]	Usage of deep learning to predict potential drug-drug interactions between Paxlovid and more than 2,000 prescription drugs.	Does not meet the inclusion criteria.
[198]	Explore resistance profiles of nirmatrelvir, ensitrelvir, and FB2001 to find that the next-generation drugs have the potential to be effective against nirmatrelvir-resistant variants.	Does not meet the inclusion criteria.
[199]	Present experience gained from	Does not meet the inclusion

Author(s)	Summary	Reason for Exclusion
	the COVID-19 pandemic and strategies for facing present and future pandemics.	criteria.
[200]	Review to present in-depth information on new molecules and repurposed drugs for the treatment of COVID-19.	Does not meet the inclusion criteria.
[201]	Discuss how administration of Paxlovid for the treatment of COVID-19 is feasible for solid organ transplant recipients with medical management.	Does not meet the inclusion criteria.
[202]	Case study of a patient who suffered from tacrolimus toxicity as a result of drug-drug interaction between Paxlovid and their home medications.	Does not meet the inclusion criteria.
[203]	Exploration of effectiveness of Favipiravir monotherapy in the treatment of COVID-19.	Does not meet the inclusion criteria.
[204]	Usage of molecular docking analysis to discover Nirmatrelvir's antiviral potency against different coronaviruses.	Does not meet the inclusion criteria.
[205]	Explore vaccination and treatment options for COVID-19 for lactating patients.	Does not meet the inclusion criteria.
[206]	No abstract available on pubmed.	Abstract unavailable.
[207]	Explore nirmatrelvir/ritonavir-associated reports of dysgeusia.	Does not meet the inclusion criteria.
[208]	Discuss potential use and dosing of nirmatrelvir/ritonavir for patients with chronic kidney disease.	Does not meet the inclusion criteria.
[209]	Explore factors associated with prolonged COVID-19 viral shedding in elderly Chinese patients.	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
[210]	Case report of a patient who tested positive again for the Omicron BA.5 variant after being treated with nirmatrelvir/ritonavir.	Does not meet the inclusion criteria.
[211]	No abstract available on pubmed.	Abstract unavailable.
[212]	No abstract available on pubmed.	Abstract unavailable.
[121]	No abstract available on pubmed.	Abstract unavailable.
[213]	Demonstration of a 7-step, 3-pot synthesis of the antiviral nirmatrelvir, arriving at the targeted drug in 70% overall yield.	Does not meet the inclusion criteria.
[214]	Summary of the currently available data on ritonavir and nirmatrelvir during pregnancy as well as review of recommendations of major societies worldwide regarding Paxlovid use during pregnancy.	Does not meet the inclusion criteria.
[215]	Analysis of Google search queries to explore public interest in COVID-19 antivirals. Does not meet inclusion criteria.	Does not meet the inclusion criteria.
[216]	Design of mutations to impair binding of nirmatrelvir and demonstrate nirmatrelvir resistance in the COVID-19 BA.1 variant and WA1 replicons.	Does not meet the inclusion criteria.
[217]	Report the case of a patient who developed symptomatic bradycardia, syncopal episodes, and sinus pause	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	subsequent to taking nirmatrelvir/ritonavir.	
[218]	Report consensus on diagnosis and treatment of COVID-19 related cough and present nirmatrelvir/ritonavir as one treatment option.	Does not meet the inclusion criteria.
[219]	No abstract available on pubmed.	Abstract unavailable.
[220]	No abstract available on pubmed.	Abstract unavailable.
[221]	Explore the frequency of COVID-19's resistance to Nirmatrelvir/ritonavir and virological/clinical rebounds and underline the importance of these findings for immunocompromised patients.	Does not meet the inclusion criteria.
[222]	Discussion of the potential of theranostics regarding COVID-19 antivirals.	Does not meet the inclusion criteria.
[223]	Case report of a patient, who is being treated with clozapine and who, after receiving Paxlovid, developed neutropenia.	Does not meet the inclusion criteria.
[224]	Commentary on the differences, therapeutic potential, and best-practices of Paxlovid and Lagevrio to inform and support community pharmacy practices.	Does not meet the inclusion criteria.
[225]	Discovery of a uniquely Y-shaped, S-O-N-O-S-bridged post-translational cross-link that connects three residues C22, C44, and K61 at their side chains while analyzing M <sup>Pro</sup> crystals.	Does not meet the inclusion criteria.
[226]	Case report of a patient who	Does not meet the inclusion

Author(s)	Summary	Reason for Exclusion
	experienced symptomatic bradycardia, likely as a result of a drug-drug interaction of ritonavir-boosted nirmatrelvir and verapamil.	criteria.
[227]	Essay on how to ease the process of obtaining Paxlovid for the treatment of COVID-19.	Does not meet the inclusion criteria.
[228]	Review of the treatment possibilities for COVID-19 in pediatric care. Come to the conclusion that molnupiravir cannot be used, while the other currently available antivirals may be administered at least to older children.	Does not meet the inclusion criteria.
[229]	No abstract available on pubmed.	Abstract unavailable.
[230]	No abstract available on pubmed.	Abstract unavailable.
[231]	Exploration of pediatric Nirmatrelvir/Ritonavir Prescribing Patterns, coming to the conclusion that Nirmatrelvir/Ritonavir is prescribed infrequently to children.	Does not meet the inclusion criteria.
[232]	No abstract available on pubmed.	Abstract unavailable.
[233]	No abstract available on pubmed.	Abstract unavailable.
[234]	No abstract available on pubmed.	Abstract unavailable.
[235]	Report a devised route to nirmatrelvir that reduces the overall sequence to a 1-pot process, relying on the green coupling reagent T3P.	Does not meet the inclusion criteria.
[236]	Presentation of some examples of new peptides and	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	peptidomimetic protease inhibitors in the field of SARS-CoV-2.	
[237]	Present the SARS-COV-2-within-host evolution of 15 patients during antiviral treatment, comparing their in vivo variability to drug-naive patients.	Does not meet the inclusion criteria.
[238]	Considers available literature and product information to assess potential drug-drug interactions between asthma medication and antiviral treatments for COVID-19.	Does not meet the inclusion criteria.
[239]	Case report of a patient who suffered from drug-drug interaction between tacrolimus and nirmatrelvir/ritonavir resulting in toxic levels of tacrolimus.	Does not meet the inclusion criteria.
[240]	Present examples of drugs co-administered with pharmacoenhancers (e.g. nirmatrelvir/ritonavir).	Does not meet the inclusion criteria.
[241]	Retrospective analysis of COVID-19 patients who were eligible for nirmatrelvir/ritonavir treatment during early 2022. Discussion of prescription rate, reasons for non-prescription as well as patient outcomes.	Does not meet the inclusion criteria.
[242]	Review which summarizes the basic features of protease of human immunodeficiency virus (HIV), hepatitis C virus (HCV), and SARS-CoV-2 and discusses the properties of their inhibitors in clinical use.	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
[243]	No abstract available on pubmed.	Abstract unavailable.
[244]	Case report of an elderly patient who was being administered Paxlovid upon admittance to the hospital due to moderate COVID-19.	Does not meet inclusion criteria.
[245]	No abstract available on pubmed.	Abstract unavailable.
[246]	Acknowledge the importance of systematic, orderly and complete collection of health data of citizens and explore how the clinical research methodology process has changed in Italy due to COVID-19.	Does not meet the inclusion criteria.
[247]	Exploration of nirmatrelvir developmental toxicity using zebrafish as in vivo model to contribute to research on administration of nirmatrelvir during pregnancy.	Does not meet the inclusion criteria.
[248]	Explore the optimal docking between M <sup>pro</sup> and antiviral drugs concluding that Paxlovid may work on new strains, including omicron as Omicrons M <sup>pro</sup> mutation has no direct effect on the protein.	Does not meet the inclusion criteria.
[249]	Performance of in vitro and in vivo studies to explore the effect of nirmatrelvir/ritonavir on chondrocyte degeneration and senescence.	Does not meet the inclusion criteria.
[250]	Presentation of difficulties when administering nirmatrelvir/ritonavir to epilepsy patients.	Does not meet the inclusion criteria.
[251]	Assessment of potency of	Does not meet the inclusion

Author(s)	Summary	Reason for Exclusion
	molnupiravir and nirmatrelvir/ritonavir against SARS-Cov-2 in two animal models (Roborovski dwarf hamster model and the ferret model of upper respiratory tract disease and transmission) confirming similar therapeutic benefit of both drugs for older adults.	criteria.
[252]	Development of compartmental model, which describes SARS-CoV-2 spread after the reopening of a French Zero-COVID territory, which displays high vaccination hesitancy, low immunity, and high levels of comorbidities.	Does not meet the inclusion criteria.
[253]	Introduction of a new method to transparently report the living evidence surveillance process that occurs between published living systematic reviews (LSR) versions. Example used is a review on nirmatrelvir/ritonavir.	Does not meet the inclusion criteria.
[254]	No abstract available on pubmed.	Abstract unavailable.
[255]	Case report of a heart transplant recipients to increase awareness of nuances when treating COVID-19 for such patients.	Does not meet the inclusion criteria.
[256]	In silico study that suggests that sage is a rich source of potent anti-coronavirus flavonoids.	Does not meet the inclusion criteria.
[257]	Presentation of various COVID-19 treatments and therapies.	Does not meet the inclusion criteria.
[258]	Usage of <i>in silico</i> mutational scanning and inhibitor docking	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	of Mpro to identify potential resistance mutations.	
[259]	Case report of three children/adolescents who received Paxlovid to treat their COVID-19 infection. Paxlovid was effective and relatively safe in these three cases.	Does not meet the inclusion criteria.
[260]	Suggest that a direct application of 2-5 oligoadenylylates, included in an inhalation spray, may be effective in treatment of severe COVID-19 infections.	Does not meet the inclusion criteria.
[261]	No abstract available on pubmed.	Abstract unavailable.
[98]	Investigation of the susceptibility of natural variants of SARS-CoV-2's main protease to monitor variance resistant to currently available vaccines and drugs.	Does not meet the inclusion criteria.
[262]	Uncover a metabolic pathway that simultaneously modulates pulmonary inflammation, tissue recovery and host metabolic health, which patients with an underlying metabolic disease suffering from COVID-19 are especially prone to.	Does not meet the inclusion criteria.
[263]	Analyze the effect of nirmatrelvir/ritonavir on the length of hospital stay, finding that Paxlovid reduces the viral clearance time and length of hospital stay for hospitalized COVID-19 patients.	Does not meet the inclusion criteria.
[264]	A review which explores the evidence of tested herocyclic	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	compounds acting on different targets against COVID-19 in order to help the development of novel COVID-19 drugs.	
[265]	Discuss and encourage the usage of convalescent plasma when treating immunocompromised COVID-19 patients.	Does not meet the inclusion criteria.
[266]	Review of currently approved drugs and most of the important drug candidates for COVID-19 related targets.	Does not meet inclusion criteria.
[267]	Description of a LC-MS/MS method for the simultaneous quantification of nirmatrelvir and ritonavir in human plasma of COVID-19 patients who are being treated with nirmatrelvir/ritonavir.	Does not meet the inclusion criteria.
[268]	No abstract available on pubmed.	Abstract unavailable.
[269]	Study of the solid-state behavior of the drugs ritonavir and lopinavir in raw materials and in milled compositions indicated that mixtures of ritonavir Forms I and II are found in different raw materials from the same manufacturer.	Does not meet the inclusion criteria.
[270]	Discussion of four patients with end-stage renal disease undergoing hemodialysis, who simultaneously received nirmatrelvir/ritonavir.	Does not meet the inclusion criteria.
[271]	Case report of a cardiac transplant patient who suffered from tracolimus toxicity induced by outpatient treatment for COVID-19 with	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	ritonavir-nirmatrelvir.	
[272]	Explore racial and ethnic disparities for nirmatrelvir/ritonavir prescriptions.	Does not meet the inclusion criteria.
[273]	Report pharmacists' experiences of managing/counseling/following up the usage of nirmatrelvir/ritonavir.	Does not meet the inclusion criteria.
[274]	Analyze nirmatrelvir/ritonavir and molnupiravir dispensing rates by zip code-level social vulnerability and highlight intervention strategies.	Does not meet the inclusion criteria.
[275]	Write an "A guide to" article to explain the development of antiviral drugs against SARS-CoV-2 and their potential in fighting COVID-19.	Does not meet the inclusion criteria.
[276]	Characterization of 14 naturally occurring M polymorphisms that are close to the binding site of these antivirals and testing of potency of nirmatrelvir and ensitrelvir against these polymorphisms.	Does not meet the inclusion criteria.
[277]	Explore the progress of circulating microRNAs in the regulation of severe inflammatory response, immune dysfunction, and thrombosis caused by SARS-CoV-2 infection.	Does not meet the inclusion criteria.
[278]	Presentation of a class of enzyme inhibitors, which could serve as a class of COVID-19 drug targets.	Does not meet the inclusion criteria.
[279]	Description of an in vitro pathway of SARS-CoV-2	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	development towards resistance against nirmatrelvir/ritonavir.	
[280]	Analyze recent mutations of M <sup>Pro</sup> coming to the conclusion that said increased mutational variability may encourage development of antiviral resistance.	Does not meet the inclusion criteria.
[281]	Present the current stage of development of covalent and noncovalent main protease (M <sup>pro</sup> ) inhibitors and the identification of nirmatrelvir-resistant mutants.	Does not meet the inclusion criteria.
[282]	Report of an optically active synthesis of nirmatrelvir that avoids a critical epimerization step.	Does not meet the inclusion criteria.
[283]	Review on COVID-19 vaccines, focusing on their protective efficacies, as well as presentation of an overview of progress in developing potent drugs and monoclonal antibodies against COVID-19.	Does not meet the inclusion criteria.
[284]	Presentation of a COVID-19 oral antiviral assessment protocol developed by an independent community pharmacy and highlighting of the ability of community pharmacists to identify and address medication-related problems.	Does not meet the inclusion criteria.
[285]	Introduction of a nirmatrelvir/ritonavir physiologically-based pharmacokinetic (PBPK) model.	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	Does not meet inclusion criteria.	
[286]	Exploration of a rare natural mutation, H172Y and its potential ability to weaken nirmatrelvir's inhibitory activity.	Does not meet the inclusion criteria.
[287]	Updated version of the preprint above.	Does not meet the inclusion criteria.
[288]	Explore the imitation of the utility of pharmacists prescriptive authority for Paxlovid due the the restrictions included in the introduced FDA framework.	Does not meet the inclusion criteria.
[289]	Suggest the replacement of hydrogens with deuterium in nirmatrelvir to slow oxidation and potentially replace ritonavir, which alters the metabolism of other drugs	Does not meet the inclusion criteria.
[290]	Clinical trial of VV116, a potential alternative to nirmatrelvir/ritonavir for oral treatment of COVID-19.	Does not meet the inclusion criteria.
[291]	No abstract available on pubmed.	Abstract unavailable.
[292]	No abstract available on pubmed.	Abstract unavailable.
[293]	No abstract available on pubmed.	Abstract unavailable.
[294]	Usage of Density Functional Theory to examine electronic and thermodynamic properties of nine COVID-19 drugs, coming to the conclusion that the electrophilic indexes are important to understand their efficiencies.	Does not meet the inclusion criteria.
[295]	Review of current knowledge	Does not meet the inclusion

Author(s)	Summary	Reason for Exclusion
	of pathogenesis and neurological disorders related to COVID-19 to determine potential of using Panax ginseng to treat neurological disorders.	criteria.
[296]	Review on the currently employed COVID-19 management approaches and discussion of available treatment and prevention strategies.	Does not meet the inclusion criteria.
[297]	Description of the uptake and delivery of intravenous monoclonal antibodies, nirmatrelvir-ritonavir, and molnupiravir in the Torres and Cape region.	Does not meet the inclusion criteria.
[298]	Review of the current challenges of available COVID-19 treatments and drug development strategies for future therapy.	Does not meet the inclusion criteria.
[299]	Clinical trial finding that nirmatrelvir/ritonavir can significantly reduce the viral shedding time in elderly Chinese patients.	Does not meet the inclusion criteria.
[300]	No abstract available on pubmed.	Abstract unavailable.
[301]	No abstract available on pubmed.	Abstract unavailable.
[302]	Presentation of a patent of novel proline derivatives for the treatment of COVID-19.	Does not meet the inclusion criteria.
[303]	Discussion of access barriers to Paxlovid in low- and middle-income countries.	Does not meet the inclusion criteria.
[304]	No abstract available on pubmed.	Abstract unavailable.

Author(s)	Summary	Reason for Exclusion
[305]	Editorial with the aim of providing an update on current knowledge on COVID-19 rebound and how this may relate to currently available drugs.	Does not meet the inclusion criteria.
[306]	No abstract available on pubmed.	Abstract unavailable.
[307]	No abstract available on pubmed.	Abstract unavailable.
[308]	No abstract available on pubmed.	Abstract unavailable.
[309]	Report of 14 cases of COVID-19 patients, who are kidney transplant recipients coming to the conclusion that nirmatrelvir/ritonavir can be used in these patients, but should be closely monitored.	Does not meet the inclusion criteria.
[310]	Report of a cohort of 483 high-risk COVID-19 patients, who were being treated nirmatrelvir-ritonavir and of whom 2 required hospitalization, while 4 experienced rebound symptoms	Does not meet the inclusion criteria.
[311]	No abstract available on pubmed.	Abstract unavailable.
[312]	No abstract available on pubmed.	Abstract unavailable.
[313]	Examine the effect of nirmatrelvir/ritonavir and Chinese herb medicine lianhuaqingwen on PCR conversion time finding that early and aggressive administration of Paxlovid can significantly reduce the PCR conversion time.	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
[314]	Investigation of the MEK1/2 inhibitor ATR-002's ability to potentiate the effect of direct-acting antivirals (DAA) against SARS-CoV-2 on viral replication.	Does not meet the inclusion criteria.
[315]	Identification of 100 naturally occurring M <sup>pro</sup> mutations, located at the nirmatrelvir binding site, which might consequently inhibit Paxlovid's efficacy.	Does not meet the inclusion criteria.
[316]	Retrospective cohort study comparing the rate of progression to severe disease after receiving bebtelovimab or nirmatrelvir-ritonavir, coming to the conclusion that betelovimab's efficacy was not significantly different from nirmatrelvir/ritonavir's.	Does not meet the inclusion criteria.
[317]	Explore how nanotechnology provides potential alternative solutions to combat COVID-19.	Does not meet the inclusion criteria.
[318]	To improve currently available COVID-19 drugs, the authors suggest a combination of host- and virus-targeting antiviral drugs as it has the potential to synergistically suppress SARS-CoV-2 infection in Calu-3 lung epithelial cells.	Does not meet the inclusion criteria.
[319]	Review of the advances and challenges related to using nirmatrelvir to treat COVID-19 in order to provide insight to both drug developers and physicians treating COVID-19 patients.	Does not meet the inclusion criteria.
[320]	Review on the progress in	Does not meet the inclusion

Author(s)	Summary	Reason for Exclusion
	development of peptide drugs for use in the treatment of COVID-19.	criteria.
[321]	Evaluation of the trimeric derivative TF27 for its anti-SARS-CoV-2 potential coming to the conclusion that trioxane compounds should be considered for pharmacologic development towards future COVID-19 medication.	Does not meet the inclusion criteria.
[322]	Exploration of plant-derived antiviral small molecules to halt the viral entrance and in consequence serve as a component of future COVID-19 drugs.	Does not meet the inclusion criteria.
[323]	Description of the protocol for the Pediatric Observational Cohort Study of the NIH's RE searching COVID to Enhance Recovery (RECOVER) Initiative, focusing on two of five cohorts.	Does not meet the inclusion criteria.
[324]	Argue that nirmatrelvir/ritonavir and molnupiravir therapy is frequently underutilized. To overcome this underutilization, they describe the implementation of a pharmacy consult service for oral COVID-19 drugs within the ambulatory care setting.	Does not meet the inclusion criteria.
[325]	Structurally analyze the coronavirus main protease, focus on frequently occurring atoms and residues to find a generalized strategy for inhibitor design given a large	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	amount of protein complex from SARS-Cov.	
[326]	Long-term study of virological outcomes of immunodeficient individuals to uncover their (lack of) ability to sustain viral clearance.	Does not meet the inclusion criteria.
[327]	Discuss the possibility to reduce the ritonavir dosage to lower unwanted side effects and introduce another potent CYP3A inhibitor, cobicistat.	Does not meet the inclusion criteria.
[328]	Query the French national pharmacovigilance database to better understand nirmatrelvir/ritonavir's drug safety profile and potentially associated drug-drug interactions.	Does not meet the inclusion criteria.
[329]	Investigation of both GC376, a transition-state analog inhibitor of the main protease of feline infectious peritonitis coronavirus, and Nirmatrelvir to find that both are capable of strongly binding SARS-CoV-2's M <sup>Pro</sup> .	Does not meet the inclusion criteria.
[330]	Pursue an AI-driven approach followed by in vitro validation, to identify five fragment like M <sup>Pro</sup> inhibitors.	Does not meet the inclusion criteria.
[331]	Introduce latest version of COVID-19Base, a database summarizing the biomedical entities of COVID-19.	Does not meet the inclusion criteria.
[332]	Review summarizing the current developments in plant-derived drugs to combat COVID-19.	Does not meet the inclusion criteria.
[333]	Argue for a global Intellectual	Does not meet the inclusion

Author(s)	Summary	Reason for Exclusion
	Property waiver to meet the ongoing inequities of COVID-19, and to ensure equitable pandemic preparedness.	criteria.
[334]	Analysis of 213 critical, aged 50 years or younger, adult COVID-19 patients in South Korea to determine clinical characteristics and risk factors for mortality.	Does not meet the inclusion criteria.
[335]	Usage of virtual screening, molecular docking, and molecular dynamics simulations to identify potent phytocompounds as inhibitors of SARS-CoV-2 targets.	Does not meet the inclusion criteria.
[336]	Retrospectively analyze routine clinical data from outpatients with COVID-19 at high risk of severe outcomes to ascertain patient eligibility status and describe coverage of antiviral drugs and neutralising monoclonal antibodies (nMAB) as treatment for COVID-19 in community settings.	Does not meet the inclusion criteria.
[337]	Identification of prevalence of 3CL <sup>Pro</sup> inhibitor mutations that are associated with reduced susceptibility to nirmatrelvir/ritonavir.	Does not meet the inclusion criteria.
[338]	Descriptive study based on the French Pharmacovigilance Database to determine adverse drug reactions in COVID-19 patients.	Does not meet the inclusion criteria.
[339]	Review on the recent literature on drugs which were at the time under advanced clinical evaluation for COVID-19	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	treatment. They include antiviral agents as well as host-directed therapies.	
[340]	Usage of a network-based bioinformatics methodology to investigate the potential side effects of known Food and Drug Administration (FDA)-approved antihypertensive drugs, Sartans as well as application of the same methodology to nirmatrelvir/ritonavir.	Does not meet the inclusion criteria.
[341]	Report the inhibitory potentials of epigenetic-targeting drugs against SARS-CoV-2's M <sup>pro</sup> .	Does not meet the inclusion criteria.
[342]	Application of in vitro screen using the 2560 compounds from the Microsource Spectrum library to identify two additional small-molecule hits that can serve as PL <sup>pro</sup> and M <sup>pro</sup> inhibitors.	Does not meet the inclusion criteria.
[343]	Application of population pharmacokinetic modeling with clinical study data was employed to explore the pharmacokinetic profile of nirmatrelvir in older adult Chinese patients with COVID-19 infection	Does not meet the inclusion criteria.
[344]	Identification of the top 100 drugs most commonly prescribed to US patients who have a high risk of progressing to severe COVID-19. Each of these 100 drugs on the one hand and nirmatrelvir/ritonavir on the other hand were then examined for their drug-drug	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	interaction risk.	
[345]	Discussion of aspects of immune response in order to understand B and T cell lymphocyte regulatory mechanisms and antibody effectiveness and senescence plus presentation of COVID-19 therapies for non-hospitalized patients including limitations for identification, collection and distribution to infected patients.	Does not meet the inclusion criteria.
[346]	Screening of prognostic signature of lung adenocarcinoma/COVID-19 patients to identify the potential therapeutic targets and molecular pathways of nirmatrelvir/ritonavir.	Does not meet the inclusion criteria.
[347]	Discover subtle structural differences of nucleotide analogs that might lead to more potent SARS-CoV-2 inhibitors than the ones used in currently available COVID-19 drugs.	Does not meet the inclusion criteria.
[348]	Summary of literature, which was available at the time of publication, on nirmatrelvir/ritonavir, focusing on dosage, drug-drug interactions, toleration as well as how nirmatrelvir/ritonavir combats COVID-19 infections. Only efficacy study included is the clinical trial by [9].	Does not meet the inclusion criteria.
[349]	Review summarizing the clinical outcomes that have been obtained from the clinical trials published on antivirals,	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	immunomodulators, and other medications, such as antibiotics, stem cells, and plasma therapy in the treatment of COVID-19.	
[350]	Review on efficacy and safety of nirmatrelvir/ritonavir in, includes studies up until July 11, 2022.	Does not meet the inclusion criteria.
[351]	Review on nirmatrelvir/ritonavir. Explains how nirmatrelvir can inhibit SARS-CoV-2's M <sup>Pro</sup> , how ritonavir increases plasma concentration of nirmatrelvir, introduces the associated clinical studies and discusses possible safety concerns, the need for more studies who explore drug-drug interactions. Different focus than our work.	Does not meet the inclusion criteria.
[352]	Meta-analysis of relevant studies up to March, 2023 to estimate nirmatrelvir/ritonavir's efficacy compared with other antiviral drugs. Hence, their focus differs from ours.	Does not meet the inclusion criteria.
[353]	Review on efficacy and safety of nirmatrelvir/ritonavir. Control group consisted of patients who did not receive any COVID-19 medication. Included articles up to January 1, 2023. Hence, the studies we present from 2023 are not taken into account here.	Does not meet the inclusion criteria.
[354]	Review on efficacy and safety of nirmatrelvir/ritonavir. Included articles up to November 10, 2022. Hence, the	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	studies we present from 2023 are not taken into account here.	
[355]	Meta-analysis on efficacy and safety of nirmatrelvir/ritonavir. Included articles up to February 15, 2023. The studies we present from 2023 are not taken into account.	Does not meet the inclusion criteria
[356]	Review which presents currently available findings in the development of small molecules for COVID-19 treatment.	Does not meet the inclusion criteria.
[357]	Review on current knowledge of Paxlovid to explore how it may be used to treat future COVID-19 patients.	Does not meet the inclusion criteria.
[3]	Examine disparities in Paxlovid treatment, finding lower rates among Black and Hispanic or Latino patients and within socially vulnerable communities. Does not meet inclusion criteria.	Does not meet the inclusion criteria.
[358]	Review which summarizes Paxlovid's development journeys, undertaken clinical studies, regulatory approvals as well as current clinical trials and safety measures to end up with a discussion of drug-drug interactions and rebound effects.	Does not meet the inclusion criteria.
[359]	No abstract available on pubmed.	Does not meet the inclusion criteria.
[360]	Report of a phase 2/phase study of nirmatrelvir/ritonavir on patients with standard risk for developing severe COVID-19.	Does not meet the inclusion criteria.

Author(s)	Summary	Reason for Exclusion
	Here, the primary endpoint was self-reported sustained alleviation of all symptoms after 4 consecutive days. This endpoint was not met.	
[361]	Study on the efficacy of Paxlovid in elderly patients considering length of hospital stay, viral shedding time and rebound as outcomes.	Does not meet the inclusion criteria.
[362]	Review and discussion of efficacy and safety of nirmatrelvir/ritonavir by presenting the literature which was available at the time of publication. Does not include novel publications from 2023.	Does not meet the inclusion criteria.

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