

Chloroquine

Indications/Dosage

expand all | collapse all

Labeled

- amebiasis
- malaria
- malaria prophylaxis

Off-Label

- coronavirus disease 2019 (COVID-19) †
- discoid lupus erythematosus †
- severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) infection †

† Off-label indication

Per the manufacturer, this drug has been shown to be active against most strains of the following microorganisms either in vitro and/or in clinical infections:

Entamoeba histolytica, *Plasmodium falciparum*, *Plasmodium malariae*, *Plasmodium ovale*, *Plasmodium vivax*.

NOTE: The safety and effectiveness in treating clinical infections due to organisms with in vitro data only have not been established in adequate and well-controlled clinical trials.

This drug may also have activity against the following microorganisms:

severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2).

NOTE: Some organisms may not have been adequately studied during clinical trials; therefore, exclusion from this list does not necessarily negate the drug's activity against the organism.

INVESTIGATIONAL USE: For the treatment of severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) infection†, the virus that causes coronavirus disease 2019 (COVID-19)†

Oral dosage

- **Adults weighing 50 kg or more**

Available data are limited and efficacy has not been established. Due to a lack of clinical data, the National Institutes of Health (NIH) COVID-19 treatment guidelines do not give recommendations for or against the use of chloroquine; however, if used, guidelines advise monitoring for adverse events including QT interval prolongation.[\[65314\]](#) 1,000 mg (600 mg base) PO on day 1 then 500 mg (300 mg base) PO once daily for 4 to 7 days in patients in which enrollment in clinical trials is not feasible is suggested by the FDA in the Emergency Use Authorization (EUA) statement.[\[65171\]](#) [\[65172\]](#) 500 mg (300 mg base) PO twice daily for 10 days is also being evaluated alone and in combination. Preliminary data suggest chloroquine may inhibit the exacerbation of pneumonia, improve lung imaging findings, promote virus-negative conversion, and shorten the disease course.[\[65119\]](#) [\[65120\]](#) [\[65123\]](#) [\[65124\]](#) [\[65125\]](#) [\[65126\]](#) [\[65148\]](#)

- **Adults weighing less than 50 kg**

Available data are limited, and efficacy has not been established. Due to a lack of clinical data, the National Institutes of Health (NIH) COVID-19 treatment guidelines do not give recommendations for or against the use of chloroquine; however, if used, guidelines advise monitoring for adverse events including QT interval prolongation.[\[65314\]](#) 500 mg (300 mg base) PO twice daily for 10 days is being evaluated alone and in combination. Preliminary data suggest chloroquine may inhibit the exacerbation of pneumonia, improve lung imaging findings, promote virus-negative conversion, and shorten the disease course.[\[65119\]](#) [\[65120\]](#) [\[65123\]](#) [\[65124\]](#) [\[65125\]](#) [\[65126\]](#) [\[65148\]](#)

- **Adolescents weighing 50 kg or more**

Available data are limited, and efficacy has not been established. Due to a lack of clinical data, the National Institutes of Health (NIH) COVID-19 treatment guidelines do not give recommendations for or against the use of chloroquine; however, if used, guidelines advise monitoring for adverse events including QT interval prolongation.[\[65314\]](#) 1,000 mg (600 mg base) PO on day 1 then 500 mg (300 mg base) PO once daily for 4 to 7 days in patients in which enrollment in clinical trials is not feasible is suggested by the FDA in the Emergency Use Authorization (EUA) statement.[\[65171\]](#) [\[65172\]](#) Based on extrapolation from pediatric dosing for other indications and comparative doses to the adult dosing regimen suggested for COVID-19, 8.3 mg (5 mg base)/kg/dose PO twice daily [Max: 500 mg/dose (300 mg base/dose)] is being used in limited pediatric dosing protocols.[\[29758\]](#) [\[63245\]](#) [\[65159\]](#) A 10-day course is being used in adult patients.[\[65125\]](#) [\[65126\]](#) [\[65148\]](#)

- **Adolescents weighing less than 50 kg**

Efficacy and optimal dosing in pediatric patients are not established. Due to a lack of clinical data, the National Institutes of Health (NIH) COVID-19 treatment guidelines do not give recommendations for or against the use of chloroquine; however, if used, guidelines advise monitoring for adverse events including QT interval prolongation.[\[65314\]](#) Based on extrapolation from pediatric dosing for other indications and comparative doses to the

adult dosing regimen suggested for COVID-19, 8.3 mg (5 mg base)/kg/dose PO twice daily [Max: 500 mg/dose (300 mg base/dose)] is being used in limited pediatric dosing protocols.[29758] [63245] [65159] A 10-day course is being used in adult patients.[65125] [65126] [65148]

- **Infants and Children**

Efficacy and optimal dosing in pediatric patients are not established. Due to a lack of clinical data, the National Institutes of Health (NIH) COVID-19 treatment guidelines do not give recommendations for or against the use of chloroquine; however, if used, guidelines advise monitoring for adverse events including QT interval prolongation.[65314] Based on extrapolation from pediatric dosing for other indications and comparative doses to the adult dosing regimen suggested for COVID-19, 8.3 mg (5 mg base)/kg/dose PO twice daily [Max: 500 mg/dose (300 mg base/dose)] is being used in limited pediatric dosing protocols.[29758] [63245] [65159] A 10-day course is being used in adult patients.[65125] [65126] [65148]

For the treatment of uncomplicated malaria due to susceptible strains of *P. falciparum*, *P. knowlesi*†, *P. malariae*, *P. ovale*, and *P. vivax*

Oral dosage

- **Adults**

16.6 mg (10 mg base)/kg/dose [Max: 1,000 mg/dose (600 mg base/dose)] PO once, then 8.3 mg (5 mg base)/kg/dose [Max: 500 mg/dose (300 mg base/dose)] PO in 6 to 8 hours, then 8.3 mg (5 mg base)/kg/dose [Max: 500 mg/dose (300 mg base/dose)] PO once daily for 2 days.[29758] For *P. vivax* or *P. ovale*, give in combination with primaquine phosphate or tafenoquine. Guidelines recommend chloroquine for uncomplicated malaria in patients with chloroquine-sensitive *P. falciparum* or *P. vivax* or in all patients with *P. malariae*, *P. knowlesi*, or *P. ovale*. [64059]

- **Infants, Children, and Adolescents**

16.6 mg (10 mg base)/kg/dose [Max: 1,000 mg/dose (600 mg base/dose)] PO once, then 8.3 mg (5 mg base)/kg/dose [Max: 500 mg/dose (300 mg base/dose)] PO in 6 to 8 hours, then 8.3 mg (5 mg base)/kg/dose [Max: 500 mg/dose (300 mg base/dose)] PO once daily for 2 days.[29758] [63245] For *P. vivax* or *P. ovale*, give in combination with primaquine phosphate or tafenoquine. Guidelines recommend chloroquine for uncomplicated malaria in patients with chloroquine-sensitive *P. falciparum* or *P. vivax* or in all patients with *P. malariae*, *P. knowlesi*, or *P. ovale*. [64059]

For malaria prophylaxis against chloroquine-sensitive *Plasmodium* species

Oral dosage

- **Adults**

500 mg (300 mg base) PO weekly on the same day of each week, starting 2 weeks before entering the endemic area and continuing for 8 weeks after leaving the area. If it is not feasible to begin therapy before entering the endemic area, use 1,000 mg (600 mg base) as initial loading dose given in 2 divided doses 6 hours apart.[\[29758\]](#) Alternatively, guidelines suggest a shorter course; start the usual dosage regimen 1 to 2 weeks prior to entry into the endemic area and continue for 4 weeks after leaving the area.[\[63990\]](#)

- **Pregnant Female Adults[†]**

500 mg (300 mg base) PO weekly for duration of pregnancy for *P. ovale* or *P. vivax* infections after completing acute treatment. After delivery, subsequent treatment with primaquine phosphate or tafenoquine is needed in patients without G6PD deficiency.[\[64059\]](#)

- **Pregnant Female Adolescents[†]**

500 mg (300 mg base) PO weekly for duration of pregnancy for *P. ovale* or *P. vivax* infections after completing acute treatment. After delivery, subsequent treatment with primaquine phosphate or tafenoquine (16 years and older) is needed in patients without G6PD deficiency.[\[64059\]](#)

- **Infants, Children, and Adolescents**

8.3 mg (5 mg base)/kg/dose [Max: 500 mg/dose (300 mg base/dose)] PO weekly on the same day of each week, starting 2 weeks before entering the endemic area and continuing for 8 weeks after leaving the area. If it is not feasible to begin therapy before entering the endemic area, use 16.6 mg (10 mg base)/kg/dose [Max: 1,000 mg/dose (600 mg base/dose)] as initial loading dose given in 2 divided doses 6 hours apart.[\[29758\]](#) [\[63245\]](#) Alternatively, guidelines suggest a shorter course; start the usual dosage regimen 1 to 2 weeks prior to entry into the endemic area and continue for 4 weeks after leaving the area.[\[63245\]](#) [\[63990\]](#)

For the treatment of extraintestinal amebiasis (adjunct treatment with an effective intestinal amebicide)

Oral dosage

- **Adults**

1,000 mg (600 mg base) PO once daily for 2 days, then 500 mg (300 mg base) PO once daily for at least 2 to 3 weeks.[\[29758\]](#)

For the treatment of discoid lupus erythematosus†

Oral dosage

- **Adults**

125 to 250 mg (75 to 150 mg base) PO once daily. Do not exceed 3.5 to 4 mg/kg/day to minimize retinal toxicity. Chloroquine is recommended in patients who fail hydroxychloroquine plus quinacrine; quinacrine may be continued with chloroquine.[62154]

Therapeutic Drug Monitoring

The following recommendations are for baseline and continuous monitoring when using chloroquine with azithromycin:

- Obtain a pre-treatment QTc using a standard 12-lead ECG, telemetry, or mobile ECG device.
- Obtain baseline electrolytes, including calcium, magnesium, and potassium; correct abnormalities.
- Determine if the patient is currently on any QT-prolonging medications that can be discontinued.[65170]
- Document high-risk cardiovascular and comorbid conditions.[65170] Assess and adjust for hepatic and renal dysfunction.[65242]

Inpatient Use

- Place telemetry prior to initiation, if possible.
- Monitor and optimize serum electrolytes daily.[65242]
- If the baseline QTc is 500 msec or more and/or the patient has an inherent tendency to develop an exaggerated QTc response (i.e., change of 60 msec or more), correct contributing electrolyte abnormalities, review and discontinue other unnecessary QTc prolonging medications, and proceed with close QTc surveillance.[65170] Some experts recommend withholding treatment for patients with a baseline QTc of 500 msec or more (or more than 530 to 550 msec in patients with a QRS interval more than 120 msec) or in those with congenital long QT syndrome.[65242]
- If the baseline QTc is 460 to 499 msec (prepubertal), 470 to 499 msec (postpubertal males), or 480 to 499 msec (postpubertal females), correct contributing electrolyte abnormalities, review and discontinue other unnecessary QTc prolonging medications, and obtain an initial on-therapy QTc daily (or 48 and 96 hours after treatment initiation).[65170] [65242]

- If the baseline QTc is less than 460 msec (prepubertal), less than 470 msec (postpubertal males), or less than 480 msec (postpubertal females), correct electrolyte abnormalities and obtain an initial on-therapy QTc daily (or 48 and 96 hours after treatment initiation).[65170] [65242]
- Obtain an initial on-therapy QTc approximately 2 to 4 hours after the first dose and then daily (some recommend 48 and 96 hours after treatment initiation).[65170] [65242]
- Discontinue azithromycin and/or reduce the antimalarial dose if the subsequent QTc is prolonged or significantly increased above the specified parameters. If the QTc remains prolonged or significantly increased, reevaluate the risk/benefit of therapy, consider consultation with an electrophysiologist, and consider hydroxychloroquine/chloroquine discontinuation.[65242]

Outpatient Use

- Do not initiate outpatient therapy in the setting of acute renal or hepatic failure.[65242]
- If the baseline QTc is 500 msec or more and/or the patient has an inherent tendency to develop an exaggerated QTc response (i.e., change of 60 msec or more), correct contributing electrolyte abnormalities, review and discontinue other unnecessary QTc prolonging medications, and proceed with close QTc surveillance.[65170] Some experts recommend withholding treatment in patients with a baseline QTc of 480 msec or more (or more than 510 to 530 msec in patients with a QRS interval more than 120 msec), congenital long QT syndrome, or a Tisdale risk score of 11 or more.[65242]
- Consider no further ECG/telemetry assessment for patients with a Tisdale risk score of 6 or less, if resource or quarantine constraints are prohibitive of monitoring. Otherwise, repeat the ECG 2 to 3 hours after dosing on day 3 of therapy. If the QTc exceeds 500 msec (or 530 to 550 msec if QRS is more than 120 msec) or increases by more than 30 to 60 msec, consider discontinuing therapy.[65242]

Maximum Dosage Limits

- **Adults**

1,000 mg/dose (600 mg base/dose) PO for malaria up to a total of 2.5 g (1.5 g base) PO in 48 hours; 500 mg/week (300 mg base/week) PO for malaria prophylaxis; 1,000 mg/day (600 mg base/day) PO for other indications.

- **Geriatric**

1,000 mg/dose (600 mg base/dose) PO for malaria up to a total of 2.5 g (1.5 g base) PO in 48 hours; 500 mg/week (300 mg base/week) PO for malaria prophylaxis; 1,000 mg/day (600 mg base/day) PO for other indications.

- **Adolescents**

16.6 mg/kg/dose (10 mg base/kg/dose) [Max: 1,000 mg (600 mg base)] PO for malaria up to a total of 41.5 mg/kg (25 mg/kg base) [Max: 2.5 g (1.5 g base)] PO in 48 hours; 8.3 mg/kg/week (5 mg base/kg/week) [Max: 500 mg/week (300 mg base/week)] PO for malaria prophylaxis.

- Children

16.6 mg/kg/dose (10 mg base/kg/dose) [Max: 1,000 mg (600 mg base)] PO for malaria up to a total of 41.5 mg/kg (25 mg/kg base) [Max: 2.5 g (1.5 g base)] PO in 48 hours; 8.3 mg/kg/week (5 mg base/kg/week) [Max: 500 mg/week (300 mg base/week)] PO for malaria prophylaxis.

- Infants

16.6 mg/kg/dose (10 mg base/kg/dose) PO for malaria up to a total of 41.5 mg/kg (25 mg/kg base) PO in 48 hours; 8.3 mg/kg/week (5 mg base/kg/week) PO for malaria prophylaxis.

- Neonates

Safety and efficacy have not been established.

Patients with Hepatic Impairment Dosing

Chloroquine concentrates in the liver. However, no specific dosage adjustment guidelines are available for patients with hepatic impairment.[\[29758\]](#)

Patients with Renal Impairment Dosing

CrCl 10 mL/minute or more: No dosage adjustment necessary.

CrCl less than 10 mL/minute: Decrease dose by 50%.[\[32569\]](#)

Intermittent hemodialysis:

Decrease dose by 50%.[\[32569\]](#)

Peritoneal dialysis:

Decrease dose by 50%.[\[32569\]](#)

Continuous renal replacement therapy:

No dosage adjustment necessary.[\[32569\]](#)

† Off-label indication

Revision Date: 04/22/2020 12:27:21 PM

References

29758 – Aralen (chloroquine) package insert. Bridgewater, NJ: Sanofi-aventis U.S. LLC.; 2018 Oct.

32569 – Aronoff GR, Bennett WM, Berns JS, et al. Drug prescribing in renal failure: dosing guidelines for adults and children, 5th ed. Philadelphia: American College of Physicians; 2007.

62154 – Okon LG, Werth VP. Cutaneous lupus erythematosus: diagnosis and treatment. *Best Pract Res Clin Rheumatol* 2013;27:391-404.

63245 – American Academy of Pediatrics. Red Book: 2018-2021 Report of the Committee on Infectious Diseases. 31st ed. Elk Grove Village, IL: American Academy of Pediatrics; 2018.

63990 – Arguin PM, Tan KR. Chapter 3. Infectious diseases related to travel. Malaria. In. Centers for Disease Control and Prevention. 2018 Yellow Book - Traveler's Health. Atlanta: U.S. Department of Health and Human Services, Public Health Service.
<https://wwwnc.cdc.gov/travel/yellowbook/2018/infectious-diseases-related-to-travel/malaria>

64059 – Centers for Disease Control and Prevention. Treatment of malaria (guidelines for clinicians). Atlanta, GA: US Department of Health and Human Services; 2019.
https://www.cdc.gov/malaria/diagnosis_treatment/treatment.html.

65119 – Gao J, Tian Z, Yang X. Breakthrough: Chloroquine Phosphate Has Shown Apparent Efficacy in Treatment of COVID-19 Associated Pneumonia in Clinical Studies. *Biosci Trends* 2020 Feb 19. [Epub ahead of print]

65120 – Wang M, Cao R, Zhang L, et al. Remdesivir and Chloroquine Effectively Inhibit the Recently Emerged Novel Coronavirus (2019-nCoV) in Vitro. *Cell Res* 2020;30(3):269-271.

65123 – World Health Organization (WHO). Coronavirus: landscape analysis of therapeutics as of 17 February 2020. Retrieved March 16, 2020. Available on the World Wide Web at
https://www.who.int/blueprint/priority-diseases/key-action/Table_of_therapeutics_Appendix_17022020.pdf?ua=1.

65124 – Colson P, Rolain J, Lagier J, et al. Chloroquine and hydroxychloroquine as available weapons to fight COVID-19. *Int J Antimicrob Agents*. 2020 (Epub ahead of print)

65125 – Dong L, Hu S, Gao J. Discovering drugs to treat coronavirus disease 2019 (COVID-19). *Drug Discov Ther* 2020;14:58-60.

65126 – Multicenter Collaboration Group of Department of Science and Technology of Guangdong Province and Health Commission of Guangdong Province for chloroquine in the treatment of novel coronavirus pneumonia. Expert consensus on chloroquine phosphate for the treatment of novel coronavirus pneumonia. *Zhonghua Jie He He Hu Xi Za Zhi* 2020 Mar;43:185-8.

65148 – Italian Society of Infectious and Tropical Diseases. Handbook for the care of people with disease-COVI 19. Edition 2.0, March 13, 2020.

65159 – American Society of Transplantation and Cellular Therapy Infectious Diseases Special Interest Group. Interim Guidelines for COVID-19: Management in Hematopoietic Cell Transplant and Cellular Therapy Patients. March 18, 2020. Available on the world wide web at: [https://higherlogicdownload.s3.amazonaws.com/ASBMT/a1e2ac9a-36d2-4e23-945c-45118b667268/UploadedImages/COVID-19 Interim Patient Guidelines 3 18 20.pdf](https://higherlogicdownload.s3.amazonaws.com/ASBMT/a1e2ac9a-36d2-4e23-945c-45118b667268/UploadedImages/COVID-19%20Interim%20Patient%20Guidelines%203%2018%2020.pdf)

65170 – Giudicessi JR, Noseworthy PA, Friedman PA, Ackerman MJ. Urgent guidance for navigating and circumventing the QTc prolonging and torsadogenic potential of possible pharmacotherapies for COVID-19 [published online ahead of print, March 25, 2020]. Mayo Clin Proc 2020;95.



65171 – Food and Drug Administration (FDA). Chloroquine phosphate or hydroxychloroquine sulfate supplied from the strategic national stockpile for treatment of 2019 coronavirus disease: emergency use authorization letter. Retrieved March 30, 2020. Available on the World Wide Web at <https://www.fda.gov/emergency-preparedness-and-response/mcm-legal-regulatory-and-policy-framework/emergency-use-authorization#2019-ncov>.

65172 – Food and Drug Administration (FDA). Fact sheet for health care providers: emergency use authorization (EUA) of chloroquine phosphate supplied from the strategic national stockpile for treatment of COVID-19 in certain hospitalized patients. Retrieved March 30, 2020. Available on the World Wide Web at <https://www.fda.gov/emergency-preparedness-and-response/mcm-legal-regulatory-and-policy-framework/emergency-use-authorization#2019-ncov>.

65242 – Simpson TF, Kovacs RJ, Stecker EC. Ventricular arrhythmia risk due to hydroxychloroquine-azithromycin treatment for COVID-19. Accessed April 10, 2020. Available at: <https://www.acc.org/latest-in-cardiology/articles/2020/03/27/14/00/ventricular-arrhythmia-risk-due-to-hydroxychloroquine-azithromycin-treatment-for-covid-19>

65314 – COVID-19 treatment guidelines panel. COVID-19 treatment guidelines. National Institutes of Health Web site. Accessed April 22, 2020. Available at: <https://covid19treatmentguidelines.nih.gov/>.

How Supplied

Chloroquine Hydrochloride Solution for injection	
Aralen 50mg/ml Solution for Injection (00024-0074) (Sanofi U.S. LLC) (off market)	
Chloroquine Phosphate Oral tablet	
Chloroquine Phosphate 250mg Tablet (00143-1195) (Hikma Pharmaceuticals USA inc.) (off market)	
Chloroquine Phosphate 250mg Tablet (00143-1195) (Hikma Pharmaceuticals USA Inc.) (off market)	
Chloroquine Phosphate 250mg Tablet (00115-2790) (Impax Generics, a division of Impax Laboratories, Inc.) (off market)	

Chloroquine Phosphate Oral tablet	
	
Chloroquine Phosphate 250mg Tablet (00115-7056) (Impax Generics, a division of Impax Laboratories, Inc.) (off market)	
Chloroquine Phosphate 250mg Tablet (64980-0177) (Rising Pharmaceuticals Inc) (off market)	
Chloroquine Phosphate 250mg Tablet (64980-0177) (Rising Pharmaceuticals Inc)	
Chloroquine Phosphate 250mg Tablet (63304-0460) (Sun Pharmaceutical Industries, Inc.) (off market)	
Aralen 500mg Tablet (00280-0084) (Bayer Corp Consumer Care Div) (off market)	
Aralen 500mg Tablet (00024-0084) (Sanofi U.S. LLC) (off market)	 
Chloroquine Phosphate 500mg Tablet (00143-2125) (Hikma Pharmaceuticals USA Inc.) (off market)	
Chloroquine Phosphate 500mg Tablet (00115-7010) (Impax Generics, a division of Impax Laboratories, Inc.) (off market)	
Chloroquine Phosphate 500mg Tablet (55289-0856) (PD-Rx Pharmaceuticals, Inc.)	
Chloroquine Phosphate 500mg Tablet (43063-0454) (PD-Rx Pharmaceuticals, Inc.)	
Chloroquine Phosphate 500mg Tablet (64980-0178) (Rising Pharmaceuticals Inc) (off market)	
Chloroquine Phosphate 500mg Tablet (64980-0178) (Rising Pharmaceuticals Inc)	

Chloroquine Phosphate Oral tablet
Chloroquine Phosphate 500mg Tablet (63304-0461) (Sun Pharmaceutical Industries, Inc.) (off market)

Description/Classification

Description

Chloroquine is a 4-aminoquinoline anti-protozoal agent indicated for the treatment and prophylaxis of susceptible malaria strains and for the treatment of extraintestinal amebiasis. Chloroquine is not active against gametocytes and the exoerythrocytic forms including the hypnozoite stage of the *Plasmodium* parasites. Resistance to chloroquine is widespread. Irreversible retinal damage has been observed with use, and postmarketing cases of life-threatening and fatal cardiomyopathy, including ventricular arrhythmias and torsade de pointes (TdP), have been reported.[29758]

Updates for coronavirus disease 2019 (COVID-19):

Available data are limited. Due to a lack of clinical data, the National Institutes of Health (NIH) COVID-19 treatment guidelines do not give recommendations for or against the use of chloroquine; however, if used, guidelines advise monitoring for adverse events including QT interval prolongation.[65314] The FDA has issued an Emergency Use Authorization (EUA) for the use of chloroquine to treat COVID-19 patients for whom clinical trial participation is not feasible. [65171][65172] Preliminary data suggest chloroquine may have clinical benefit in the treatment of COVID-19 due to SARS-CoV-2. In results for more than 100 patients, chloroquine was superior to control in inhibiting the exacerbation of pneumonia.[65119][65124][65126] Additional data regarding clinical efficacy for COVID-19 are being evaluated.[65123]

Classifications

- [General Anti-infectives Systemic](#)
 - [Antiparasitic Agents, Insecticides, and Repellants](#)
 - [Antiprotozoals](#)
 - [Agents for Amoebiasis and Other Protozoal Diseases](#)
 - [Antimalarials](#)

Revision Date: 04/22/2020 11:23:53 PM

References

29758 – Aralen (chloroquine) package insert. Bridgewater, NJ: Sanofi-aventis U.S. LLC.; 2018 Oct.

65119 – Gao J, Tian Z, Yang X. Breakthrough: Chloroquine Phosphate Has Shown Apparent Efficacy in Treatment of COVID-19 Associated Pneumonia in Clinical Studies. Biosci Trends 2020 Feb 19. [Epub ahead of print]

65123 – World Health Organization (WHO). Coronavirus: landscape analysis of therapeutics as of 17 February 2020. Retrieved March 16, 2020. Available on the World Wide Web at https://www.who.int/blueprint/priority-diseases/key-action/Table_of_therapeutics_Appendix_17022020.pdf?ua=1.

65124 – Colson P, Rolain J, Lagier J, et al. Chloroquine and hydroxychloroquine as available weapons to fight COVID-19. Int J Antimicrob Agents. 2020 (Epub ahead of print)

65126 – Multicenter Collaboration Group of Department of Science and Technology of Guangdong Province and Health Commission of Guangdong Province for chloroquine in the treatment of novel coronavirus pneumonia. Expert consensus on chloroquine phosphate for the treatment of novel coronavirus pneumonia. Zhonghua Jie He He Hu Xi Za Zhi 2020 Mar;43:185-8.

65171 – Food and Drug Administration (FDA). Chloroquine phosphate or hydroxychloroquine sulfate supplied from the strategic national stockpile for treatment of 2019 coronavirus disease: emergency use authorization letter. Retrieved March 30, 2020. Available on the World Wide Web at <https://www.fda.gov/emergency-preparedness-and-response/mcm-legal-regulatory-and-policy-framework/emergency-use-authorization#2019-ncov>.

65172 – Food and Drug Administration (FDA). Fact sheet for health care providers: emergency use authorization (EUA) of chloroquine phosphate supplied from the strategic national stockpile for treatment of COVID-19 in certain hospitalized patients. Retrieved March 30, 2020. Available on the World Wide Web at <https://www.fda.gov/emergency-preparedness-and-response/mcm-legal-regulatory-and-policy-framework/emergency-use-authorization#2019-ncov>.

65314 – COVID-19 treatment guidelines panel. COVID-19 treatment guidelines. National Institutes of Health Web site. Accessed April 22, 2020. Available at: <https://covid19treatmentguidelines.nih.gov/>.

Administration Information

General Administration Information

For storage information, see the specific product information within the How Supplied section.

Route-Specific Administration

Oral Administration

- May administer with meals in patients who experience gastrointestinal side effects. [63990]

Oral Solid Formulations

- To mitigate bitter tablet taste for children, tablets may be pulverized and enclosed in gelatin capsules. If the child is unable to swallow the capsules or tablets, the gelatin capsules may be opened and the contents mixed with a small amount of something sweet, such as applesauce, chocolate syrup, or jelly.[\[63990\]](#)

Extemporaneous Compounding-Oral

NOTE: Chloroquine extemporaneous suspension is not FDA-approved.

Extemporaneous chloroquine suspension has been compounded using the following formulations:

- Chloroquine phosphate 15 mg/mL in 1:1 Ora-Sweet and Ora-Plus, 1:1 Ora-Sweet SF and Ora-Plus, or Cherry Syrup:
 - Pulverize three 500-mg chloroquine phosphate tablets into a fine powder in a mortar.
 - Add approximately 15 mL of vehicle, which may be a 1:1 mixture of Ora-Sweet and Ora-Plus, 1:1 mixture of Ora-Sweet SF and Ora-Plus, or cherry syrup.
 - Add the vehicle in geometric portions almost to volume and mix thoroughly after each addition.
 - Transfer the contents of the mortar to a calibrated bottle.
 - Add enough vehicle to bring the final volume to 100 mL.
 - Label "Shake Well Before Using" and "Protect from Light"
 - *Storage:* The suspension is stable for 60 days when stored without light at 5 and 25 degrees C.[\[62145\]](#)
- Chloroquine base 10 mg/mL in Cherry Syrup:
 - Pulverize two 500-mg chloroquine phosphate tablets in a mortar after removing the film coating.
 - Levigate with a small amount of sterile water.
 - Add by geometric proportions a significant amount of cherry syrup, and levigate until a uniform mixture is obtained.
 - Transfer the contents of the mortar to a conical graduated cylinder.
 - Add enough cherry syrup to bring the final volume to 60 mL.
 - Pour the suspension into an amber glass bottle and shake vigorously.
 - *Storage:* The suspension is stable for up to 4 weeks under refrigeration (4 degrees C), at room temperature (22 to 25 degrees C), and at 29 degrees C.[\[62150\]](#)
- Chloroquine base 15 mg/mL in Glycerin or Distilled Water, Cologel (Lilly), and Simple Syrup/Cherry Syrup:
 - Pulverize two 500-mg chloroquine phosphate tablets in a mortar.
 - Levigate with a small amount of glycerin or distilled water.
 - Add 13 mL of Cologel, and levigate until a uniform mixture is obtained.
 - Add by geometric proportion a significant amount of a 2:1 simple syrup/cherry syrup mixture, and levigate until a uniform mixture is obtained.
 - Transfer the contents of the mortar to a conical graduated cylinder.
 - Add enough of the syrup mixture to bring the final volume to 40 mL.
 - Pour the suspension into an amber glass bottle and shake vigorously.
 - Label "Shake Well" and "Refrigerate".

- *Storage:* The suspension is stable for 3 days when stored in the refrigerator. [62172]
-

Clinical Pharmaceutics Information

From Trissel's 2TM Clinical Pharmaceutics Database

Chloroquine

1. pH Range

pH 5.5 to 6.6 (as the hydrochloride salt)

References

McEvoy GK (ed). AHFS Drug Information (current edition). Bethesda, MD: American Society of Health-System Pharmacists.

2. Stability

Chloroquine sulfate in intact containers stored as directed by the manufacturer is stable until the labeled expiration date. Infusion Solutions: Chloroquine sulfate 0.5 mg/mL in sodium chloride 0.9% was reported by Martens et al. to be stable for 24 hours at 21 degree C protected from exposure to light.

References

Anon. Manufacturer's information and labeling. (Package insert).

Martens HJ, De Goede PN, van Loenen AC. Sorption of various drugs in polyvinyl chloride, glass, and polyethylene-lined infusion containers. *Am J Hosp Pharm.* 1990; 47:369-73

3. Light Exposure

Chloroquine injection should be protected from exposure to light during long-term storage.

References

Anon. Manufacturer's information and labeling. (Package insert).

Anon. BIAM; l'Universite et l'Industrie Pharmaceutique, Banque de donnees automatisee sur les medicaments (Biam), Paris, France. Accessed at: <http://www2.biam2.org>.

Anon. South African Electronic Package Inserts, Malahyde Information Systems.
Accessed at <http://home.intekom.com/pharm/intramed>.

4. Filtration

Geary et al. reported that chloroquine exhibited binding to cellulose acetate filters. About 60% or more of the drug was lost from 10 mL of a chloroquine 0.32-mg/mL solution passed through Millipore and Nalgene 0.45-micron cellulose acetate filters. No drug loss occurred when the chloroquine solution was passed through a polycarbonate filter. The clinical implications, if any, of binding to filters from such a low concentration are uncertain.

References

Geary TG, Akood MA, Jensen JB. Characteristics of chloroquine binding to glass and plastic. *Am J Trop Med Hyg.* 1984; 32:19-23

5. Sorption Leaching

Geary et al. reported that chloroquine at low concentrations exhibits sorption to glass. As much as 30 to 40% was lost to glass test tubes from a 0.32 mg/mL solution. No loss due to sorption to polycarbonate, polystyrene, or polypropylene plastics was found. The clinical implications, if any, of sorption from such a low concentration are uncertain. Moreover, Martens et al. were unable to confirm this result. They found no sorption to glass bottles, polyvinyl chloride (PVC) plastic bags, and polyethylene-lined laminated bags from a chloroquine 0.5 mg/mL (as sulfate) solution.

References

D'Arcy PF. Drug interactions with medical plastics. *Drug Intell Clin Pharm.* 1083; 17:726-31

Geary TG, Akood MA, Jensen JB. Characteristics of chloroquine binding to glass and plastic. *Am J Trop Med Hyg.* 1984; 32:19-23

Martens HJ, De Goede PN, van Loenen AC. Sorption of various drugs in polyvinyl chloride, glass, and polyethylene-lined infusion containers. *Am J Hosp Pharm.* 1990; 47:369-73

6. Stability Max

Maximum reported stability period: In NS- 24 hours at room temperature

References

Martens HJ, De Goede PN, van Loenen AC. Sorption of various drugs in polyvinyl chloride, glass, and polyethylene-lined infusion containers. *Am J Hosp Pharm.* 1990; 47:369-73

Compounding Drug Information

From Trissel's 2TM Clinical Pharmaceutics Database

Chloroquine

1. Identity/Properties

Chloroquine is a white to slightly yellow, odorless crystalline powder with a bitter taste. The hydrochloride, a white crystalline substance, is prepared using hydrochloric acid. Chloroquine hydrochloride 123 mg is approximately equivalent to 100 mg of chloroquine. The phosphate is a white or almost white, odorless, hygroscopic crystalline powder with a bitter taste. Chloroquine phosphate 161 mg is approximately equivalent to 100 mg of chloroquine. Solubility: Chloroquine is only slightly soluble in water. The hydrochloride and the phosphate are freely soluble in water. The phosphate has an aqueous solubility of about 250 mg/mL but is almost insoluble in ethanol. pH: Chloroquine hydrochloride injection has a pH between 5.5 and 6.5. A 1% chloroquine phosphate solution has a pH of about 4.5.

References

Anon. Manufacturer's information and labeling. (Package insert and bulk material data sheet).

Anon. The Merck Index, Whitehouse Station, New Jersey: Merck & Co., Inc. Current edition.

Anon. Martindale The Complete Drug Reference. London: The Pharmaceutical Press. Current edition and selected information from prior editions.

Anon. The United States Pharmacopeia. Rockville, Maryland: The United States Pharmacopeial Convention. Current edition.

McEvoy GK (ed). AHFS Drug Information (current edition). Bethesda, MD: American Society of Health-System Pharmacists.

2. General Stability Info

Chloroquine products should be stored in well-closed containers. The injection (as the hydrochloride) should be stored at controlled room temperature and protected from freezing and temperatures exceeding 40degree C. The phosphate is light sensitive, discoloring upon light exposure.

References

Anon. Martindale The Complete Drug Reference. London: The Pharmaceutical Press. Current edition and selected information from prior editions.

Anon. The United States Pharmacopeia. Rockville, Maryland: The United States Pharmacopeial Convention. Current edition.

McEvoy GK (ed). AHFS Drug Information (current edition). Bethesda, MD: American Society of Health-System Pharmacists.

3. Injection, extemporaneous

Injections, like other sterile drugs, should be prepared in a suitable clean air environment using appropriate aseptic procedures. When prepared from non-sterile components, an appropriate and effective sterilization method must be employed. Allen reported on a compounded formulation of chloroquine phosphate 64.5-mg/mL injection. The injection had the following formula: Chloroquine phosphate- 6.45 g Benzyl alcohol- 2 g Sterile water for injection- qs 100 mL The recommended method of preparation is to dissolve the chloroquine phosphate powder in about 90 mL of sterile water for injection. The benzyl alcohol is then added and stirred until dissolved. Sterile water for injection sufficient to bring the volume to 100 mL is added and the solution is mixed well. The solution is to be filtered through a suitable 0.2-micron sterilizing filter and packaged in sterile containers. If no sterility test is performed, the USP specifies a beyond-use date of 24 hours at room temperature or three days stored under refrigeration because of concern for inadvertent microbiological contamination during preparation. However, if an official USP sterility test for each batch of drug is performed, the author recommended a beyond-use date of six months at room temperature because this formula is similar or the same as a commercial medication in some countries with an expiration date of two years or more.

References

Allen LV Jr. Chloroquine phosphate 64.5-mg/mL injection. *Int J Pharmaceut Compound.* 2009; 13:154

4. Oral Liquid

Study 1: Closson reported that chloroquine hydrochloride injection (Aralen HCl, Sanofi Winthrop) was added to simple syrup to make a 20-mg/mL oral pediatric dosage form. The product was incubated at 49degree C for 63 hours; no visible changes in physical appearance or consistency occurred. The product then was frozen at -6degree C for eight hours; it became a white frozen solid and reliquified upon warming to its original colorless, slightly hazy appearance. No chemical analysis was performed. Study 2: Allen and Erickson evaluated the stability of three chloroquine phosphate 15-mg/mL oral liquids extemporaneously compounded from tablets. Vehicles used in this study were (1) an equal parts mixture of Ora-Sweet and Ora-Plus (Paddock), (2) an equal parts mixture of Ora-Sweet SF and Ora-Plus (Paddock), and (3) cherry syrup (Robinson Laboratories) mixed 1:4 with simple syrup. Three chloroquine phosphate 500-mg tablets (Sanofi Winthrop) were crushed and comminuted to fine powder using a mortar and pestle. About 15 mL of the test vehicle was added to the powder and mixed to yield a uniform paste. Additional vehicle was added geo-metrically and brought to the final volume of 100 mL, mixing thoroughly after each addition. The process was repeated for each of the three test suspension vehicles. Samples of each of the finished suspensions were packaged in 120-mL amber poly-ethylene terephthalate plastic prescription bottles and stored at 5degree C and 25degree C. Because the

phosphate salt is freely soluble in water, the drug is in solution in these products. No visual changes or changes in odor were detected during the study. Stability-indicating HPLC analysis found little or no drug loss in any of the liquid products stored at either temperature after 60 days of storage. Study 3: Odusote and Nasipuri evaluated the stability of three syrup formulations (See Table 1 below) containing chloroquine phosphate 16 mg/mL prepared from bulk powder. The sucrose syrup formulation was prepared by adding the chloroquine phosphate powder and sodium benzoate to heated syrup and stirring until complete dissolution occurred. The flavor and color were then added. The other two formulations were prepared by dispersing the methylcellulose in some hot water followed by ice-cold water and subsequently keeping the mixture in a freezer. The other materials then were added and mixed well. After preparation, the syrups were packaged in amber bottles and stored at 5, 25, and 40 degree C for 12 weeks. Chloroquine content was assessed spectrophotometrically. No change in chloroquine concentration and no observable physical change were found in samples from any storage temperature during storage. If clear bottles were used instead of amber, exposure to light resulted in about 6% drug loss in eight weeks. If the syrup pH was adjusted from the original pH 4.5 to 4.9 down to pH 3.5 with citric acid (used to help mask the bitter taste), the chloroquine concentration remained constant over 12 weeks at all temperatures. However, if 1% sodium carboxymethylcellulose was substituted as the viscosity agent, a white turbidity or precipitate appeared (depending on concentration) along with a sudden drop in viscosity, indicating an interaction with the chloroquine phosphate. Consequently, Odusote and Nasipuri recommended using only methylcellulose as the viscosity-imparting agent. Table 1. Chloroquine Phosphate Formulations Tested for Stability by Odusote and Nasipuri

Formula 1: Chloroquine phosphate 1.6 g Sodium benzoate 0.2 g Essence of lemon grass 0.5 mL Yellow food color 0.2 mL Sucrose syrup 84% (w/v) qs 100mL

Formula 2: Chloroquine phosphate 1.6 g Sodium benzoate 0.2 g Saccharin sodium 0.05 g Essence of lemon grass 0.5 mL Yellow food color 0.2 mL Methylcellulose 1.12% solution qs 100 mL

Formula 3: Chloroquine phosphate 1.6 g Sodium benzoate 0.2 g Talin 0.05 g Essence of lemon grass 0.5 mL Yellow food color 0.2 mL Methylcellulose 1.12% solution qs 100 mL

Study 4: Mirochnick et al. attempted to determine the stability of a chloroquine phosphate suspension prepared from tablets. Although no loss of drug was found by HPLC analysis, substantial increases in drug concentration at various time points indicated a nonuniform dispersion of the drug might have existed. Study 5: Van Doorne et al. evaluated the suitability of several antimicrobial preservatives for use in chloroquine phosphate 16-mg/mL syrup containing sucrose 66%. Chloroform had been used previously, but it is carcinogenic, potentially toxic to liver and kidneys, and is volatile resulting in loss of protection over time. Benzalkonium chloride is unsuitable because of its taste and incompatibilities. The best result was obtained using sorbic acid 1.5 g/L along with citric acid 2 g/L to reduce the pH to 4. Methylparaben 1.8 g/L with propylparaben 0.2 g/L also was acceptable, although the latter system was not as effective against *Aspergillus niger*. Study 6: Chandibhamar et al. evaluated the stability of a taste-masked suspension of chloroquine. Chloroquine phosphate 1.7 g was dissolved in 40 mL of simple syrup containing glycerin 5% (v/v). Then 30 mL of hot syrup containing pamoate sodium 1.3 g and sodium bicarbonate 0.56 g was slowly mixed in at a rate of 5 mL per minute with constant stirring of 30 to 80 RPM. A precipitate of chloroquine pamoate was produced. The suspension was adjusted to pH 6.0 and sodium benzoate 100 mg, amaranth, and raspberry flavor were added. The suspension was brought to 100 mL with additional syrup. The suspension exhibited a very slow rate of sedimentation. Spectroscopic analysis found the suspension remained stable over at least 42 days at 25 degree C with over 98% of the ion pair remaining and only 1.75% of free chloroquine present. Elevated temperature of 45 degree C resulted

in an approximate doubling of the rate of free chloroquine formation. Bioavailability of this suspension was comparable with chloroquine phosphate liquid.

References

Allen LV Jr, Erickson MA. Stability of alprazolam, chloroquine phosphate, cisapride, enalapril maleate, and hydralazine hydrochloride in extemporaneously compounded oral liquids. *Am J Health-Syst Pharm.* 1998; 55:1915-20

Chandibhammar V, Yadav MR, Murthy RSR. Studies on the development of taste-masked suspension of chloroquine. *Boll Chim Farm.* 2004; 143:377-82

Closson RG. Liquid dosage form of chloroquine. *Drug Intell Clin Pharm.* 1988; 22:347

Mironchnick M, Barnett E, Clarke DF, et al. Stability of chloroquine in an extemporaneously prepared suspension stored at three temperatures. *Pediatr Infect Dis J.* 1994; 13:817-8

Oduote MO, Nasipuri RN. Effect of pH and storage conditions on the stability of a novel chloroquine phosphate syrup formulation. *Pharm Ind.* 1988; 50:367-9

Van Doorne H, Wieringa NF, Bosch EH, et al. The suitability of some preservatives in chloroquine phosphate syrup. *Pharm Weekbl Sci Ed.* 1988; 10:170-2

5. Topical

Brouwers et al. developed a topical gel containing chloroquine phosphate for use as a microbicide against HIV-1 infection. The gel was prepared by adding a mixture of hydroxyethyl cellulose 1.6% wt/wt and glycerol 2.5% to a solution of methyl- and propylparabens 0.18% and 0.02%, respectively. When mixed, a clear and homogeneous gel formed. The pH of the gel was decreased by adding lactic acid 0.05% and adjusting to pH 4.5 by adding sodium hydroxide 1 M. Chloroquine phosphate powder was added to the gel in varying amounts of 0.3, 1.3, 3, 10, and 30 mg/g of gel and mixed thoroughly to assure complete dissolution and uniformity. Entrapped air was removed by using reduced pressure. The completed gels were packaged in capped syringes. The gels were clear and homogeneous with an osmolality of 300 mOsm/kg, a pH of 4.6, and a viscosity of 1.4 Pa s. Samples containing chloroquine phosphate 3 mg/g were stored at 40degree C and 75% relative humidity for three months. Little or no change in gel mass, pH, and osmolality occurred. Viscosity decreased by 14%, which is similar to changes observed in previously reported observations for hydroxyethyl cellulose gels. HPLC analysis found that little or no change in chloroquine concentrations occurred over the three-month test period.

References

Brouwers J, Vermeire K, Schols D, et al. Development and in vitro evaluation of chloroquine gels as microbicides against HIV-1 infection. *Virology.* 2008; 378:306-10

References

62145 – Allen LV, Erickson MA. Stability of alprazolam, chloroquine phosphate, cisapride, enalapril maleate, and hydralazine hydrochloride in extemporaneously compounded oral liquids. *Am J Health-Syst Pharm* 1998;55:1915-20.

62150 – Mirochnick M, Barnett E, Clarke DF, et al. Stability of chloroquine in an extemporaneously prepared suspension stored at three temperatures. *Pediatr Infect Dis J* 1994;13:827-8.

62172 – ASHP (American Society of Hospital Pharmacists). Committee on extemporaneous formulations. *Handbook on Extemporaneous Formulations*. Bethesda MD: ASHP, 1987.

63990 – Arguin PM, Tan KR. Chapter 3. Infectious diseases related to travel. Malaria. In. Centers for Disease Control and Prevention. 2018 Yellow Book - Traveler's Health. Atlanta: U.S. Department of Health and Human Services, Public Health Service.

<https://wwwnc.cdc.gov/travel/yellowbook/2018/infectious-diseases-related-to-travel/malaria>

Adverse Reactions

- abdominal pain
- acute generalized exanthematous pustulosis (AGEP)
- agitation
- agranulocytosis
- alopecia
- anaphylactic shock
- anaphylactoid reactions
- angioedema
- anorexia
- anxiety
- aplastic anemia
- AV block
- blurred vision
- bundle-branch block
- cardiomyopathy
- confusion
- corneal deposits
- corneal opacification
- delirium
- depression
- diarrhea
- Drug Reaction with Eosinophilia and Systemic Symptoms (DRESS)
- dyskinesia
- elevated hepatic enzymes
- erythema multiforme
- exfoliative dermatitis
- hair discoloration
- hallucinations
- headache
- hearing loss
- hemolysis
- hemolytic anemia
- hepatitis
- hypoglycemia
- hyporeflexia
- hypotension
- insomnia
- macular degeneration
- myasthenia
- myopathy
- nausea
- neutropenia
- night blindness
- pancytopenia
- photosensitivity
- pruritus
- psoriasis
- psychosis
- QT prolongation
- retinopathy
- scotomata
- seizures
- skin discoloration
- Stevens-Johnson syndrome
- suicidal ideation
- thrombocytopenia
- tinnitus
- torsade de pointes
- torticollis
- toxic epidermal necrolysis
- urticaria
- ventricular fibrillation

- ventricular tachycardia
 - visual impairment
 - vomiting
 - weakness
-

Irreversible maculopathy and macular degeneration have been reported with chloroquine or other 4-aminoquinoline compounds during postmarketing use. Irreversible retinopathy with retinal pigment changes (bull's eye appearance) and visual field defects (paracentral scotomas) have been reported in patients receiving long-term or high-dose 4-aminoquinoline therapy. Visual impairment (i.e., blurred vision and difficulty in focusing or accommodation), nyctalopia (night blindness), scotomatous vision with field defects of paracentral, pericentral ring types, and typically temporal scotomata (e.g., difficulty in reading with words tending to disappear, seeing half an object, misty vision, and fog before the eyes), and reversible corneal opacification (corneal deposits) have been reported. For patients with significant risk factors, monitoring should include annual examinations which include best corrected distance visual acuity (BCVA), automated threshold visual field (VF), and spectral domain optical coherence tomography (SD-OCT). For individuals without significant risk factors, annual exams can usually be deferred until 5 years of treatment. Discontinue chloroquine if ocular toxicity is suspected, and monitor the patient closely as retinal changes and visual disturbances may progress after cessation of therapy.[\[29758\]](#)

Erythema multiforme, Stevens-Johnson syndrome, toxic epidermal necrolysis, exfoliative dermatitis, pleomorphic skin eruptions, skin and mucosal pigmentary changes (skin discoloration), lichen planus-like eruptions, pruritus, drug reaction with eosinophilia and systemic symptoms (DRESS), photosensitivity, hair loss (alopecia), bleaching of hair pigment (hair discoloration), urticaria, anaphylactoid reactions or anaphylactic shock, and angioedema have been reported during postmarketing use of chloroquine or other 4-aminoquinoline compounds. An acute attack of psoriasis can be precipitated by chloroquine in predisposed patients.[\[29758\]](#)

Chloroquine has been associated with acute generalized exanthematous pustulosis (AGEP). The non-follicular, pustular, erythematous rash starts suddenly and is associated with fever above 38 degrees C. Drugs are the main cause of AGEP. A period of 2 to 3 weeks after an inciting drug exposure appears necessary for a first episode of AGEP. Unintentional reexposure may cause a second episode within 2 days.[\[27736\]](#)

Hematological adverse reactions have been reported during postmarketing use of chloroquine or other 4-aminoquinoline compounds and include reversible agranulocytosis, aplastic anemia, pancytopenia, neutropenia, and thrombocytopenia. Chloroquine may cause hemolysis and hemolytic anemia in patients with glucose-6-phosphate dehydrogenase deficiency (G6PD deficiency).[\[29758\]](#)

During postmarketing use, chloroquine and/or other 4-aminoquinoline compounds have been associated with sensorimotor disorders as well as skeletal muscle myopathy or neuromyopathy leading to progressive weakness (myasthenia) and atrophy of proximal muscle groups, depressed tendon reflexes (hyporeflexia), and abnormal nerve conduction. Periodically test knee and ankle reflexes to detect any evidence of muscular weakness. Discontinue chloroquine if weakness develops.[\[29758\]](#)

Nerve type deafness, tinnitus, and reduced hearing (hearing loss) in patients with preexisting auditory damage have been reported during postmarketing use of chloroquine or other 4-aminoquinoline compounds. Discontinue chloroquine with any hearing defects, and monitor the patient closely.[\[29758\]](#)

Cardiovascular adverse reactions associated with chloroquine or other 4-aminoquinoline compounds during postmarketing include cardiomyopathy (which may result in cardiac failure and in some cases fatal outcome), electrocardiogram (ECG) changes (particularly, inversion or flattening of the T-wave with widening of the QRS complex), and hypotension. Cardiac arrhythmias, conduction disorders such as bundle-branch block and AV block, QT prolongation, torsade de pointes (TdP), ventricular arrhythmias (e.g., ventricular tachycardia, ventricular fibrillation) have been reported, including fatal cases. The risk is greater with higher doses, although cases have been reported with therapeutic doses. Chronic toxicity should be considered when conduction disorders, such as bundle-branch block or AV block, are diagnosed. Additionally, cases of cardiomyopathy resulting in cardiac failure with some cases of fatal outcome have been reported with chloroquine. Prompt discontinuation of chloroquine may prevent life-threatening complications if cardiotoxicity is suspected.[28225] [28229] [28230] [28231] [29758]

Adverse gastrointestinal effects noted with chloroquine or other 4-aminoquinoline compounds during postmarketing include hepatitis, elevated hepatic enzymes, nausea, vomiting, abdominal pain/cramps, diarrhea, and anorexia.[29758] Gastric effects can be minimized by taking chloroquine with food.[61673]

Nervous system adverse reactions associated with chloroquine or other 4-aminoquinoline compounds during postmarketing include headache (usually mild and transient), seizures, polyneuropathy, acute extrapyramidal symptoms (e.g., dystonia, dyskinesia, tongue protrusion, torticollis), and neuropsychiatric changes including psychosis, delirium, anxiety, agitation, insomnia, hallucinations, confusion, personality changes, depression, and suicidal ideation/behavior. Extrapyramidal symptoms usually resolve after treatment discontinuation and/or symptomatic treatment.[29758]

Chloroquine has been shown to cause severe hypoglycemia including loss of consciousness that could be life threatening in patients treated with or without antidiabetic medications. Monitor blood glucose and adjust treatment as necessary in patients presenting with clinical symptoms of hypoglycemia during chloroquine treatment.[29758]

Revision Date: 08/02/2017 10:28:02 AM

References

27736 – Beylot C, Doutré M, Beylot-Barry M. Acute generalized exanthematous pustulosis. *Semin Cutan Med Surg* 1996;15(4):244-249.

28225 – CredibleMeds. Drugs to avoid in congenital long QT. Available on the World Wide Web at <http://www.crediblemeds.org>.

28229 – Demazière J, Fourcade JM, Busseuil CT, et al. The hazards of chloroquine self prescription in west Africa. *J Toxicol Clin Toxicol* 1995;33:369-70.

28230 – Mansfield RJ, Thomas RD. Recurrent syncope. Drug induced long QT syndrome. *Postgrad Med J* 2001;77:344, 352-3.

28231 – Pinski SL, Eguía LE, Trohman RG. What is the minimal pacing rate that prevents torsades de pointes? Insights from patients with permanent pacemakers. *Pacing Clin Electrophysiol* 2002;25:1612-5.

29758 – Aralen (chloroquine) package insert. Bridgewater, NJ: Sanofi-aventis U.S. LLC.; 2018 Oct.

61673 – Anon. Drugs for Parasitic Infections. Treatment Guidelines from The Medical Letter 2013;11S:e1-31.

Contraindications/Precautions

Absolute contraindications are italicized.

- *chloroquine hypersensitivity*
 - *ocular disease*
 - accidental exposure
 - alcoholism
 - antimicrobial resistance
 - apheresis
 - Asian patients
 - AV block
 - bradycardia
 - breast-feeding
 - cardiomyopathy
 - celiac disease
 - children
 - diabetes mellitus
 - Fabry disease
 - females
 - fever
 - G6PD deficiency
 - geriatric
 - hearing impairment
 - heart failure
 - hepatic disease
 - human immunodeficiency virus (HIV) infection
 - hydroxychloroquine hypersensitivity
 - hyperparathyroidism
 - hypocalcemia
 - hypoglycemia
 - hypokalemia
 - hypomagnesemia
 - hypothermia
 - hypothyroidism
 - infants
 - infertility
 - long QT syndrome
 - myocardial infarction
 - neonates
 - neurological disease
 - pheochromocytoma
 - porphyria
 - pregnancy
 - psoriasis
 - QT prolongation
 - renal failure
 - renal impairment
 - rheumatoid arthritis
 - seizure disorder
 - seizures
 - sickle cell disease
 - sleep deprivation
 - stroke
 - systemic lupus erythematosus (SLE)
-

Chloroquine is reported in the literature to be a weak genotoxic agent that may elicit both gene mutations and chromosomal/DNA breaks. Mechanisms may involve DNA intercalation or induction of oxidative stress. Both positive and negative results have been reported with in vitro reverse gene mutation assays and with in vivo animal studies. The chromosomal effects were not observed when chloroquine was administered to animals orally.[29758]

Antimicrobial resistance to chloroquine therapy is widespread in *P. falciparum* and is reported in *P. vivax*. Prior to chloroquine use, it should be ascertained whether chloroquine is appropriate for use based on resistance patterns. Information regarding the geographic areas where resistance to chloroquine occurs is available from the Centers for Disease Control and Prevention.[29758]

Use of chloroquine for indications other than acute malaria is contraindicated in patients with *ocular disease*, specifically those who have retinal or visual field changes of any etiology. Irreversible retinal damage has been observed in some patients who received chloroquine. Significant risk factors for retinal damage include daily doses of chloroquine phosphate more than 2.3 mg/kg of actual body weight, duration of use more than 5 years, subnormal glomerular filtration (renal impairment or renal failure), use of some concomitant drug products such as tamoxifen, and concurrent macular disease. Baseline ophthalmological examination should be performed within the first year of starting chloroquine and should include best corrected distance visual acuity (BCVA), automated threshold visual field (VF) of the central 10 degrees (with retesting if an abnormality is noted), and spectral domain optical coherence tomography (SD-OCT). In Asian patients, retinal toxicity may first be noticed outside the macula, and VF testing should be performed in the central 24 degrees instead of the central 10 degrees. For patients with significant risk factors, monitoring should include annual examinations which include BCVA, VF, and SD-OCT. For individuals without significant risk factors, annual exams can usually be deferred until 5 years of treatment. Discontinue chloroquine if ocular toxicity is suspected, and monitor the patient closely as retinal changes and visual disturbances may progress after cessation of therapy.[29758] The use of chloroquine should be approached with caution in patients with Fabry disease, particularly those with ocular symptoms. The drug can cause a keratopathy that is clinically and ultrastructurally indistinguishable from keratopathy caused by Fabry disease; this drug-induced keratopathy is reversible with drug cessation. In addition, chloroquine poses a theoretical risk of decreased intracellular alpha-galactosidase A activity in Fabry disease patients. Chloroquine has been reported to induce clinical symptoms that mimic those of Fabry disease, including formation of inclusion bodies that are biochemically and ultrastructurally similar in most of the cells affected by Fabry disease (e.g., striated muscle, smooth muscle, etc.). The distinguishing factor is that the ultrastructural features of chloroquine toxicity in striated muscle, curvilinear bodies, are not present in renal cells.[30609]

Chloroquine is contraindicated in patients with known *chloroquine hypersensitivity*, or with a known allergy to 4-aminoquinolines. Patients with hydroxychloroquine hypersensitivity may have cross sensitivity to chloroquine.[29758]

QT prolongation, torsade de pointes (TdP), and ventricular arrhythmias have been reported with chloroquine use. The risk is greater with higher doses, although cases have been reported with therapeutic doses.[28225] [28229] [28230] [28231] [29758] Use chloroquine with caution in patients with conditions that may increase the risk of QT prolongation including congenital long QT syndrome, bradycardia, AV block, heart failure, stress-related cardiomyopathy, myocardial infarction, stroke, hypomagnesemia, hypokalemia, hypocalcemia, or in patients receiving medications known to prolong the QT interval or cause electrolyte imbalances. Females, geriatric patients, patients with sleep deprivation, pheochromocytoma, sickle cell disease, hypothyroidism, hyperparathyroidism, hypothermia, systemic inflammation (e.g., human immunodeficiency virus (HIV) infection, fever, and some autoimmune diseases including rheumatoid arthritis, systemic lupus erythematosus (SLE), and celiac disease) and patients undergoing apheresis procedures (e.g., plasmapheresis [plasma exchange], cytophoresis) may also be at increased risk for QT prolongation.[28432] [28457] [29758] [56592].[65180] In patients taking chloroquine with another drug that also prolongs the QT interval (see Therapeutic Drug Monitoring for recommendations specific to azithromycin with chloroquine used together for COVID-19), obtain a pre-treatment QTc using a standard 12-lead ECG, telemetry, or mobile ECG device. Obtain baseline electrolytes, including calcium, magnesium, and potassium. Determine if the patient is currently on any QT-prolonging medications that can be discontinued. Document high-risk cardiovascular and comorbid conditions. If the baseline QTc is 500 msec or more and/or the patient has an inherent tendency to develop an exaggerated QTc response (i.e., change of 60 msec or more), correct contributing electrolyte abnormalities, review and discontinue other unnecessary QTc prolonging medications, and proceed with close QTc surveillance. Obtain an initial on-therapy QTc

approximately 2 to 4 hours after the first dose and then again at 48 and 96 hours after treatment initiation. If the baseline QTc is 460 to 499 msec (prepubertal), 470 to 499 msec (postpubertal males), or 480 to 499 msec (postpubertal females), correct contributing electrolyte abnormalities, review and discontinue other unnecessary QTc prolonging medications, and obtain an initial on-therapy QTc 48 and 96 hours after treatment initiation. If the baseline QTc is less than 460 msec (prepubertal), less than 470 msec (postpubertal males), or less than 480 msec (postpubertal females), correct electrolyte abnormalities and obtain an initial on-therapy QTc 48 and 96 hours after treatment initiation.[\[65170\]](#) Consider chronic chloroquine toxicity when conduction disorders, such as bundle-branch block or AV block, are diagnosed. Cases of cardiomyopathy resulting in cardiac failure, sometimes fatal, have been reported with chloroquine. Prompt discontinuation of chloroquine may prevent life-threatening complications if cardiotoxicity is suspected.[\[29758\]](#)

Chloroquine should not be used in patients with psoriasis unless the benefit to the patient outweighs the potential risks because it may precipitate a severe attack of psoriasis.[\[29758\]](#)

Chloroquine should be used with caution in patients with hepatic disease or alcoholism because the drug is metabolized in the liver and accumulation can occur producing toxic effects. Patients receiving other hepatotoxic drugs also should be treated with caution.[\[29758\]](#)

Chloroquine should be used with caution in patients with neurological disease including preexisting hearing impairment or seizure disorder. Polyneuritis, ototoxicity, seizures, neuromyopathy, and acute extrapyramidal symptoms (dystonia, dyskinesia, tongue protrusion, torticollis) have occurred with chloroquine therapy. Symptoms of muscle weakness and response of knee and ankle reflexes should be investigated regularly. If muscle weakness, extrapyramidal symptoms, or any defects in hearing occur during chloroquine therapy, the drug should be discontinued immediately and the patient observed closely.[\[29758\]](#)

Chloroquine can exacerbate porphyria or may cause hemolysis in patients with glucose-6-phosphate dehydrogenase deficiency (G6PD deficiency). Use chloroquine with caution in patients with these conditions. Blood monitoring for hemolytic anemia in G6PD deficiency patients may be necessary, particularly with concomitant use of other medications associated with hemolysis.[\[29758\]](#)

Use chloroquine with caution in patients with hypoglycemia or diabetes mellitus. Chloroquine can cause severe, life-threatening hypoglycemia in patients with or without antidiabetic medications. Warn patients about the risk of hypoglycemia and the associated clinical signs and symptoms. Monitor blood glucose and adjust treatment as necessary in patients presenting with clinical symptoms of hypoglycemia during chloroquine treatment.[\[29758\]](#)

Children are especially sensitive to the 4-aminoquinoline compounds. Fatalities have been reported after accidental exposure of chloroquine; some cases involved relatively small doses (e.g., 0.75 g or 1 g in a 3-year-old child). Strongly warn patients to keep chloroquine out of the reach of pediatric patients, including neonates, infants, children, and adolescents.[\[29758\]](#)

Weigh the benefit of chloroquine prophylaxis or treatment of malaria against the potential risk to the fetus, and consider the drug's potential to remain in the body for several months after discontinuation of therapy.[\[29758\]](#) [\[63990\]](#) [\[64059\]](#) In humans at recommended doses for prophylaxis and treatment of malaria, observational studies as well as a meta-analysis, including a small number of prospective studies with chloroquine during pregnancy, have shown no increase in the rate of birth defects or spontaneous abortions.[\[29758\]](#) Guidelines recommend chloroquine as a treatment option for acute malaria and for prophylaxis in pregnant women during all trimesters. Chloroquine crosses the placenta, but the potential damage to the mother from malaria is greater than the drug's risk to the fetus. Weekly prophylactic doses appear to have minimal adverse effects when administered during pregnancy.[\[63990\]](#) [\[64059\]](#) Animal studies showed embryo-fetal

developmental toxicity at doses 3 to 16 times the maximum recommended therapeutic dose and the potential of genotoxicity in some test systems. Autoradiographic studies have shown accumulation in the eyes and ears when chloroquine is administered at the start or end of gestation in animal studies.[29758]

Use caution when administering chloroquine to breast-feeding women. Chloroquine is excreted into breast milk. The excretion of chloroquine and the major metabolite, desethylchloroquine, in breast milk was investigated in 11 lactating mothers following a single oral dose of chloroquine (600 mg base). The maximum daily dose of the drug that the infant received from breast-feeding was about 0.7% of the maternal start dose of the drug in malaria chemotherapy. Separate chemoprophylaxis for an infant is required.[29758] However, previous American Academy of Pediatrics (AAP) recommendations consider chloroquine usually compatible with breast-feeding, and chloroquine has an established dosage in infants.[27500]

Chloroquine should be used with caution in males because animal studies suggest that infertility is possible; after 30 days of oral treatment, testosterone levels and weight of testes, epididymis, seminal vesicles, and prostate decreased.[29758]

Revision Date: 04/10/2020 04:22:06 PM

References

- 27500** – American Academy of Pediatrics (AAP) Committee on Drugs. Transfer of drugs and other chemicals into human milk. *Pediatrics* 2001;108(3):776-789.
- 28225** – CredibleMeds. Drugs to avoid in congenital long QT. Available on the World Wide Web at <http://www.crediblemeds.org>.
- 28229** – Demaziere J, Fourcade JM, Busseuil CT, et al. The hazards of chloroquine self prescription in west Africa. *J Toxicol Clin Toxicol* 1995;33:369-70.
- 28230** – Mansfield RJ, Thomas RD. Recurrent syncope. Drug induced long QT syndrome. *Postgrad Med J* 2001;77:344, 352-3.
- 28231** – Pinski SL, Eguia LE, Trohman RG. What is the minimal pacing rate that prevents torsades de pointes? Insights from patients with permanent pacemakers. *Pacing Clin Electrophysiol* 2002;25:1612-5.
- 28432** – Roden, DM. Drug-induced prolongation of the QT interval. *New Engl J Med* 2004;350:1013-22.
- 28457** – Crouch MA, Limon L, Cassano AT. Clinical relevance and management of drug-related QT interval prolongation. *Pharmacotherapy* 2003;23:881-908.
- 29758** – Aralen (chloroquine) package insert. Bridgewater, NJ: Sanofi-aventis U.S. LLC.; 2018 Oct.
- 30609** – Albay D, Adler SG, Philipose J, et al. Chloroquine-induced lipodosis mimicking Fabry disease. *Mod Pathol* 2005;18:733-8.
- 56592** – van Noord C, Eijgelsheim M, Stricker BH. Drug- and non-drug-associated QT interval prolongation. *Br J Clin Pharmacol* 2010;70(1):16-23.

63990 – Arguin PM, Tan KR. Chapter 3. Infectious diseases related to travel. Malaria. In. Centers for Disease Control and Prevention. 2018 Yellow Book - Traveler's Health. Atlanta: U.S. Department of Health and Human Services, Public Health Service.
<https://wwwnc.cdc.gov/travel/yellowbook/2018/infectious-diseases-related-to-travel/malaria>

64059 – Centers for Disease Control and Prevention. Treatment of malaria (guidelines for clinicians). Atlanta, GA: US Department of Health and Human Services; 2019.
https://www.cdc.gov/malaria/diagnosis_treatment/treatment.html.

65170 – Giudicessi JR, Noseworthy PA, Friedman PA, Ackerman MJ. Urgent guidance for navigating and circumventing the QTc prolonging and torsadogenic potential of possible pharmacotherapies for COVID-19 [published online ahead of print, March 25, 2020]. *Mayo Clin Proc* 2020;95.

65180 – Woosley RL, Heise CW, Gallo T, et al. QTFactors List. Oro Valley, AZ: AZCERT, Inc.; Accessed March 31, 2020. Available on the World Wide Web at: <https://crediblemeds.org/ndfa-list/>

Mechanism of Action

Chloroquine, a 4-aminoquinoline, is an anti-protozoal agent. The precise mechanism is unknown. Chloroquine may exert its effect against *Plasmodium* species by concentrating in the acid vesicles of the parasite and by inhibiting polymerization of heme. It can also inhibit certain enzymes by its interaction with DNA. Chloroquine is not active against gametocytes and the exoerythrocytic forms, including the hypnozoite stage (*P. vivax* and *P. ovale*) of the *Plasmodium* parasites. Organisms with reduced susceptibilities to hydroxychloroquine also show reduced susceptibilities to chloroquine.[29758]

Although the mechanisms underlying the antiinflammatory and immunomodulatory effects of chloroquine are unknown, several possible mechanisms of action have been proposed. It is unclear if these mechanisms work similarly for rheumatic and autoimmune diseases. Potential mechanisms include reduced cytokine production, inhibition of immune effector cells, inhibition of platelet function, protection of the cell surface from external disturbances, competitive binding to nucleic acid ligands or toll-like receptors (TLRs), interference with lysosomal function, reduction of leakage of lysosomal enzymes, and interference with endosomal NADPH oxidase (NOX).[61727] [61728] [61729]

There are several potential mechanisms by which chloroquine may be active against severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2). These include inhibition of viral enzymes or processes such as viral DNA and RNA polymerase, viral protein glycosylation, virus assembly, new virus particle transport, and virus release. Other mechanisms may also involve ACE2 cellular receptor inhibition, acidification at the surface of the cell membrane inhibiting fusion of the virus, and immunomodulation of cytokine release.[61729] [61732] [65120] [65121] [65139] [65140] [65141] [65142] [65143]

References

- 29758** – Aralen (chloroquine) package insert. Bridgewater, NJ: Sanofi-aventis U.S. LLC.; 2018 Oct.
- 61727** – Muller-Calleja N, Manukyan D, Canisius A, et al. Hydroxychloroquine inhibits proinflammatory signalling pathways by targeting endosomal NADPH oxidase. *Ann Rheum Dis* 2016. Epub ahead of print, doi: 10.1136/annrheumdis-2016-210012.
- 61728** – Kuznik A, Bencina M, Svajger U, et al. Mechanism of endosomal TLR inhibition by antimalarial drugs and imidazoquinolines. *J Immunol* 2011;186:4794-804.
- 61729** – Fox R. Anti-malarial drugs: possible mechanisms of action in autoimmune disease and prospects for drug development. *Lupus* 1996;S1:S4-10.
- 61732** – Ben-Zvi H, Kivity S, Langevitz P, et al. Hydroxychloroquine: from malaria to autoimmunity. *Clin Rev Allergy Immunol* 2012;42:145-53.
- 65120** – Wang M, Cao R, Zhang L, et al. Remdesivir and Chloroquine Effectively Inhibit the Recently Emerged Novel Coronavirus (2019-nCoV) in Vitro. *Cell Res* 2020;30(3):269-271.
- 65121** – Yao X, Fei Y, Zhang M, et al. In Vitro Antiviral Activity and Projection of Optimized Dosing Design of Hydroxychloroquine for the Treatment of Severe Acute Respiratory Syndrome Coronavirus 2 (SARS-CoV-2). *Clin Infect Dis* 2020 Mar 9. [Epub ahead of print]
- 65139** – Fox RI. Mechanism of action of hydroxychloroquine as an antirheumatic drug. *Semin Arthritis Rheum* 1993;23(2 S1):82-91.
- 65140** – Savarino A, Trani LD, Donatelli I, et al. New insights into the antiviral effects of chloroquine. *The Lancet* 2006;6:67-9.
- 65141** – Vincent MJ, Bergeron E, Benjannet S, et al. Chloroquine is a potent inhibitor of SARS coronavirus infection and spread. *Virology* 2005;2:69.
- 65142** – Cortegiani A, Ingoglia G, Ippolito M, et al. A systematic review on the efficacy and safety of chloroquine for the treatment of COVID-19. *J Crit Care* 2020. [In press]
- 65143** – Cohen SN, Yielding KL. Inhibition of DNA and RNA polymerase reactions by chloroquine. *Proc Natl Acad Sci U S A* 1965; 54: 521-7.

Pharmacokinetics

Chloroquine is administered orally. It is widely distributed into body tissues, with higher concentrations in the liver, kidneys, spleen, and lungs. Leukocytes also concentrate the drug. Smaller amounts of the drug are found in the brain and spinal cord. Cells containing melanin in the eyes and skin bind strongly to chloroquine. The drug also concentrates in erythrocytes and is bound to platelets and granulocytes. It is about 55% bound to plasma protein.[\[29758\]](#) [\[61731\]](#) [\[62151\]](#)

Excretion of chloroquine is largely through urine, but this is a slow process and may be increased by acidification of the urine. Chloroquine undergoes appreciable degradation in the body, and the major metabolite is desethylchloroquine. Slightly more than half of a dose is excreted in urine as unchanged drug and about 25% as the major metabolite; bisdesethylchloroquine and other metabolic products are found in small amounts. A small portion of the unabsorbed drug is excreted in the feces. Elimination appears to take place in a biphasic manner. The elimination half-life is 108 to 291 hours.[29758] [61731] [62151]

Affected cytochrome P450 isoenzymes and drug transporters: CYP2C8, CYP2D6, CYP3A4, P-gp

In vitro data suggest that chloroquine is metabolized primarily by CYP2C8 and CYP3A4, and to a much lesser extent, by CYP2D6.[65236] [65239] It has also been shown to be an inhibitor of the drug transporter P-glycoprotein (P-gp).[65237]

Route-Specific Pharmacokinetics

- **Oral Route**

After oral administration, chloroquine is rapidly and almost completely absorbed from the gastrointestinal tract.[29758][62151] The T_{max} is 2.7 to 6.9 hours with a C_{max} of 283 to 1,430 ng/mL and an AUC of 8.2 to 140 mcg x hour/mL.[62151] A single study showed that the AUC in patients with malaria was higher than in normal volunteers (281 vs. 122 mcg x mL/L x hour).[61760]

Revision Date: 04/13/2020 02:16:15 PM

References

29758 – Aralen (chloroquine) package insert. Bridgewater, NJ: Sanofi-aventis U.S. LLC.; 2018 Oct.

61731 – Mackenzie AH. Pharmacologic actions of 4-aminoquinoline compounds. Am J Med 1983;75:5-10.

61760 – Furst DE. Pharmacokinetics of hydroxychloroquine and chloroquine during treatment of rheumatic diseases. Lupus 1996;5(S1):S11-15.

62151 – Guidelines for the Treatment of Malaria. 3rd edition. Geneva: World Health Organization; 2015. Available on the worldwide web: <https://www.ncbi.nlm.nih.gov/books/NBK294440/>.

65236 – Schrezenmeier E, Dorner T. Mechanisms of action of hydroxychloroquine and chloroquine: implications for rheumatology. Rheumatology 2020;16:155-166.

65237 – Tiberghien F, Loo F. Ranking of P-glycoprotein substrates and inhibitors by a calcein-AM fluorometry screening assay. Anti-Cancer Drugs 1996;7:568-578.

65239 – Projean D, Baune B, Farinotti R, et al. In vitro metabolism of chloroquine: Identification of CYP2C8, CYP3A4, and CYP2D6 as the main isoforms catalyzing N-desethylchloroquine formation. *Drug Metab Dispos* 2003;31:748-754.

Pregnancy/Breast-feeding

Pregnancy

Weigh the benefit of chloroquine prophylaxis or treatment of malaria against the potential risk to the fetus, and consider the drug's potential to remain in the body for several months after discontinuation of therapy.[29758] [63990] [64059] In humans at recommended doses for prophylaxis and treatment of malaria, observational studies as well as a meta-analysis, including a small number of prospective studies with chloroquine during pregnancy, have shown no increase in the rate of birth defects or spontaneous abortions.[29758] Guidelines recommend chloroquine as a treatment option for acute malaria and for prophylaxis in pregnant women during all trimesters. Chloroquine crosses the placenta, but the potential damage to the mother from malaria is greater than the drug's risk to the fetus. Weekly prophylactic doses appear to have minimal adverse effects when administered during pregnancy.[63990] [64059] Animal studies showed embryo-fetal developmental toxicity at doses 3 to 16 times the maximum recommended therapeutic dose and the potential of genotoxicity in some test systems. Autoradiographic studies have shown accumulation in the eyes and ears when chloroquine is administered at the start or end of gestation in animal studies.[29758]

Breast-Feeding

Use caution when administering chloroquine to breast-feeding women. Chloroquine is excreted into breast milk. The excretion of chloroquine and the major metabolite, desethylchloroquine, in breast milk was investigated in 11 lactating mothers following a single oral dose of chloroquine (600 mg base). The maximum daily dose of the drug that the infant received from breast-feeding was about 0.7% of the maternal start dose of the drug in malaria chemotherapy. Separate chemoprophylaxis for an infant is required.[29758] However, previous American Academy of Pediatrics (AAP) recommendations consider chloroquine usually compatible with breast-feeding, and chloroquine has an established dosage in infants.[27500]

Revision Date: 04/15/2019 05:39:21 PM

References

27500 – American Academy of Pediatrics (AAP) Committee on Drugs. Transfer of drugs and other chemicals into human milk. *Pediatrics* 2001;108(3):776-789.

29758 – Aralen (chloroquine) package insert. Bridgewater, NJ: Sanofi-aventis U.S. LLC.; 2018 Oct.

63990 – Arguin PM, Tan KR. Chapter 3. Infectious diseases related to travel. Malaria. In. Centers for Disease Control and Prevention. 2018 Yellow Book - Traveler's Health. Atlanta: U.S. Department of Health and Human Services, Public Health Service.
<https://wwwnc.cdc.gov/travel/yellowbook/2018/infectious-diseases-related-to-travel/malaria>

64059 – Centers for Disease Control and Prevention. Treatment of malaria (guidelines for clinicians). Atlanta, GA: US Department of Health and Human Services; 2019.
https://www.cdc.gov/malaria/diagnosis_treatment/treatment.html.

Interactions

Level 1 (Severe)

- Cisapride
- Dronedarone
- Pimozide
- Thioridazine

Level 2 (Major)

- Acarbose
- Acetohexamide
- Aclidinium; Formoterol
- Albiglutide
- Albuterol
- Albuterol; Ipratropium
- Alfuzosin
- Alogliptin
- Alogliptin; Metformin
- Alogliptin; Pioglitazone
- Alpha-glucosidase Inhibitors
- Aluminum Hydroxide
- Aluminum Hydroxide; Magnesium Carbonate
- Aluminum Hydroxide; Magnesium Hydroxide
- Aluminum Hydroxide; Magnesium Hydroxide; Simethicone
- Aluminum Hydroxide; Magnesium Trisilicate
- Amiodarone
- Amitriptyline
- Amitriptyline; Chlordiazepoxide
- Amoxicillin; Clarithromycin; Lansoprazole
- Amoxicillin; Clarithromycin; Omeprazole
- Anagrelide
- Antacids
- Apomorphine
- Arformoterol
- Aripiprazole
- Arsenic Trioxide
- Artemether; Lumefantrine
- Asenapine
- Aspirin, ASA; Citric Acid; Sodium Bicarbonate
- Atomoxetine
- Azithromycin
- Bedaquiline
- Bismuth Subcitrate Potassium; Metronidazole; Tetracycline
- Bismuth Subsalicylate; Metronidazole; Tetracycline
- Budesonide; Formoterol
- Buprenorphine
- Buprenorphine; Naloxone
- Calcium Carbonate
- Calcium Carbonate; Magnesium Hydroxide
- Calcium Carbonate; Risedronate
- Calcium Carbonate; Simethicone
- Canagliflozin
- Canagliflozin; Metformin
- Ceritinib
- Chlorpromazine
- Chlorpropamide
- Cimetidine
- Ciprofloxacin
- Citalopram
- Clarithromycin
- Clofazimine
- Clomipramine
- Clozapine
- Codeine; Phenylephrine; Promethazine
- Codeine; Promethazine
- Crizotinib
- Cyclosporine
- Dapagliflozin
- Dapagliflozin; Metformin
- Dapagliflozin; Saxagliptin
- Dasatinib
- Degarelix
- Desflurane

- Desipramine
- Deutetrabenazine
- Dextromethorphan; Promethazine
- Dextromethorphan; Quinidine
- Dipeptidyl Peptidase-4 Inhibitors
- Disopyramide
- Dofetilide
- Dolasetron
- Dolutegravir; Rilpivirine
- Donepezil
- Donepezil; Memantine
- Doxepin
- Droperidol
- Dulaglutide
- Efavirenz
- Efavirenz; Emtricitabine; Tenofovir
- Efavirenz; Lamivudine; Tenofovir Disoproxil Fumarate
- Eliglustat
- Empagliflozin
- Empagliflozin; Linagliptin
- Empagliflozin; Linagliptin; Metformin
- Empagliflozin; Metformin
- Emtricitabine; Rilpivirine; Tenofovir alafenamide
- Emtricitabine; Rilpivirine; Tenofovir disoproxil fumarate
- Encorafenib
- Enflurane
- Entrectinib
- Eribulin
- Ertugliflozin
- Ertugliflozin; Metformin
- Ertugliflozin; Sitagliptin
- Erythromycin
- Erythromycin; Sulfisoxazole
- Escitalopram
- Exenatide
- Ezogabine
- Fingolimod
- Flecainide
- Fluconazole
- Fluoxetine
- Fluoxetine; Olanzapine
- Fluphenazine
- Fluticasone; Salmeterol
- Fluticasone; Umeclidinium; Vilanterol
- Fluticasone; Vilanterol
- Fluvoxamine
- Formoterol
- Formoterol; Mometasone
- Foscarnet
- Gemifloxacin
- Gemtuzumab Ozogamicin
- Gilteritinib
- Glasdegib
- Glimepiride
- Glimepiride; Pioglitazone
- Glimepiride; Rosiglitazone
- Glipizide
- Glipizide; Metformin
- Glyburide
- Glyburide; Metformin
- Glycopyrrolate; Formoterol
- Goserelin
- Granisetron
- Halogenated Anesthetics
- Haloperidol
- Halothane
- Histrelin
- Hydroxyzine
- Ibutilide
- Iloperidone
- Imipramine
- Incretin Mimetics
- Indacaterol
- Indacaterol; Glycopyrrolate
- Inotuzumab Ozogamicin
- Insulin Aspart
- Insulin Aspart; Insulin Aspart Protamine
- Insulin Degludec
- Insulin Degludec; Liraglutide
- Insulin Detemir
- Insulin Glargine
- Insulin Glargine; Lixisenatide
- Insulin Glulisine
- Insulin Lispro
- Insulin Lispro; Insulin Lispro Protamine
- Insulin, Inhaled
- Insulins
- Isoflurane
- Isophane Insulin (NPH)
- Itraconazole
- Ivosidenib
- Ketoconazole
- Lanthanum Carbonate
- Lapatinib
- Lefamulin
- Lente Insulin
- Lenvatinib
- Leuprolide
- Leuprolide; Norethindrone
- Levalbuterol
- Levofloxacin
- Linagliptin
- Linagliptin; Metformin

- Liraglutide
- Lithium
- Lixisenatide
- Lofexidine
- Lomefloxacin
- Long-acting beta-agonists
- Loperamide
- Loperamide; Simethicone
- Lopinavir; Ritonavir
- Macimorelin
- Magnesium Hydroxide
- Maprotiline
- Mefloquine
- Meglitinides
- Meperidine; Promethazine
- Metaproterenol
- Metformin
- Metformin; Pioglitazone
- Metformin; Repaglinide
- Metformin; Rosiglitazone
- Metformin; Saxagliptin
- Metformin; Sitagliptin
- Methadone
- Metronidazole
- Midostaurin
- Mifepristone
- Miglitol
- Mirtazapine
- Moxifloxacin
- Nateglinide
- Nilotinib
- Norfloxacin
- Nortriptyline
- Octreotide
- Ofloxacin
- Olanzapine
- Olodaterol
- Omeprazole; Sodium Bicarbonate
- Ondansetron
- Osilodrostat
- Osimertinib
- Oxaliplatin
- Paliperidone
- Panobinostat
- Pasireotide
- Pazopanib
- Penicillamine
- Pentamidine
- Perphenazine
- Perphenazine; Amitriptyline
- Phenylephrine; Promethazine
- Pimavanserin
- Pioglitazone
- Pirbuterol
- Pitolisant
- Posaconazole
- Pramlintide
- Primaquine
- Procainamide
- Prochlorperazine
- Promethazine
- Propafenone
- Protriptyline
- Quetiapine
- Quinidine
- Quinine
- Rabies Vaccine
- Ranolazine
- Regular Insulin
- Regular Insulin; Isophane Insulin (NPH)
- Repaglinide
- Ribociclib
- Ribociclib; Letrozole
- Rilpivirine
- Risperidone
- Romidepsin
- Rosiglitazone
- Salmeterol
- Saquinavir
- Saxagliptin
- Semaglutide
- Sertraline
- Sevoflurane
- SGLT2 Inhibitors
- Short-acting beta-agonists
- Simvastatin; Sitagliptin
- Siponimod
- Sitagliptin
- Sodium Bicarbonate
- Solifenacin
- Sorafenib
- Sotalol
- Sulfonylureas
- Sunitinib
- Tacrolimus
- Tamoxifen
- Telavancin
- Telithromycin
- Terbutaline
- Tetrabenazine
- Thiazolidinediones
- Tiotropium; Olodaterol
- Tolazamide
- Tolbutamide
- Tolterodine
- Toremfene

- Trazodone
- Tricyclic antidepressants
- Trifluoperazine
- Trimipramine
- Triptorelin
- Ultralente Insulin
- Umeclidinium; Vilanterol
- Vandetanib
- Vardenafil
- Vemurafenib
- Venlafaxine
- Vigabatrin
- Voriconazole
- Vorinostat
- Ziprasidone

Level 3 (Moderate)

- Ampicillin
- Articaine; Epinephrine
- Atropine; Hyoscyamine; Phenobarbital; Scopolamine
- Belladonna Alkaloids; Ergotamine; Phenobarbital
- Betrixaban
- Bupivacaine
- Bupivacaine Liposomal
- Bupivacaine; Lidocaine
- Carbamazepine
- Chlorprocaine
- Cobimetinib
- Dabigatran
- Dapsone
- Digoxin
- Edoxaban
- Interferon Alfa-2a
- Interferon Alfa-2b
- Interferon Alfa-2b; Ribavirin
- Interferon Alfa-n3
- Interferon Alfacon-1
- Interferon Beta-1a
- Interferon Beta-1b
- Interferon Gamma-1b
- Interferons
- Lidocaine
- Mepivacaine
- Mepivacaine; Levonordefrin
- Peginterferon Alfa-2a
- Peginterferon Alfa-2b
- Peginterferon beta-1a
- Penicillin G Benzathine; Penicillin G Procaine
- Penicillin G Procaine
- Phenobarbital
- Phenytoin
- Ponatinib
- Prilocaine
- Prilocaine; Epinephrine
- Ropivacaine
- Succinylcholine
- Tetracaine
- Trametinib

Level 4 (Minor)

- Praziquantel
- Telbivudine

Acarbose: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the alpha-glucosidase inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Acetohexamide: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the sulfonylureas, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Acclidinium; Formoterol: (Major) Avoid coadministration of chloroquine with long-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG

at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists (LABAs) as compared to short-acting beta-agonists.. [\[28229\]](#) [\[29758\]](#) [\[41231\]](#) [\[44026\]](#) [\[44063\]](#) [\[44979\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Albiglutide: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the incretin mimetics, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Albuterol: (Major) Avoid coadministration of chloroquine with short-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists as compared to short-acting beta-agonists. [\[28229\]](#) [\[28532\]](#) [\[29758\]](#) [\[49951\]](#) [\[51793\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Albuterol; Ipratropium: (Major) Avoid coadministration of chloroquine with short-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists as compared to short-acting beta-agonists. [\[28229\]](#) [\[28532\]](#) [\[29758\]](#) [\[49951\]](#) [\[51793\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Alfuzosin: (Major) Avoid coadministration of chloroquine with alfuzosin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Alfuzosin may prolong the QT interval in a dose-dependent manner. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28261\]](#) [\[65157\]](#) [\[65170\]](#)

Alogliptin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the dipeptidyl peptidase-4 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Alogliptin; Metformin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including metformin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#) (Major) Careful

monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the dipeptidyl peptidase-4 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Alogliptin; Pioglitazone: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the dipeptidyl peptidase-4 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the thiazolidinediones, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Alpha-glucosidase Inhibitors: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the alpha-glucosidase inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Aluminum Hydroxide: (Major) Chloroquine absorption may be reduced by antacids. Administer chloroquine and antacids at least 4 hours apart. [29758] [30285]

Aluminum Hydroxide; Magnesium Carbonate: (Major) Chloroquine absorption may be reduced by antacids. Administer chloroquine and antacids at least 4 hours apart. [29758] [30285]

Aluminum Hydroxide; Magnesium Hydroxide: (Major) Chloroquine absorption may be reduced by antacids. Administer chloroquine and antacids at least 4 hours apart. [29758] [30285]

Aluminum Hydroxide; Magnesium Hydroxide; Simethicone: (Major) Chloroquine absorption may be reduced by antacids. Administer chloroquine and antacids at least 4 hours apart. [29758] [30285]

Aluminum Hydroxide; Magnesium Trisilicate: (Major) Chloroquine absorption may be reduced by antacids. Administer chloroquine and antacids at least 4 hours apart. [29758] [30285]

Amiodarone: (Major) Avoid coadministration of chloroquine with amiodarone due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Amiodarone, a Class III antiarrhythmic agent, is associated with a well-established risk of QT prolongation and torsade de pointes (TdP). Although the frequency of TdP is less with amiodarone than with other Class III agents, amiodarone is still associated with a risk of TdP. Due to the extremely long half-life of amiodarone, a drug interaction is possible for days to weeks after discontinuation of amiodarone. [28224] [28229] [28230] [28231] [28432] [28457] [65157] [65170]

Amitriptyline: (Major) Avoid coadministration of chloroquine with tricyclic antidepressants (TCAs) due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. TCAs share pharmacologic properties similar to the Class IA antiarrhythmic agents and may prolong the QT interval, particularly in

overdose or with higher-dose prescription therapy (elevated serum concentrations). [\[28225\]](#) [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28415\]](#) [\[28416\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Amitriptyline; Chlordiazepoxide: (Major) Avoid coadministration of chloroquine with tricyclic antidepressants (TCAs) due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. TCAs share pharmacologic properties similar to the Class IA antiarrhythmic agents and may prolong the QT interval, particularly in overdose or with higher-dose prescription therapy (elevated serum concentrations). [\[28225\]](#) [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28415\]](#) [\[28416\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Amoxicillin; Clarithromycin; Lansoprazole: (Major) Avoid coadministration of chloroquine with clarithromycin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Clarithromycin is associated with an established risk for QT prolongation and TdP. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28238\]](#) [\[29758\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Amoxicillin; Clarithromycin; Omeprazole: (Major) Avoid coadministration of chloroquine with clarithromycin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Clarithromycin is associated with an established risk for QT prolongation and TdP. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28238\]](#) [\[29758\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Ampicillin: (Moderate) Administer oral ampicillin 2 hours before or 2 hours after chloroquine. In a study of healthy volunteers, chloroquine significantly reduced the bioavailability of ampicillin. The reduction of ampicillin bioavailability could be attributed to slower gastric emptying and enhancement of gut motility produced by chloroquine. [\[29758\]](#) [\[61761\]](#)

Anagrelide: (Major) Avoid coadministration of chloroquine with anagrelide due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. TdP and ventricular tachycardia have been reported with anagrelide. In addition, dose-related increases in mean QTc and heart rate were observed in healthy subjects. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[30163\]](#) [\[65157\]](#) [\[65170\]](#)

Antacids: (Major) Chloroquine absorption may be reduced by antacids. Administer chloroquine and antacids at least 4 hours apart. [\[29758\]](#) [\[30285\]](#)

Apomorphine: (Major) Avoid coadministration of chloroquine with apomorphine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Limited data indicate that QT prolongation is possible

with apomorphine administration; the change in QTc interval is not significant in most patients receiving dosages within the manufacturer's guidelines. [28229] [28230] [28231] [28661] [29758] [65157] [65170]

Arformoterol: (Major) Avoid coadministration of chloroquine with long-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists (LABAs) as compared to short-acting beta-agonists.. [28229] [29758] [41231] [44026] [44063] [44979] [59321] [65157] [65170]

Aripiprazole: (Major) Avoid coadministration of chloroquine with aripiprazole due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. QT prolongation has occurred during therapeutic use of aripiprazole and following overdose. [28229] [28230] [28231] [29758] [42845] [53394] [60196] [65157] [65170]

Arsenic Trioxide: (Major) Avoid coadministration of chloroquine with arsenic trioxide due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. TdP, QT interval prolongation, and complete atrioventricular block have been reported with arsenic trioxide use. [28226] [28229] [28230] [28231] [65157] [65170]

Artemether; Lumefantrine: (Major) Avoid coadministration of chloroquine with artemether; lumefantrine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Artemether; lumefantrine is associated with prolongation of the QT interval. [28229] [28230] [28231] [29758] [35401] [65157] [65170]

Articaine; Epinephrine: (Moderate) Coadministration of articaine with oxidizing agents, such as chloroquine, may increase the risk of developing methemoglobinemia. Monitor patients closely for signs and symptoms of methemoglobinemia if coadministration is necessary. If methemoglobinemia occurs or is suspected, discontinue articaine and any other oxidizing agents. Depending on the severity of symptoms, patients may respond to supportive care; more severe symptoms may require treatment with methylene blue, exchange transfusion, or hyperbaric oxygen. [28996]

Asenapine: (Major) Avoid coadministration of chloroquine with asenapine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT

prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Asenapine has also been associated with QT prolongation. [28229] [28230] [28231] [29758] [36343] [65157] [65170]

Aspirin, ASA; Citric Acid; Sodium Bicarbonate: (Major) Chloroquine absorption may be reduced by antacids. Administer chloroquine and antacids at least 4 hours apart. [29758] [30284] [30285]

Atomoxetine: (Major) Avoid coadministration of chloroquine with atomoxetine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. QT prolongation has occurred during therapeutic use of atomoxetine and following overdose. [28229] [28230] [28231] [28405] [29758] [59321] [65157] [65170]

Atropine; Hyoscyamine; Phenobarbital; Scopolamine: (Moderate) Coadministration of chloroquine and phenobarbital may decrease exposure of chloroquine which may reduce its efficacy. Chloroquine may be an in vitro CYP3A4 substrate and phenobarbital is a strong CYP3A4 inducer. [65210] [65237] [65239]

Azithromycin: (Major) Avoid coadministration of chloroquine with azithromycin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances (See Therapeutic Drug Monitoring for recommendations specific to COVID-19). Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. QT prolongation and torsade de pointes (TdP) have been spontaneously reported during azithromycin postmarketing surveillance. [28229] [28230] [28231] [28855] [29758] [43974] [65157] [65170]

Bedaquiline: (Major) Avoid coadministration of chloroquine with bedaquiline due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Discontinue bedaquiline if evidence of serious ventricular arrhythmia or QTcF interval greater than 500 msec. Bedaquiline prolongs the QT interval. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. [28229] [28230] [28231] [29758] [52746] [65157] [65170]

Belladonna Alkaloids; Ergotamine; Phenobarbital: (Moderate) Coadministration of chloroquine and phenobarbital may decrease exposure of chloroquine which may reduce its efficacy. Chloroquine may be an in vitro CYP3A4 substrate and phenobarbital is a strong CYP3A4 inducer. [65210] [65237] [65239]

Betrixaban: (Moderate) Use caution if chloroquine is coadministered with betrixaban due to the potential for increased betrixaban exposure which may increase the risk of bleeding. Betrixaban is a P-gp substrate; limited data suggests that chloroquine is a P-gp inhibitor. [62037] [65210]

Bismuth Subcitrate Potassium; Metronidazole; Tetracycline: (Major) Avoid coadministration of chloroquine with metronidazole due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de

pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Potential QT prolongation has been reported in limited case reports with metronidazole. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[57377\]](#) [\[57378\]](#) [\[65157\]](#) [\[65170\]](#)

Bismuth Subsalicylate; Metronidazole; Tetracycline: (Major) Avoid coadministration of chloroquine with metronidazole due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Potential QT prolongation has been reported in limited case reports with metronidazole. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[57377\]](#) [\[57378\]](#) [\[65157\]](#) [\[65170\]](#)

Budesonide; Formoterol: (Major) Avoid coadministration of chloroquine with long-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists (LABAs) as compared to short-acting beta-agonists.. [\[28229\]](#) [\[29758\]](#) [\[41231\]](#) [\[44026\]](#) [\[44063\]](#) [\[44979\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Bupivacaine Liposomal: (Moderate) Coadministration of bupivacaine with oxidizing agents, such as chloroquine, may increase the risk of developing methemoglobinemia. Monitor patients closely for signs and symptoms of methemoglobinemia if coadministration is necessary. If methemoglobinemia occurs or is suspected, discontinue bupivacaine and any other oxidizing agents. Depending on the severity of symptoms, patients may respond to supportive care; more severe symptoms may require treatment with methylene blue, exchange transfusion, or hyperbaric oxygen. [\[52331\]](#)

Bupivacaine: (Moderate) Coadministration of bupivacaine with oxidizing agents, such as chloroquine, may increase the risk of developing methemoglobinemia. Monitor patients closely for signs and symptoms of methemoglobinemia if coadministration is necessary. If methemoglobinemia occurs or is suspected, discontinue bupivacaine and any other oxidizing agents. Depending on the severity of symptoms, patients may respond to supportive care; more severe symptoms may require treatment with methylene blue, exchange transfusion, or hyperbaric oxygen. [\[52331\]](#)

Bupivacaine; Lidocaine: (Moderate) Coadministration of bupivacaine with oxidizing agents, such as chloroquine, may increase the risk of developing methemoglobinemia. Monitor patients closely for signs and symptoms of methemoglobinemia if coadministration is necessary. If methemoglobinemia occurs or is suspected, discontinue bupivacaine and any other oxidizing agents. Depending on the severity of symptoms, patients may respond to supportive care; more severe symptoms may require treatment with methylene blue, exchange transfusion, or hyperbaric oxygen. [\[52331\]](#) (Moderate) Coadministration of lidocaine with oxidizing agents, such as chloroquine, may increase the risk of developing methemoglobinemia. Monitor patients closely for signs and symptoms of methemoglobinemia if coadministration is necessary. If methemoglobinemia occurs or is suspected, discontinue lidocaine and any other oxidizing agents. Depending on the severity of symptoms, patients may respond to supportive care; more severe symptoms may require treatment with methylene blue, exchange transfusion, or hyperbaric oxygen. [\[43383\]](#)

Buprenorphine: (Major) Avoid coadministration of chloroquine with buprenorphine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Buprenorphine has been associated with QT prolongation and has a possible risk of TdP. [28229] [28230] [28231] [29758] [41235] [59321] [60270] [65157] [65170]

Buprenorphine; Naloxone: (Major) Avoid coadministration of chloroquine with buprenorphine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Buprenorphine has been associated with QT prolongation and has a possible risk of TdP. [28229] [28230] [28231] [29758] [41235] [59321] [60270] [65157] [65170]

Calcium Carbonate: (Major) Chloroquine absorption may be reduced by antacids. Administer chloroquine and antacids at least 4 hours apart. [29758] [30284] [30285]

Calcium Carbonate; Magnesium Hydroxide: (Major) Chloroquine absorption may be reduced by antacids. Administer chloroquine and antacids at least 4 hours apart. [29758] [30284] [30285]
(Major) Chloroquine absorption may be reduced by antacids. Administer chloroquine and antacids at least 4 hours apart. [29758] [30285]

Calcium Carbonate; Risedronate: (Major) Chloroquine absorption may be reduced by antacids. Administer chloroquine and antacids at least 4 hours apart. [29758] [30284] [30285]

Calcium Carbonate; Simethicone: (Major) Chloroquine absorption may be reduced by antacids. Administer chloroquine and antacids at least 4 hours apart. [29758] [30284] [30285]

Canagliflozin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the SGLT2 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Canagliflozin; Metformin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including metformin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the SGLT2 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Carbamazepine: (Moderate) Coadministration of chloroquine and carbamazepine may decrease exposure of chloroquine which may reduce its efficacy. Chloroquine may be an in vitro CYP3A4 substrate and carbamazepine is a strong CYP3A4 inducer. [65210] [65237] [65239]

Ceritinib: (Major) Avoid coadministration of chloroquine with ceritinib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Interrupt, dose reduce, or discontinue

ceritinib if QT prolongation occurs. Ceritinib causes concentration-dependent QT prolongation. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. [28229] [28230] [28231] [29758] [34353] [57094] [65157] [65170]

Chlorprocaine: (Moderate) Coadministration of chlorprocaine with oxidizing agents, such as chloroquine, may increase the risk of developing methemoglobinemia. Monitor patients closely for signs and symptoms of methemoglobinemia if coadministration is necessary. If methemoglobinemia occurs or is suspected, discontinue chlorprocaine and any other oxidizing agents. Depending on the severity of symptoms, patients may respond to supportive care; more severe symptoms may require treatment with methylene blue, exchange transfusion, or hyperbaric oxygen. [29062]

Chlorpromazine: (Major) Avoid coadministration of chloroquine with chlorpromazine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Chlorpromazine is associated with an established risk of QT prolongation and TdP. [28229] [28230] [28231] [29758] [43065] [59321] [65157] [65170]

Chlorpropamide: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the sulfonylureas, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Cimetidine: (Major) Avoid concomitant use of chloroquine and cimetidine as cimetidine may inhibit the metabolism of chloroquine, increasing its plasma concentration. [29758] [34335] [34353] [61759] [61760]

Ciprofloxacin: (Major) Avoid coadministration of chloroquine with ciprofloxacin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Rare cases of QT prolongation and TdP have been reported with ciprofloxacin during postmarketing surveillance. [28229] [28230] [28231] [29758] [43411] [46655] [65157] [65170]

Cisapride: (Severe) Coadministration of chloroquine with cisapride is contraindicated due to the increased risk of QT prolongation. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. QT prolongation and ventricular arrhythmias, including TdP and death, have been reported with cisapride. [28229] [28230] [28231] [29758] [47221] [65157] [65170]

Citalopram: (Major) Avoid coadministration of chloroquine with citalopram due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Citalopram causes dose-dependent QT interval prolongation. [28229] [28230] [28231] [28269] [29758] [65157] [65170]

Clarithromycin: (Major) Avoid coadministration of chloroquine with clarithromycin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Clarithromycin is associated with an established risk for QT prolongation and TdP. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28238\]](#) [\[29758\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Clofazimine: (Major) Avoid coadministration of chloroquine with clofazimine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. QT prolongation and TdP have been reported in patients receiving clofazimine in combination with QT prolonging medications. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[63936\]](#) [\[65157\]](#) [\[65170\]](#)

Clomipramine: (Major) Avoid coadministration of chloroquine with tricyclic antidepressants (TCAs) due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. TCAs share pharmacologic properties similar to the Class IA antiarrhythmic agents and may prolong the QT interval, particularly in overdose or with higher-dose prescription therapy (elevated serum concentrations). [\[28225\]](#) [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28415\]](#) [\[28416\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Clozapine: (Major) Avoid coadministration of chloroquine with clozapine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Treatment with clozapine has been associated with QT prolongation, TdP, cardiac arrest, and sudden death. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28262\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Cobimetinib: (Moderate) Concurrent use of chloroquine and cobimetinib is not recommended as there is an increased risk of retinal toxicity. [\[29758\]](#) [\[60281\]](#)

Codeine; Phenylephrine; Promethazine: (Major) Avoid coadministration of chloroquine with promethazine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Promethazine is associated with a possible risk for QT prolongation. [\[28225\]](#) [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[55578\]](#) [\[65157\]](#) [\[65170\]](#)

Codeine; Promethazine: (Major) Avoid coadministration of chloroquine with promethazine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is

increased with higher chloroquine doses. Promethazine is associated with a possible risk for QT prolongation. [\[28225\]](#) [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[55578\]](#) [\[65157\]](#) [\[65170\]](#)

Crizotinib: (Major) Avoid coadministration of chloroquine with crizotinib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. An interruption of therapy, dose reduction, or discontinuation of therapy may be necessary for crizotinib if QT prolongation occurs. Crizotinib can cause concentration-dependent QT prolongation. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. [\[28229\]](#) [\[28231\]](#) [\[29758\]](#) [\[45458\]](#) [\[65157\]](#) [\[65170\]](#)

Cyclosporine: (Major) Close monitoring of serum cyclosporine concentrations is recommended during coadministration of chloroquine. Sudden increases in cyclosporine concentrations have been reported after the addition of chloroquine. Discontinue chloroquine if necessary. [\[29758\]](#)

Dabigatran: (Moderate) Use caution if chloroquine is coadministered with dabigatran due to the potential for increased dabigatran exposure which may increase the risk of bleeding. Dabigatran is a P-gp substrate; limited data suggests that chloroquine is a P-gp inhibitor. [\[42121\]](#) [\[65210\]](#)

Dapagliflozin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the SGLT2 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Dapagliflozin; Metformin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including metformin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#) (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the SGLT2 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Dapagliflozin; Saxagliptin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the dipeptidyl peptidase-4 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#) (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the SGLT2 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Dapsone: (Moderate) Coadministration of dapsone with chloroquine may increase the risk of developing methemoglobinemia. Advise patients to discontinue treatment and seek immediate medical attention with any signs or symptoms of methemoglobinemia. [\[60612\]](#)

Dasatinib: (Major) Avoid coadministration of chloroquine with dasatinib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. In vitro studies have shown that dasatinib has the potential to prolong the QT interval. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[32387\]](#) [\[65157\]](#) [\[65170\]](#)

Degarelix: (Major) Avoid coadministration of chloroquine with degarelix due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Androgen deprivation therapy (i.e., degarelix) may also prolong the QT/QTc interval. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[46869\]](#) [\[65157\]](#) [\[65170\]](#)

Desflurane: (Major) Avoid coadministration of chloroquine with halogenated anesthetics due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Halogenated anesthetics are known to prolong the QT interval. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28457\]](#) [\[28458\]](#) [\[28754\]](#) [\[28755\]](#) [\[28756\]](#) [\[29758\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Desipramine: (Major) Avoid coadministration of chloroquine with tricyclic antidepressants (TCAs) due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. TCAs share pharmacologic properties similar to the Class IA antiarrhythmic agents and may prolong the QT interval, particularly in overdose or with higher-dose prescription therapy (elevated serum concentrations). [\[28225\]](#) [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28415\]](#) [\[28416\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Deutetrabenazine: (Major) Avoid coadministration of chloroquine with deutetrabenazine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. For patients taking a deutetrabenazine dosage more than 24 mg/day, assess the QTc interval before and after increasing the deutetrabenazine dosage or other medications that prolong the QTc interval. Clinically relevant QTc prolongation may occur with deutetrabenazine. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[61845\]](#) [\[65157\]](#) [\[65170\]](#)

Dextromethorphan; Promethazine: (Major) Avoid coadministration of chloroquine with promethazine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Promethazine is associated with a possible risk for QT prolongation. [\[28225\]](#) [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[55578\]](#) [\[65157\]](#) [\[65170\]](#)

Dextromethorphan; Quinidine: (Major) Avoid coadministration of chloroquine with quinidine due to the increased risk of QT prolongation or other drug toxicities. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Quinidine

administration is also associated with QT prolongation and TdP. [28229] [28230] [28231] [29758] [42280] [47357] [65157] [65170]

Digoxin: (Moderate) Monitor serum digoxin concentrations in patients receiving digoxin and chloroquine as coadministration may result in increased serum digoxin concentrations. Digoxin serum concentrations have been reported to increase when hydroxychloroquine was added. Although this interaction has not been reported with chloroquine, chloroquine may similarly increase the plasma concentration of digoxin. [41806] [60957]

Dipeptidyl Peptidase-4 Inhibitors: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the dipeptidyl peptidase-4 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Disopyramide: (Major) Avoid coadministration of chloroquine with disopyramide due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Disopyramide administration is associated with QT prolongation and TdP. [28228] [28229] [28230] [28231] [29758] [65157] [65170]

Dofetilide: (Major) Avoid coadministration of chloroquine with dofetilide due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Dofetilide, a Class III antiarrhythmic agent, is associated with a well-established risk of QT prolongation and TdP. [28221] [28229] [28230] [28231] [28432] [28457] [29758] [65157] [65170]

Dolasetron: (Major) Avoid coadministration of chloroquine with dolasetron due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Dolasetron has been associated with a dose-dependent prolongation of the QT, PR, and QRS intervals on an electrocardiogram. [28229] [28230] [28231] [29758] [34353] [42844] [55935] [65157] [65170]

Dolutegravir; Rilpivirine: (Major) Avoid coadministration of chloroquine with rilpivirine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Supratherapeutic doses of rilpivirine (75 to 300 mg/day) have caused QT prolongation. [28229] [28230] [28231] [29758] [44376] [65157] [65170]

Donepezil: (Major) Avoid coadministration of chloroquine with donepezil due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased

with higher chloroquine doses. Case reports indicate that QT prolongation and TdP can occur during donepezil therapy. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[59321\]](#) [\[59322\]](#) [\[65157\]](#) [\[65170\]](#)

Donepezil; Memantine: (Major) Avoid coadministration of chloroquine with donepezil due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Case reports indicate that QT prolongation and TdP can occur during donepezil therapy. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[59321\]](#) [\[59322\]](#) [\[65157\]](#) [\[65170\]](#)

Doxepin: (Major) Avoid coadministration of chloroquine with tricyclic antidepressants (TCAs) due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. TCAs share pharmacologic properties similar to the Class IA antiarrhythmic agents and may prolong the QT interval, particularly in overdose or with higher-dose prescription therapy (elevated serum concentrations). [\[28225\]](#) [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28415\]](#) [\[28416\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Dronedaron: (Severe) Coadministration of chloroquine with dronedarone is contraindicated due to the increased risk of QT prolongation. Dronedaron administration is associated with a dose-related increase in the QTc interval. The increase in QTc is approximately 10 milliseconds at doses of 400 mg twice daily (the FDA-approved dose) and up to 25 milliseconds at doses of 1,600 mg twice daily. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[36101\]](#) [\[65157\]](#) [\[65170\]](#)

Droperidol: (Major) Avoid coadministration of chloroquine with droperidol due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Initiate droperidol at a low dose and increase the dose as needed to achieve the desired effect. Droperidol administration is associated with an established risk for QT prolongation and torsade de pointes (TdP). Some cases have occurred in patients with no known risk factors for QT prolongation and some cases have been fatal. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. [\[28229\]](#) [\[28231\]](#) [\[28235\]](#) [\[28236\]](#) [\[28237\]](#) [\[28737\]](#) [\[29758\]](#) [\[51289\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Dulaglutide: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the incretin mimetics, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Edoxaban: (Moderate) Use caution if chloroquine is coadministered with edoxaban due to the potential for increased edoxaban exposure which may increase the risk of bleeding. Edoxaban is a P-gp substrate; limited data suggests that chloroquine is a P-gp inhibitor. [\[58685\]](#) [\[65210\]](#)

Efavirenz: (Major) Avoid coadministration of chloroquine with efavirenz due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased

with higher chloroquine doses. QTc prolongation has been observed with the use of efavirenz. [28229] [28230] [28231] [28442] [29758] [65157] [65170]

Efavirenz; Emtricitabine; Tenofovir: (Major) Avoid coadministration of chloroquine with efavirenz due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. QTc prolongation has been observed with the use of efavirenz. [28229] [28230] [28231] [28442] [29758] [65157] [65170]

Efavirenz; Lamivudine; Tenofovir Disoproxil Fumarate: (Major) Avoid coadministration of chloroquine with efavirenz due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. QTc prolongation has been observed with the use of efavirenz. [28229] [28230] [28231] [28442] [29758] [65157] [65170]

Eliglustat: (Major) Avoid coadministration of chloroquine with eliglustat due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Eliglustat is predicted to cause PR, QRS, and/or QT prolongation at significantly elevated plasma concentrations. [28229] [28230] [28231] [29758] [57803] [65157] [65170]

Empagliflozin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the SGLT2 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Empagliflozin; Linagliptin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the dipeptidyl peptidase-4 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the SGLT2 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Empagliflozin; Linagliptin; Metformin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including metformin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the dipeptidyl peptidase-4 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the SGLT2 inhibitors, are coadministered. A

decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Empagliflozin; Metformin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including metformin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the SGLT2 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Emtricitabine; Rilpivirine; Tenofovir alafenamide: (Major) Avoid coadministration of chloroquine with rilpivirine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Supratherapeutic doses of rilpivirine (75 to 300 mg/day) have caused QT prolongation. [28229] [28230] [28231] [29758] [44376] [65157] [65170]

Emtricitabine; Rilpivirine; Tenofovir disoproxil fumarate: (Major) Avoid coadministration of chloroquine with rilpivirine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Supratherapeutic doses of rilpivirine (75 to 300 mg/day) have caused QT prolongation. [28229] [28230] [28231] [29758] [44376] [65157] [65170]

Encorafenib: (Major) Avoid coadministration of chloroquine with encorafenib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Encorafenib has been associated with dose-dependent QT prolongation. [28229] [28230] [28231] [29758] [63317] [65157] [65170]

Enflurane: (Major) Avoid coadministration of chloroquine with halogenated anesthetics due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Halogenated anesthetics are known to prolong the QT interval. [28229] [28230] [28231] [28457] [28458] [28754] [28755] [28756] [29758] [59321] [65157] [65170]

Entrectinib: (Major) Avoid coadministration of chloroquine with entrectinib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased

with higher chloroquine doses. Entrectinib has been associated with QT prolongation. [28229] [28230] [28231] [29758] [64567] [65157] [65170]

Eribulin: (Major) Avoid coadministration of chloroquine with eribulin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Eribulin has been associated with QT prolongation. [28229] [28230] [28231] [29758] [42449] [65157] [65170]

Ertugliflozin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the SGLT2 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Ertugliflozin; Metformin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including metformin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the SGLT2 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Ertugliflozin; Sitagliptin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the dipeptidyl peptidase-4 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the SGLT2 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Erythromycin: (Major) Avoid coadministration of chloroquine with erythromycin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Erythromycin is associated with QT prolongation and TdP. [28229] [28230] [28231] [29758] [43258] [59321] [65157] [65170]

Erythromycin; Sulfisoxazole: (Major) Avoid coadministration of chloroquine with erythromycin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Erythromycin is associated with QT prolongation and TdP. [28229] [28230] [28231] [29758] [43258] [59321] [65157] [65170]

Escitalopram: (Major) Avoid coadministration of chloroquine with escitalopram due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with

an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Escitalopram has been associated with a risk of QT prolongation and TdP. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28270\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Exenatide: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the incretin mimetics, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Ezogabine: (Major) Avoid coadministration of chloroquine with ezogabine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Ezogabine has been associated with QT prolongation. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[44800\]](#) [\[65157\]](#) [\[65170\]](#)

Fingolimod: (Major) Avoid coadministration of chloroquine with fingolimod due to the increased risk of QT prolongation. If use together is necessary, overnight monitoring with continuous ECG in a medical facility is advised after the first dose of fingolimod; monitor ECG closely throughout therapy, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Fingolimod initiation results in decreased heart rate and may prolong the QT interval. Fingolimod has not been studied in patients treated with drugs that prolong the QT interval, but drugs that prolong the QT interval have been associated with cases of TdP in patients with bradycardia. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Flecainide: (Major) Avoid coadministration of chloroquine with flecainide due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Flecainide is a Class IC antiarrhythmic associated with a possible risk for QT prolongation and/or TdP; flecainide increases the QT interval, but largely due to prolongation of the QRS interval. Although causality for TdP has not been established for flecainide, patients receiving concurrent drugs that have the potential for QT prolongation may have an increased risk of developing proarrhythmias. [\[23774\]](#) [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28752\]](#) [\[29758\]](#) [\[51070\]](#) [\[65157\]](#) [\[65170\]](#)

Fluconazole: (Major) Avoid coadministration of chloroquine with fluconazole due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Fluconazole has been associated with QT prolongation and rare cases of TdP. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28674\]](#) [\[29758\]](#) [\[60686\]](#) [\[65157\]](#) [\[65170\]](#)

Fluoxetine: (Major) Avoid coadministration of chloroquine with fluoxetine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased

with higher chloroquine doses. QT prolongation and TdP have been reported in patients treated with fluoxetine. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[32127\]](#) [\[44058\]](#) [\[44059\]](#) [\[65157\]](#) [\[65170\]](#)

Fluoxetine; Olanzapine: (Major) Avoid coadministration of chloroquine with fluoxetine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. QT prolongation and TdP have been reported in patients treated with fluoxetine. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[32127\]](#) [\[44058\]](#) [\[44059\]](#) [\[65157\]](#) [\[65170\]](#)

(Major) Avoid coadministration of chloroquine with olanzapine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Limited data, including some case reports, suggest that olanzapine may be associated with a significant prolongation of the QTc interval. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28785\]](#) [\[29758\]](#) [\[32732\]](#) [\[32734\]](#) [\[32745\]](#) [\[65157\]](#) [\[65170\]](#)

Fluphenazine: (Major) Avoid coadministration of chloroquine with fluphenazine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Fluphenazine is associated with a possible risk for QT prolongation. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28415\]](#) [\[29758\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Fluticasone; Salmeterol: (Major) Avoid coadministration of chloroquine with long-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists (LABAs) as compared to short-acting beta-agonists.. [\[28229\]](#) [\[29758\]](#) [\[41231\]](#) [\[44026\]](#) [\[44063\]](#) [\[44979\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Fluticasone; Umeclidinium; Vilanterol: (Major) Avoid coadministration of chloroquine with long-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists (LABAs) as compared to short-acting beta-agonists.. [\[28229\]](#) [\[29758\]](#) [\[41231\]](#) [\[44026\]](#) [\[44063\]](#) [\[44979\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Fluticasone; Vilanterol: (Major) Avoid coadministration of chloroquine with long-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring,

avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists (LABAs) as compared to short-acting beta-agonists.. [\[28229\]](#) [\[29758\]](#) [\[41231\]](#) [\[44026\]](#) [\[44063\]](#) [\[44979\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Fluvoxamine: (Major) Avoid coadministration of chloroquine with fluvoxamine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. QT prolongation and TdP has been reported during fluvoxamine postmarketing use. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[50507\]](#) [\[65157\]](#) [\[65170\]](#)

Formoterol: (Major) Avoid coadministration of chloroquine with long-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists (LABAs) as compared to short-acting beta-agonists.. [\[28229\]](#) [\[29758\]](#) [\[41231\]](#) [\[44026\]](#) [\[44063\]](#) [\[44979\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Formoterol; Mometasone: (Major) Avoid coadministration of chloroquine with long-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists (LABAs) as compared to short-acting beta-agonists.. [\[28229\]](#) [\[29758\]](#) [\[41231\]](#) [\[44026\]](#) [\[44063\]](#) [\[44979\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Foscarnet: (Major) Avoid coadministration of chloroquine with foscarnet due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Both QT prolongation and TdP have been reported during postmarketing use of foscarnet. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28377\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Gemifloxacin: (Major) Avoid coadministration of chloroquine with gemifloxacin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Gemifloxacin may prolong the QT interval in some patients. The maximal change in the QTc interval occurs approximately 5 to 10 hours following

oral administration of gemifloxacin. The likelihood of QTc prolongation may increase with increasing gemifloxacin dose; therefore, do not exceed the recommended dose, especially in patients with renal or hepatic impairment where the C_{max} and AUC are slightly higher. [28229] [28230] [28231] [28419] [28420] [28424] [29758] [65157] [65170]

Gemtuzumab Ozogamicin: (Major) Avoid coadministration of chloroquine with gemtuzumab due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Although QT interval prolongation has not been reported with gemtuzumab, it has been reported with other drugs that contain calicheamicin. [28229] [28230] [28231] [29758] [62292] [65157] [65170]

Gilteritinib: (Major) Avoid coadministration of chloroquine with gilteritinib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Gilteritinib has been associated with QT prolongation. [28229] [28230] [28231] [29758] [63787] [65157] [65170]

Glasdegib: (Major) Avoid coadministration of chloroquine with glasdegib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Glasdegib therapy may result in QT prolongation and ventricular arrhythmias including ventricular fibrillation and ventricular tachycardia. [28229] [28230] [28231] [29758] [63777] [65157] [65170]

Glimepiride: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the sulfonylureas, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Glimepiride; Pioglitazone: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the sulfonylureas, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the thiazolidinediones, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Glimepiride; Rosiglitazone: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the sulfonylureas, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the thiazolidinediones, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Glipizide: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the sulfonylureas, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Glipizide; Metformin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including metformin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the sulfonylureas, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Glyburide: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the sulfonylureas, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Glyburide; Metformin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including metformin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the sulfonylureas, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Glycopyrrolate; Formoterol: (Major) Avoid coadministration of chloroquine with long-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists (LABAs) as compared to short-acting beta-agonists.. [28229] [29758] [41231] [44026] [44063] [44979] [59321] [65157] [65170]

Goserelin: (Major) Avoid coadministration of chloroquine with goserelin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Androgen deprivation therapy (i.e., goserelin) may also prolong the QT/QTc interval. [28229] [28230] [28231] [28592] [29758] [65157] [65170]

Granisetron: (Major) Avoid coadministration of chloroquine with granisetron due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased

with higher chloroquine doses. Granisetron has been associated with QT prolongation. [28229] [28230] [28231] [29758] [31723] [65157] [65170]

Halogenated Anesthetics: (Major) Avoid coadministration of chloroquine with halogenated anesthetics due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Halogenated anesthetics are known to prolong the QT interval. [28229] [28230] [28231] [28457] [28458] [28754] [28755] [28756] [29758] [59321] [65157] [65170]

Haloperidol: (Major) Avoid coadministration of chloroquine with haloperidol due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. QT prolongation and TdP have been observed during haloperidol treatment. Excessive doses (particularly in the overdose setting) or IV administration of haloperidol may be associated with a higher risk of QT prolongation. [23500] [23779] [28229] [28230] [28231] [28307] [28415] [29758] [59321] [65157] [65170]

Halothane: (Major) Avoid coadministration of chloroquine with halogenated anesthetics due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Halogenated anesthetics are known to prolong the QT interval. [28229] [28230] [28231] [28457] [28458] [28754] [28755] [28756] [29758] [59321] [65157] [65170]

Histrelin: (Major) Avoid coadministration of chloroquine with histrelin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Androgen deprivation therapy (i.e., histrelin) may prolong the QT/QTc interval. [28229] [28230] [28231] [29758] [30369] [65157] [65170]

Hydroxyzine: (Major) Avoid coadministration of chloroquine with hydroxyzine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Postmarketing data indicate that hydroxyzine causes QT prolongation and TdP. [28229] [28230] [28231] [29758] [47129] [61470] [65157] [65170]

Ibutilide: (Major) Avoid coadministration of chloroquine with ibutilide due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Ibutilide administration can cause QT prolongation and TdP;

proarrhythmic events should be anticipated. The potential for proarrhythmic events with ibutilide increases with the use of other drugs that prolong the QT interval. [28229] [28230] [28231] [29758] [41830] [65157] [65170]

Iloperidone: (Major) Avoid coadministration of chloroquine with iloperidone due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Iloperidone has also been associated with QT prolongation. [28229] [28230] [28231] [29758] [36146] [65157] [65170]

Imipramine: (Major) Avoid coadministration of chloroquine with tricyclic antidepressants (TCAs) due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. TCAs share pharmacologic properties similar to the Class IA antiarrhythmic agents and may prolong the QT interval, particularly in overdose or with higher-dose prescription therapy (elevated serum concentrations). [28225] [28229] [28230] [28231] [28415] [28416] [29758] [65157] [65170]

Incretin Mimetics: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the incretin mimetics, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Indacaterol: (Major) Avoid coadministration of chloroquine with long-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists (LABAs) as compared to short-acting beta-agonists.. [28229] [29758] [41231] [44026] [44063] [44979] [59321] [65157] [65170]

Indacaterol; Glycopyrrolate: (Major) Avoid coadministration of chloroquine with long-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists (LABAs) as compared to short-acting beta-agonists.. [28229] [29758] [41231] [44026] [44063] [44979] [59321] [65157] [65170]

Inotuzumab Ozogamicin: (Major) Avoid coadministration of chloroquine with inotuzumab due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with

an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Inotuzumab has also been associated with QT interval prolongation. [28229] [28230] [28231] [29758] [62245] [65157] [65170]

Insulin Aspart: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including insulin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Insulin Aspart; Insulin Aspart Protamine: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including insulin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Insulin Degludec: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including insulin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Insulin Degludec; Liraglutide: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including insulin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the incretin mimetics, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Insulin Detemir: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including insulin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Insulin Glargine: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including insulin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Insulin Glargine; Lixisenatide: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including insulin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the incretin mimetics, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Insulin Glulisine: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including insulin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Insulin Lispro: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including insulin, are coadministered. A decreased dose of the antidiabetic

agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Insulin Lispro; Insulin Lispro Protamine: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including insulin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Insulin, Inhaled: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including insulin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Insulins: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including insulin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Interferon Alfa-2a: (Moderate) Concurrent use of chloroquine and interferons is not recommended as there is an increased risk of retinal toxicity. [29758] [47391]

Interferon Alfa-2b: (Moderate) Concurrent use of chloroquine and interferons is not recommended as there is an increased risk of retinal toxicity. [29758] [47391]

Interferon Alfa-2b; Ribavirin: (Moderate) Concurrent use of chloroquine and interferons is not recommended as there is an increased risk of retinal toxicity. [29758] [47391]

Interferon Alfacon-1: (Moderate) Concurrent use of chloroquine and interferons is not recommended as there is an increased risk of retinal toxicity. [29758] [47391]

Interferon Alfa-n3: (Moderate) Concurrent use of chloroquine and interferons is not recommended as there is an increased risk of retinal toxicity. [29758] [47391]

Interferon Beta-1a: (Moderate) Concurrent use of chloroquine and interferons is not recommended as there is an increased risk of retinal toxicity. [29758] [47391]

Interferon Beta-1b: (Moderate) Concurrent use of chloroquine and interferons is not recommended as there is an increased risk of retinal toxicity. [29758] [47391]

Interferon Gamma-1b: (Moderate) Concurrent use of chloroquine and interferons is not recommended as there is an increased risk of retinal toxicity. [29758] [47391]

Interferons: (Moderate) Concurrent use of chloroquine and interferons is not recommended as there is an increased risk of retinal toxicity. [29758] [47391]

Isoflurane: (Major) Avoid coadministration of chloroquine with halogenated anesthetics due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Halogenated anesthetics are known to prolong the QT interval. [28229] [28230] [28231] [28457] [28458] [28754] [28755] [28756] [29758] [59321] [65157] [65170]

Isophane Insulin (NPH): (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including insulin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Itraconazole: (Major) Avoid coadministration of chloroquine with itraconazole due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Itraconazole has also been associated with prolongation of the QT interval. [27983] [28229] [28230] [28231] [29758] [40233] [57441] [63821] [65157] [65170]

Ivosidenib: (Major) Avoid coadministration of chloroquine with ivosidenib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. An interruption of therapy and dose reduction of ivosidenib may be necessary if QT prolongation occurs. Prolongation of the QTc interval and ventricular arrhythmias have been reported in patients treated with ivosidenib. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. [28229] [28230] [28231] [29758] [65157] [65170]

Ketoconazole: (Major) Avoid coadministration of chloroquine with systemic ketoconazole due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Ketoconazole has been associated with prolongation of the QT interval. [27982] [28229] [28230] [28231] [29758] [59321] [65157] [65170]

Lanthanum Carbonate: (Major) Oral compounds known to interact with antacids, like chloroquine, should not be taken within 4 hours of dosing with lanthanum carbonate. If these agents are used concomitantly, space the dosing intervals appropriately. Monitor serum concentrations and clinical condition. [29758] [30284] [30285] [44406]

Lapatinib: (Major) Avoid coadministration of chloroquine with lapatinib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Lapatinib has been associated with concentration-dependent QT prolongation; ventricular arrhythmias and TdP have been reported in postmarketing experience with lapatinib. [28229] [28230] [28231] [29758] [33192] [65157] [65170]

Lefamulin: (Major) Avoid coadministration of chloroquine with lefamulin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Lefamulin has a concentration dependent QTc prolongation effect. The potential to prolong the QT interval of the ECG between lefamulin and other drugs that effect cardiac conduction is unknown. [28229] [28230] [28231] [29758] [64576] [65157] [65170]

Lente Insulin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including insulin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Lenvatinib: (Major) Avoid coadministration of chloroquine with lenvatinib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Prolongation of the QT interval has been reported with lenvatinib therapy. [28229] [28230] [28231] [29758] [58782] [65157] [65170]

Leuprolide: (Major) Avoid coadministration of chloroquine with leuprolide due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Androgen deprivation therapy (i.e., leuprolide) may prolong the QT/QTc interval. [28229] [28230] [28231] [29758] [43800] [65157] [65170]

Leuprolide; Norethindrone: (Major) Avoid coadministration of chloroquine with leuprolide due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Androgen deprivation therapy (i.e., leuprolide) may prolong the QT/QTc interval. [28229] [28230] [28231] [29758] [43800] [65157] [65170]

Levalbuterol: (Major) Avoid coadministration of chloroquine with short-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists as compared to short-acting beta-agonists. [28229] [28532] [29758] [49951] [51793] [59321] [65157] [65170]

Levofloxacin: (Major) Avoid coadministration of chloroquine with levofloxacin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Levofloxacin has been associated with prolongation of the QT interval and infrequent cases of arrhythmia. Additionally, rare cases of TdP have been spontaneously reported during postmarketing surveillance in patients receiving levofloxacin. [28229] [28230] [28231] [28421] [29758] [61195] [63729] [65157] [65170]

Lidocaine: (Moderate) Coadministration of lidocaine with oxidizing agents, such as chloroquine, may increase the risk of developing methemoglobinemia. Monitor patients closely for signs and symptoms of methemoglobinemia if coadministration is necessary. If methemoglobinemia occurs

or is suspected, discontinue lidocaine and any other oxidizing agents. Depending on the severity of symptoms, patients may respond to supportive care; more severe symptoms may require treatment with methylene blue, exchange transfusion, or hyperbaric oxygen. [\[43383\]](#)

Linagliptin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the dipeptidyl peptidase-4 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Linagliptin; Metformin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including metformin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#) (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the dipeptidyl peptidase-4 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Liraglutide: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the incretin mimetics, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Lithium: (Major) Avoid coadministration of chloroquine with lithium due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Lithium has also been associated with QT prolongation. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[59809\]](#) [\[59810\]](#) [\[59811\]](#) [\[65157\]](#) [\[65170\]](#)

Lixisenatide: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the incretin mimetics, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Lofexidine: (Major) Avoid coadministration of chloroquine with lofexidine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Lofexidine also prolongs the QT interval. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[63161\]](#) [\[65157\]](#) [\[65170\]](#)

Lomefloxacin: (Major) Avoid coadministration of chloroquine with lomefloxacin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Cases of QT prolongation and TdP have been reported with lomefloxacin. [\[29758\]](#) [\[29948\]](#) [\[59321\]](#)

Long-acting beta-agonists: (Major) Avoid coadministration of chloroquine with long-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG

at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists (LABAs) as compared to short-acting beta-agonists.. [\[28229\]](#) [\[29758\]](#) [\[41231\]](#) [\[44026\]](#) [\[44063\]](#) [\[44979\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Loperamide: (Major) Avoid coadministration of chloroquine with loperamide due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. At high doses, loperamide has been associated with serious cardiac toxicities, including syncope, ventricular tachycardia, QT prolongation, TdP, and cardiac arrest. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[30106\]](#) [\[60864\]](#) [\[65157\]](#) [\[65170\]](#)

Loperamide; Simethicone: (Major) Avoid coadministration of chloroquine with loperamide due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. At high doses, loperamide has been associated with serious cardiac toxicities, including syncope, ventricular tachycardia, QT prolongation, TdP, and cardiac arrest. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[30106\]](#) [\[60864\]](#) [\[65157\]](#) [\[65170\]](#)

Lopinavir; Ritonavir: (Major) Avoid coadministration of chloroquine and lopinavir due the risk of additive QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); fatalities have been reported. The risk of QT prolongation is increased with higher chloroquine doses. Lopinavir is also associated with QT prolongation. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28341\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Macimorelin: (Major) Avoid coadministration of chloroquine with macimorelin due to the increased risk of QT prolongation and torsade de pointes-type ventricular tachycardia. Allow a sufficient washout period before starting macimorelin. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Treatment with macimorelin has been associated with an increase in the corrected QT (QTc) interval. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[62723\]](#) [\[65157\]](#) [\[65170\]](#)

Magnesium Hydroxide: (Major) Chloroquine absorption may be reduced by antacids. Administer chloroquine and antacids at least 4 hours apart. [\[29758\]](#) [\[30285\]](#)

Maprotiline: (Major) Avoid coadministration of chloroquine with maprotiline due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased

with higher chloroquine doses. Maprotiline has been reported to prolong the QT interval, particularly in overdose or with higher-dose prescription therapy (elevated serum concentrations). Cases of long QT syndrome and TdP tachycardia have been described with maprotiline use, but rarely occur when the drug is used alone in normal prescribed doses and in the absence of other known risk factors for QT prolongation. Limited data are available regarding the safety of maprotiline in combination with other QT-prolonging drugs. [28225] [28229] [28230] [28231] [29758] [51740] [65157] [65170]

Mefloquine: (Major) Avoid coadministration of chloroquine with mefloquine due to the increased risk of QT prolongation and seizures. These drugs are both analogs of quinine. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. There is evidence that the use of halofantrine after mefloquine causes a significant lengthening of the QTc interval. Mefloquine alone has not been reported to cause QT prolongation. Also, both drugs may lower the seizure threshold. [28229] [28230] [28231] [28301] [29758] [65157] [65170]

Meglitinides: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the meglitinides, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Meperidine; Promethazine: (Major) Avoid coadministration of chloroquine with promethazine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Promethazine is associated with a possible risk for QT prolongation. [28225] [28229] [28230] [28231] [29758] [55578] [65157] [65170]

Mepivacaine: (Moderate) Coadministration of mepivacaine with oxidizing agents, such as chloroquine, may increase the risk of developing methemoglobinemia. Monitor patients closely for signs and symptoms of methemoglobinemia if coadministration is necessary. If methemoglobinemia occurs or is suspected, discontinue mepivacaine and any other oxidizing agents. Depending on the severity of symptoms, patients may respond to supportive care; more severe symptoms may require treatment with methylene blue, exchange transfusion, or hyperbaric oxygen. [29100]

Mepivacaine; Levonordefrin: (Moderate) Coadministration of mepivacaine with oxidizing agents, such as chloroquine, may increase the risk of developing methemoglobinemia. Monitor patients closely for signs and symptoms of methemoglobinemia if coadministration is necessary. If methemoglobinemia occurs or is suspected, discontinue mepivacaine and any other oxidizing agents. Depending on the severity of symptoms, patients may respond to supportive care; more severe symptoms may require treatment with methylene blue, exchange transfusion, or hyperbaric oxygen. [29100]

Metaproterenol: (Major) Avoid coadministration of chloroquine with short-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with

adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists as compared to short-acting beta-agonists. [28229] [28532] [29758] [49951] [51793] [59321] [65157] [65170]

Metformin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including metformin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Metformin; Pioglitazone: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including metformin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the thiazolidinediones, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Metformin; Repaglinide: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including metformin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the meglitinides, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Metformin; Rosiglitazone: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including metformin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the thiazolidinediones, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Metformin; Saxagliptin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including metformin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the dipeptidyl peptidase-4 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Metformin; Sitagliptin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including metformin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758] (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the dipeptidyl peptidase-4 inhibitors, are coadministered. A decreased dose of the antidiabetic

agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Methadone: (Major) Avoid coadministration of chloroquine with methadone due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Methadone is considered to be associated with an increased risk for QT prolongation and TdP, especially at higher doses (more than 200 mg/day but averaging approximately 400 mg/day in adult patients). Most cases involve patients being treated for pain with large, multiple daily doses of methadone, although cases have been reported in patients receiving doses commonly used for maintenance treatment of opioid addiction. [28229] [28230] [28231] [28319] [28320] [28321] [28322] [29758] [33136] [65157] [65170]

Metronidazole: (Major) Avoid coadministration of chloroquine with metronidazole due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Potential QT prolongation has been reported in limited case reports with metronidazole. [28229] [28230] [28231] [29758] [57377] [57378] [65157] [65170]

Midostaurin: (Major) Avoid coadministration of chloroquine with midostaurin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. QT prolongation was reported in patients who received midostaurin in clinical trials. [28229] [28230] [28231] [29758] [61906] [65157] [65170]

Mifepristone: (Major) Avoid coadministration of chloroquine with mifepristone due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Mifepristone is associated with dose-related prolongation of the QT interval. [28229] [28230] [28231] [29758] [48697] [65157] [65170]

Miglitol: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the alpha-glucosidase inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Mirtazapine: (Major) Avoid coadministration of chloroquine with mirtazapine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Mirtazapine has been associated with dose-dependent prolongation of the QT interval. TdP has been reported postmarketing, primarily in overdose or in patients with other risk factors for QT prolongation. [28229] [28230] [28231] [29758] [40942] [65157] [65170]

Moxifloxacin: (Major) Avoid coadministration of chloroquine with moxifloxacin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Quinolones have been associated with a risk of QT prolongation. TdP has been reported during postmarketing surveillance of moxifloxacin. These reports generally involved patients with concurrent medical conditions or concomitant medications that may have been contributory. [28229] [28230] [28231] [28423] [29758] [59321] [65157] [65170]

Nateglinide: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the meglitinides, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Nilotinib: (Major) Avoid coadministration of chloroquine with nilotinib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Sudden death and QT interval prolongation have occurred in patients who received nilotinib therapy. [28229] [28230] [28231] [29758] [58766] [65157] [65170]

Norfloxacin: (Major) Avoid coadministration of chloroquine with norfloxacin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Quinolones have been associated with a risk of QT prolongation and torsade de pointes (TdP). Although extremely rare, TdP has been reported during postmarketing surveillance of norfloxacin. These reports generally involved patients with concurrent medical conditions or concomitant medications that may have been contributory. [28225] [28229] [28230] [28231] [28432] [28457] [29758] [29818] [65157] [65170]

Nortriptyline: (Major) Avoid coadministration of chloroquine with tricyclic antidepressants (TCAs) due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. TCAs share pharmacologic properties similar to the Class IA antiarrhythmic agents and may prolong the QT interval, particularly in overdose or with higher-dose prescription therapy (elevated serum concentrations). [28225] [28229] [28230] [28231] [28415] [28416] [29758] [65157] [65170]

Octreotide: (Major) Avoid coadministration of chloroquine with octreotide due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Arrhythmias, sinus bradycardia, and conduction disturbances have occurred during octreotide therapy. Since bradycardia is a risk factor for development of TdP, the potential occurrence of bradycardia during octreotide administration could theoretically increase

the risk of TdP in patients receiving drugs that prolong the QT interval. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28432\]](#) [\[29113\]](#) [\[29758\]](#) [\[30624\]](#) [\[64969\]](#) [\[65157\]](#) [\[65170\]](#)

Ofloxacin: (Major) Avoid coadministration of chloroquine with ofloxacin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Quinolones have been associated with a risk of QT prolongation and TdP. Although extremely rare, torsade de pointes has been reported during postmarketing surveillance of ofloxacin. These reports generally involved patients with concurrent medical conditions or concomitant medications that may have been contributory. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[30738\]](#) [\[65157\]](#) [\[65170\]](#)

Olanzapine: (Major) Avoid coadministration of chloroquine with olanzapine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Limited data, including some case reports, suggest that olanzapine may be associated with a significant prolongation of the QTc interval. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28785\]](#) [\[29758\]](#) [\[32732\]](#) [\[32734\]](#) [\[32745\]](#) [\[65157\]](#) [\[65170\]](#)

Olodaterol: (Major) Avoid coadministration of chloroquine with long-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists (LABAs) as compared to short-acting beta-agonists.. [\[28229\]](#) [\[29758\]](#) [\[41231\]](#) [\[44026\]](#) [\[44063\]](#) [\[44979\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Omeprazole; Sodium Bicarbonate: (Major) Chloroquine absorption may be reduced by antacids. Administer chloroquine and antacids at least 4 hours apart. [\[29758\]](#) [\[30284\]](#) [\[30285\]](#)

Ondansetron: (Major) Avoid coadministration of chloroquine with ondansetron due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Ondansetron has been associated with a dose-related increase in the QT interval and postmarketing reports of TdP. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[31266\]](#) [\[32722\]](#) [\[34335\]](#) [\[34353\]](#) [\[65157\]](#) [\[65170\]](#)

Osilodrostat: (Major) Avoid coadministration of chloroquine with osilodrostat due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Osilodrostat is associated with dose-dependent QT prolongation. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[65098\]](#) [\[65157\]](#) [\[65170\]](#)

Osimertinib: (Major) Avoid coadministration of chloroquine with osimertinib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. An interruption of osimertinib therapy with dose reduction or discontinuation may be necessary if QT prolongation occurs. Concentration-dependent QTc prolongation occurred during clinical trials of osimertinib. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. [28229] [28230] [28231] [29758] [60297] [65157] [65170]

Oxaliplatin: (Major) Avoid coadministration of chloroquine with oxaliplatin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. QT prolongation and ventricular arrhythmias including fatal TdP have been reported with oxaliplatin use in postmarketing experience. [28229] [28230] [28231] [28415] [29758] [41958] [65157] [65170]

Paliperidone: (Major) Avoid coadministration of chloroquine with paliperidone due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Paliperidone has been associated with QT prolongation; TdP and ventricular fibrillation have been reported in the setting of overdose. [28229] [28230] [28231] [29758] [40936] [65157] [65170]

Panobinostat: (Major) Avoid coadministration of chloroquine with panobinostat due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. QT prolongation has also been reported with panobinostat. [28229] [28230] [28231] [29758] [58821] [65157] [65170]

Pasireotide: (Major) Avoid coadministration of chloroquine with pasireotide due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. QT prolongation has occurred with pasireotide at therapeutic and supra-therapeutic doses. [28229] [28230] [28231] [29758] [65157] [65170]

Pazopanib: (Major) Avoid coadministration of chloroquine with pazopanib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Pazopanib has also been reported to prolong the QT interval. [28229] [28230] [28231] [29758] [37098] [65157] [65170]

Peginterferon Alfa-2a: (Moderate) Concurrent use of chloroquine and interferons is not recommended as there is an increased risk of retinal toxicity. [29758] [47391]

Peginterferon Alfa-2b: (Moderate) Concurrent use of chloroquine and interferons is not recommended as there is an increased risk of retinal toxicity. [29758] [47391]

Peginterferon beta-1a: (Moderate) Concurrent use of chloroquine and interferons is not recommended as there is an increased risk of retinal toxicity. [29758] [47391]

Penicillamine: (Major) Do not use penicillamine concurrently with antimalarials due to an increased risk of severe hematologic and renal adverse reactions. [28834]

Penicillin G Benzathine; Penicillin G Procaine: (Moderate) Coadministration of penicillin G procaine with oxidizing agents, such as chloroquine, may increase the risk of developing methemoglobinemia. Monitor patients closely for signs and symptoms of methemoglobinemia if coadministration is necessary. If methemoglobinemia occurs or is suspected, discontinue penicillin G procaine and any other oxidizing agents. Depending on the severity of symptoms, patients may respond to supportive care; more severe symptoms may require treatment with methylene blue, exchange transfusion, or hyperbaric oxygen. [31217]

Penicillin G Procaine: (Moderate) Coadministration of penicillin G procaine with oxidizing agents, such as chloroquine, may increase the risk of developing methemoglobinemia. Monitor patients closely for signs and symptoms of methemoglobinemia if coadministration is necessary. If methemoglobinemia occurs or is suspected, discontinue penicillin G procaine and any other oxidizing agents. Depending on the severity of symptoms, patients may respond to supportive care; more severe symptoms may require treatment with methylene blue, exchange transfusion, or hyperbaric oxygen. [31217]

Pentamidine: (Major) Avoid coadministration of chloroquine with pentamidine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Systemic pentamidine has also been associated with QT prolongation. [23620] [23778] [28229] [28230] [28231] [28419] [28879] [29758] [65157] [65170]

Perphenazine: (Major) Avoid coadministration of chloroquine with perphenazine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Perphenazine is associated with a possible risk for QT prolongation. [28229] [28230] [28231] [28415] [29758] [65157] [65170]

Perphenazine; Amitriptyline: (Major) Avoid coadministration of chloroquine with perphenazine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Perphenazine is associated with a possible risk for QT prolongation. [28229] [28230] [28231] [28415] [29758] [65157] [65170] (Major) Avoid coadministration of chloroquine with tricyclic antidepressants (TCAs) due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT

prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. TCAs share pharmacologic properties similar to the Class IA antiarrhythmic agents and may prolong the QT interval, particularly in overdose or with higher-dose prescription therapy (elevated serum concentrations). [\[28225\]](#) [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28415\]](#) [\[28416\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Phenobarbital: (Moderate) Coadministration of chloroquine and phenobarbital may decrease exposure of chloroquine which may reduce its efficacy. Chloroquine may be an in vitro CYP3A4 substrate and phenobarbital is a strong CYP3A4 inducer. [\[65210\]](#) [\[65237\]](#) [\[65239\]](#)

Phenylephrine; Promethazine: (Major) Avoid coadministration of chloroquine with promethazine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Promethazine is associated with a possible risk for QT prolongation. [\[28225\]](#) [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[55578\]](#) [\[65157\]](#) [\[65170\]](#)

Phenytoin: (Moderate) Coadministration of chloroquine and phenytoin may decrease exposure of chloroquine which may reduce its efficacy. Chloroquine may be an in vitro CYP3A4 substrate and phenytoin is a strong CYP3A4 inducer. [\[65210\]](#) [\[65237\]](#) [\[65239\]](#)

Pimavanserin: (Major) Avoid coadministration of chloroquine with pimavanserin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Pimavanserin also prolongs the QT interval. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[60748\]](#) [\[65157\]](#) [\[65170\]](#)

Pimozide: (Severe) Coadministration of chloroquine with pimozide is contraindicated due to the increased risk of QT prolongation. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Pimozide is associated with a well-established risk of QT prolongation and TdP. [\[28225\]](#) [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[43463\]](#) [\[65170\]](#)

Pioglitazone: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the thiazolidinediones, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Pirbuterol: (Major) Avoid coadministration of chloroquine with short-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists as compared to short-acting beta-agonists. [\[28229\]](#) [\[28532\]](#) [\[29758\]](#) [\[49951\]](#) [\[51793\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Pitolisant: (Major) Avoid coadministration of chloroquine with pitolisant due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Pitolisant also prolongs the QT interval. [28229] [28230] [28231] [29758] [64562] [65157] [65170]

Ponatinib: (Moderate) Concurrent use of chloroquine and ponatinib is not recommended as there is an increased risk of retinal toxicity. [29758] [52603]

Posaconazole: (Major) Avoid coadministration of chloroquine with posaconazole due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Posaconazole has been associated with QT prolongation and TdP. [28229] [28230] [28231] [29758] [32723] [65157] [65170]

Pramlintide: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including pramlintide, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Praziquantel: (Minor) Concomitant administration of chloroquine and praziquantel can reduce praziquantel bioavailability and maximum serum concentrations. The mechanism of the interaction is not certain. Clinicians should be alert to the possibility of praziquantel failure if chloroquine is used. [27846] [29758]

Prilocaine: (Moderate) Coadministration of prilocaine with oxidizing agents, such as chloroquine, may increase the risk of developing methemoglobinemia. Monitor patients closely for signs and symptoms of methemoglobinemia if coadministration is necessary. If methemoglobinemia occurs or is suspected, discontinue prilocaine and any other oxidizing agents. Depending on the severity of symptoms, patients may respond to supportive care; more severe symptoms may require treatment with methylene blue, exchange transfusion, or hyperbaric oxygen. [29064]

Prilocaine; Epinephrine: (Moderate) Coadministration of prilocaine with oxidizing agents, such as chloroquine, may increase the risk of developing methemoglobinemia. Monitor patients closely for signs and symptoms of methemoglobinemia if coadministration is necessary. If methemoglobinemia occurs or is suspected, discontinue prilocaine and any other oxidizing agents. Depending on the severity of symptoms, patients may respond to supportive care; more severe symptoms may require treatment with methylene blue, exchange transfusion, or hyperbaric oxygen. [29064]

Primaquine: (Major) Avoid coadministration of chloroquine with primaquine due to the increased risk of QT prolongation or other toxicities. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Primaquine may also prolong the QT interval. [28229] [28230] [28231] [29758] [41984] [65157] [65170]

Procainamide: (Major) Avoid coadministration of chloroquine with procainamide due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess

initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Procainamide is associated with a well-established risk of QT prolongation and TdP. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Prochlorperazine: (Major) Avoid coadministration of chloroquine with prochlorperazine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Prochlorperazine is associated with a possible risk for QT prolongation. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28415\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Promethazine: (Major) Avoid coadministration of chloroquine with promethazine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Promethazine is associated with a possible risk for QT prolongation. [\[28225\]](#) [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[55578\]](#) [\[65157\]](#) [\[65170\]](#)

Propafenone: (Major) Avoid coadministration of chloroquine with propafenone due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Propafenone is a Class IC antiarrhythmic which increases the QT interval, but largely due to prolongation of the QRS interval. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28287\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Protriptyline: (Major) Avoid coadministration of chloroquine with tricyclic antidepressants (TCAs) due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. TCAs share pharmacologic properties similar to the Class IA antiarrhythmic agents and may prolong the QT interval, particularly in overdose or with higher-dose prescription therapy (elevated serum concentrations). [\[28225\]](#) [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28415\]](#) [\[28416\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Quetiapine: (Major) Avoid coadministration of chloroquine with quetiapine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Limited data, including some case reports, suggest that quetiapine may be associated with a significant prolongation of the QTc interval in rare instances. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29118\]](#) [\[29758\]](#) [\[33068\]](#) [\[33072\]](#) [\[33074\]](#) [\[65157\]](#) [\[65170\]](#)

Quinidine: (Major) Avoid coadministration of chloroquine with quinidine due to the increased risk of QT prolongation or other drug toxicities. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any

non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Quinidine administration is also associated with QT prolongation and TdP. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[42280\]](#) [\[47357\]](#) [\[65157\]](#) [\[65170\]](#)

Quinine: (Major) Avoid coadministration of chloroquine with quinine due to the increased risk of QT prolongation or other drug toxicities. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Quinine has been associated with QT prolongation and rare cases of TdP. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[31403\]](#) [\[65157\]](#) [\[65170\]](#)

Rabies Vaccine: (Major) If administered concurrently, antimalarials can impair the immunologic response to the rabies vaccine, thereby, decreasing its protective effect. If possible, administration of antimalarials should be avoided during use of the rabies vaccine for postexposure prophylaxis. When antimalarials must be administered to persons also receiving the rabies vaccine for postexposure prophylaxis, a serum rabies antibody titer should be obtained on day 14 (day of the 4th vaccination) to ensure an acceptable antibody response has been induced. [\[40848\]](#) [\[40849\]](#)

Ranolazine: (Major) Avoid coadministration of chloroquine with ranolazine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Ranolazine is associated with dose- and plasma concentration-related increases in the QTc interval. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[31938\]](#) [\[65157\]](#) [\[65170\]](#)

Regular Insulin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including insulin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Regular Insulin; Isophane Insulin (NPH): (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including insulin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Repaglinide: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the meglitinides, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Ribociclib: (Major) Avoid coadministration of chloroquine with ribociclib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Ribociclib has been shown to prolong the QT interval in a concentration-dependent manner. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[61816\]](#) [\[65157\]](#) [\[65170\]](#)

Ribociclib; Letrozole: (Major) Avoid coadministration of chloroquine with ribociclib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess

initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Ribociclib has been shown to prolong the QT interval in a concentration-dependent manner. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[61816\]](#) [\[65157\]](#) [\[65170\]](#)

Rilpivirine: (Major) Avoid coadministration of chloroquine with rilpivirine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Supratherapeutic doses of rilpivirine (75 to 300 mg/day) have caused QT prolongation. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[44376\]](#) [\[65157\]](#) [\[65170\]](#)

Risperidone: (Major) Avoid coadministration of chloroquine with risperidone due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Risperidone has been associated with a possible risk for QT prolongation and/or TdP, primarily in the overdose setting. [\[22256\]](#) [\[28229\]](#) [\[28414\]](#) [\[29758\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Romidepsin: (Major) Avoid coadministration of chloroquine with romidepsin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Romidepsin has been reported to prolong the QT interval. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[37292\]](#) [\[65157\]](#) [\[65170\]](#)

Ropivacaine: (Moderate) Coadministration of ropivacaine with oxidizing agents, such as chloroquine, may increase the risk of developing methemoglobinemia. Monitor patients closely for signs and symptoms of methemoglobinemia if coadministration is necessary. If methemoglobinemia occurs or is suspected, discontinue ropivacaine and any other oxidizing agents. Depending on the severity of symptoms, patients may respond to supportive care; more severe symptoms may require treatment with methylene blue, exchange transfusion, or hyperbaric oxygen. [\[52330\]](#)

Rosiglitazone: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the thiazolidinediones, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Salmeterol: (Major) Avoid coadministration of chloroquine with long-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval.

This risk may be more clinically significant with long-acting beta-agonists (LABAs) as compared to short-acting beta-agonists.. [\[28229\]](#) [\[29758\]](#) [\[41231\]](#) [\[44026\]](#) [\[44063\]](#) [\[44979\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Saquinavir: (Major) Avoid coadministration of chloroquine with saquinavir boosted with ritonavir due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Saquinavir boosted with ritonavir increases the QT interval in a dose-dependent fashion, which may increase the risk for serious arrhythmias such as TdP. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28995\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Saxagliptin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the dipeptidyl peptidase-4 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Semaglutide: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the incretin mimetics, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Sertraline: (Major) Avoid coadministration of chloroquine with sertraline due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. The risk of sertraline-induced QT prolongation is generally considered to be low in clinical practice. Its effect on QTc interval is minimal (typically less than 5 msec), and the drug has been used safely in patients with cardiac disease (e.g., recent myocardial infarction, unstable angina, chronic heart failure). [\[28229\]](#) [\[28343\]](#) [\[29758\]](#) [\[64391\]](#) [\[64392\]](#) [\[64394\]](#) [\[64396\]](#) [\[65157\]](#) [\[65170\]](#)

Sevoflurane: (Major) Avoid coadministration of chloroquine with halogenated anesthetics due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Halogenated anesthetics are known to prolong the QT interval. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28457\]](#) [\[28458\]](#) [\[28754\]](#) [\[28755\]](#) [\[28756\]](#) [\[29758\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

SGLT2 Inhibitors: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the SGLT2 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Short-acting beta-agonists: (Major) Avoid coadministration of chloroquine with short-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when

associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists as compared to short-acting beta-agonists. [28229] [28532] [29758] [49951] [51793] [59321] [65157] [65170]

Simvastatin; Sitagliptin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the dipeptidyl peptidase-4 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Siponimod: (Major) Avoid coadministration of chloroquine with siponimod due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Siponimod therapy prolonged the QT interval at recommended doses in a clinical study. [28229] [28230] [28231] [29758] [64031] [65157] [65170]

Sitagliptin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the dipeptidyl peptidase-4 inhibitors, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Sodium Bicarbonate: (Major) Chloroquine absorption may be reduced by antacids. Administer chloroquine and antacids at least 4 hours apart. [29758] [30284] [30285]

Solifenacin: (Major) Avoid coadministration of chloroquine with solifenacin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Solifenacin has been associated with dose-dependent prolongation of the QT interval. TdP has been reported with postmarketing use, although causality was not determined. [28229] [28230] [28231] [29758] [30515] [65157] [65170]

Sorafenib: (Major) Avoid coadministration of chloroquine with sorafenib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. An interruption or discontinuation of sorafenib therapy may be necessary if QT prolongation occurs. Sorafenib has been associated with QT prolongation. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses [28229] [28230] [28231] [29758] [31832] [65157] [65170]

Sotalol: (Major) Avoid coadministration of chloroquine with sotalol due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Sotalol administration is associated with QT prolongation and TdP. Proarrhythmic events should be anticipated after initiation of therapy and after each upward dosage adjustment. [28229] [28230] [28231] [28234] [29758] [65157] [65170]

Succinylcholine: (Moderate) Chloroquine may enhance the neuromuscular blocking action of succinylcholine. [42039]

Sulfonylureas: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the sulfonylureas, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Sunitinib: (Major) Avoid coadministration of chloroquine with sunitinib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Sunitinib can also prolong the QT interval. [28229] [28230] [28231] [29758] [65157] [65170]

Tacrolimus: (Major) Avoid coadministration of chloroquine with tacrolimus due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Tacrolimus may prolong the QT interval and cause TdP. [28229] [28230] [28231] [28611] [29758] [55401] [60497] [65157] [65170]

Tamoxifen: (Major) Avoid coadministration of chloroquine with tamoxifen due to the increased risk of QT prolongation and retinal toxicity. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Tamoxifen has been reported to prolong the QT interval, usually in overdose or when used in high doses. Rare case reports of QT prolongation have also been described when tamoxifen is used at lower doses. [28229] [28230] [28231] [29758] [61872] [63589] [65157] [65170]

Telavancin: (Major) Avoid coadministration of chloroquine with telavancin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Telavancin has also been associated with QT prolongation. [28229] [28230] [28231] [29758] [36615] [65157] [65170]

Telbivudine: (Minor) Monitor patients for signs or symptoms of unexplained muscle pain, tenderness, or weakness during concomitant treatment with chloroquine and telbivudine. Interrupt telbivudine therapy if myopathy is suspected and discontinue telbivudine if myopathy is confirmed. It is unknown if the risk of myopathy during treatment with telbivudine is increased with coadministration of other drugs associated with myopathy, like chloroquine. [32827]

Telithromycin: (Major) Avoid coadministration of chloroquine with telithromycin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is

increased with higher chloroquine doses. Telithromycin is also associated with QT prolongation and TdP. [\[28156\]](#) [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Terbutaline: (Major) Avoid coadministration of chloroquine with short-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists as compared to short-acting beta-agonists. [\[28229\]](#) [\[28532\]](#) [\[29758\]](#) [\[49951\]](#) [\[51793\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Tetrabenazine: (Major) Avoid coadministration of chloroquine with tetrabenazine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Tetrabenazine causes a small increase in the corrected QT interval (QTc). [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[34389\]](#) [\[65157\]](#) [\[65170\]](#)

Tetracaine: (Moderate) Coadministration of tetracaine with oxidizing agents, such as chloroquine, may increase the risk of developing methemoglobinemia. Monitor patients closely for signs and symptoms of methemoglobinemia if coadministration is necessary. If methemoglobinemia occurs or is suspected, discontinue tetracaine and any other oxidizing agents. Depending on the severity of symptoms, patients may respond to supportive care; more severe symptoms may require treatment with methylene blue, exchange transfusion, or hyperbaric oxygen. [\[31353\]](#)

Thiazolidinediones: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the thiazolidinediones, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [\[29758\]](#)

Thioridazine: (Severe) Coadministration of chloroquine with thioridazine is contraindicated due to the increased risk of QT prolongation. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Thioridazine is associated with a well-established risk of QT prolongation and TdP. [\[28225\]](#) [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28293\]](#) [\[29758\]](#) [\[65170\]](#)

Tiotropium; Olodaterol: (Major) Avoid coadministration of chloroquine with long-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists (LABAs) as compared to short-acting beta-agonists.. [\[28229\]](#) [\[29758\]](#) [\[41231\]](#) [\[44026\]](#) [\[44063\]](#) [\[44979\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Tolazamide: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the sulfonylureas, are coadministered. A decreased dose of the

antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Tolbutamide: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including the sulfonylureas, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Tolterodine: (Major) Avoid coadministration of chloroquine with tolterodine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Tolterodine has been associated with dose-dependent prolongation of the QT interval, especially in poor CYP2D6 metabolizers. [28229] [28230] [28231] [29758] [31112] [65157] [65170]

Toremifene: (Major) Avoid coadministration of chloroquine with toremifene due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Toremifene has been shown to prolong the QTc interval in a dose- and concentration-related manner. [28229] [28230] [28231] [28822] [29758] [65157] [65170]

Trametinib: (Moderate) Concurrent use of chloroquine and trametinib is not recommended as there is an increased risk of retinal toxicity. [29758] [60372]

Trazodone: (Major) Avoid coadministration of chloroquine with trazodone due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Trazodone can prolong the QT/QTc interval at therapeutic doses. In addition, there are postmarketing reports of TdP. [28229] [28230] [28231] [29758] [38831] [63609] [65157] [65170]

Tricyclic antidepressants: (Major) Avoid coadministration of chloroquine with tricyclic antidepressants (TCAs) due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. TCAs share pharmacologic properties similar to the Class IA antiarrhythmic agents and may prolong the QT interval, particularly in overdose or with higher-dose prescription therapy (elevated serum concentrations). [28225] [28229] [28230] [28231] [28415] [28416] [29758] [65157] [65170]

Trifluoperazine: (Major) Avoid coadministration of chloroquine with trifluoperazine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Trifluoperazine is associated with a possible risk for QT prolongation. [28229] [28230] [28231] [28415] [29758] [65157] [65170]

Trimipramine: (Major) Avoid coadministration of chloroquine with tricyclic antidepressants (TCAs) due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. TCAs share pharmacologic properties similar to the Class IA antiarrhythmic agents and may prolong the QT interval, particularly in overdose or with higher-dose prescription therapy (elevated serum concentrations). [28225] [28229] [28230] [28231] [28415] [28416] [29758] [65157] [65170]

Triptorelin: (Major) Avoid coadministration of chloroquine with triptorelin due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Androgen deprivation therapy (i.e., triptorelin) may prolong the QT interval. [28229] [28230] [28231] [29758] [45411] [65157] [65170]

Ultralente Insulin: (Major) Careful monitoring of blood glucose is recommended when chloroquine and antidiabetic agents, including insulin, are coadministered. A decreased dose of the antidiabetic agent may be necessary as severe hypoglycemia has been reported in patients treated concomitantly with chloroquine and an antidiabetic agent. [29758]

Umeclidinium; Vilanterol: (Major) Avoid coadministration of chloroquine with long-acting beta-agonists due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Beta-agonists may be associated with adverse cardiovascular effects including QT interval prolongation, usually at higher doses, when associated with hypokalemia, or when used with other drugs known to prolong the QT interval. This risk may be more clinically significant with long-acting beta-agonists (LABAs) as compared to short-acting beta-agonists.. [28229] [29758] [41231] [44026] [44063] [44979] [59321] [65157] [65170]

Vandetanib: (Major) Avoid coadministration of chloroquine with vandetanib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. An interruption of vandetanib therapy or dose reduction may be necessary for QT prolongation. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Vandetanib can prolong the QT interval in a concentration-dependent manner; TdP and sudden death have been reported in patients receiving vandetanib. [28229] [28230] [28231] [29758] [43901] [65157] [65170]

Vardenafil: (Major) Avoid coadministration of chloroquine with vardenafil due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Vardenafil is associated with QT prolongation. Both therapeutic and suprathreshold doses of vardenafil produce an increase in QTc interval. [28216] [28229] [29758] [41124] [59321] [65157] [65170]

Vemurafenib: (Major) Avoid coadministration of chloroquine with vemurafenib due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Vemurafenib has also been associated with QT prolongation. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[45335\]](#) [\[59321\]](#) [\[65157\]](#) [\[65170\]](#)

Venlafaxine: (Major) Avoid coadministration of chloroquine with venlafaxine due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Venlafaxine administration is associated with a possible risk of QT prolongation; TdP has been reported with postmarketing use. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[33715\]](#) [\[65157\]](#) [\[65170\]](#)

Vigabatrin: (Major) Vigabatrin should not be used with chloroquine due to potential retinal toxicity associated with both drugs, unless the benefits of treatment clearly outweigh the risks. [\[29758\]](#) [\[36250\]](#)

Voriconazole: (Major) Avoid coadministration of chloroquine with voriconazole due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Voriconazole has been associated with QT prolongation and rare cases of TdP. [\[28158\]](#) [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Vorinostat: (Major) Avoid coadministration of chloroquine with vorinostat due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Vorinostat therapy is also associated with a risk of QT prolongation. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[29758\]](#) [\[32789\]](#) [\[65157\]](#) [\[65170\]](#)

Ziprasidone: (Major) Avoid coadministration of chloroquine with ziprasidone due to the increased risk of QT prolongation. If use together is necessary, obtain an ECG at baseline to assess initial QT interval and determine frequency of subsequent ECG monitoring, avoid any non-essential QT prolonging drugs, and correct electrolyte imbalances. Chloroquine is associated with an increased risk of QT prolongation and torsade de pointes (TdP); the risk of QT prolongation is increased with higher chloroquine doses. Clinical trial data indicate that ziprasidone causes QT prolongation; there are postmarketing reports of TdP in patients with multiple confounding factors. [\[28229\]](#) [\[28230\]](#) [\[28231\]](#) [\[28233\]](#) [\[29758\]](#) [\[65157\]](#) [\[65170\]](#)

Revision Date: 04/15/2020 02:37:00 AM

References

22256 – Risperdal Consta (risperidone long-acting injection) package insert. Titusville, NJ: Janssen Pharmaceuticals, Inc.; 2020 Jan.

- 23500** – Wilt JL, Minnema AM, Johnson RF, et al. Torsade de pointes associated with the use of intravenous haloperidol. *Ann Intern Med* 1993;119:391-4.
- 23620** – Green PT, Reents S, Harman E, et al. Pentamidine-induced torsades de pointes in a renal transplant recipient with *Pneumocystis carinii* pneumonia. *S Med J* 1990;83:481-4.
- 23774** – Lui HK, Lee G, Dietrich P, et al. Flecainide-induced QT prolongation and ventricular tachycardia. *Am Heart J* 1982;103:567-9.
- 23778** – Wharton JM, Demopoulos PA, Goldschlager N. Torsade de pointes during administration of pentamidine isethionate. *Am J Med* 1987;83:571-6.
- 23779** – Kriwisky M, Perry GY, Tarchitsky D, et al. Haloperidol-induced torsades de pointes. *Chest* 1990;98:482-3.
- 27846** – Masimirembwa CM, Naik YS, Hasler JA. The effect of chloroquine on the pharmacokinetics and metabolism of praziquantel in rats and humans. *Biopharm Drug Dispos* 1994;15:33-43.
- 27982** – Ketoconazole tablets package insert. Morgantown, WV: Mylan Pharmaceuticals, Inc.; 2017 Sept.
- 27983** – Sporanox (itraconazole) capsules package insert. Titusville, NJ: Janssen Pharmaceuticals, Inc.; 2019 Mar.
- 28156** – Ketek (telithromycin) package insert. Bridgewater, NJ: Sanofi-Aventis Pharmaceuticals; 2015 Oct.
- 28158** – VFEND (voriconazole) tablets, suspension, and injection package insert. New York, NY: Pfizer Inc; 2019 Jan.
- 28216** – Levitra (vardenafil) package insert. Kenilworth, NJ: Schering-Plough; 2017 Aug.
- 28221** – Tikosyn (dofetilide) package insert. New York, NY: Pfizer Labs; 2019 Aug.
- 28224** – Cordarone (amiodarone) tablets package insert. Philadelphia, PA: Wyeth Pharmaceuticals Inc.; 2018 Oct.
- 28225** – CredibleMeds. Drugs to avoid in congenital long QT. Available on the World Wide Web at <http://www.crediblemeds.org>.
- 28226** – Trisenox (arsenic trioxide) package insert. Frazer, PA: Cephalon, Inc; 2010 Jun.
- 28228** – Norpace and Norpace CR (disopyramide) package insert. Chicago, IL: G.D. Searle LLC division of Pfizer Inc; 2016 Aug.
- 28229** – Demaziere J, Fourcade JM, Busseuil CT, et al. The hazards of chloroquine self prescription in west Africa. *J Toxicol Clin Toxicol* 1995;33:369-70.
- 28230** – Mansfield RJ, Thomas RD. Recurrent syncope. Drug induced long QT syndrome. *Postgrad Med J* 2001;77:344, 352-3.
- 28231** – Pinski SL, Eguia LE, Trohman RG. What is the minimal pacing rate that prevents torsades de pointes? Insights from patients with permanent pacemakers. *Pacing Clin Electrophysiol* 2002;25:1612-5.

- 28233** – Geodon (ziprasidone) package insert. New York, NY: Pfizer; 2020 Jan.
- 28234** – Betapace (sotalol) package insert. Wayne, NJ: Berlex Laboratories; 2011 Aug.
- 28235** – Richards JR, Schneir AB. Droperidol in the emergency department: is it safe? *J Emerg Med* 2003;24:441-7.
- 28236** – Kao LW, Kirk MA, Evers SJ, et al. Droperidol, QT prolongation, and sudden death: what is the evidence? *Ann Emerg Med* 2003;41:546-58.
- 28237** – Food and Drug Administration Press Office. FDA strengthens warnings for droperidol. FDA Talk Paper. December 5, 2001. Accessed: April 16, 2004. Available on the World Wide Web at: <http://www.fda.gov/bbs/topics/answers/2001/ans01123.html>
- 28238** – Biaxin (clarithromycin) package insert. North Chicago, IL: AbbVie, Inc.; 2019 Sep.
- 28261** – Uroxatral (alfuzosin) package insert. Cary, NC: Covis Pharmaceuticals, Inc.; 2013 Sep.
- 28262** – Clozaril (clozapine) tablets package insert. Rosemont, PA: HLS Therapeutics (USA), Inc. (Clozaril is a registered trademark of Novartis AG); 2017 Feb.
- 28269** – Celexa (citalopram) package insert. Irvine, CA: Allergan USA, Inc.; 2019 Jan.
- 28270** – Lexapro (escitalopram) package insert. Irvine, CA: Allergan USA, Inc.; 2019 Jan.
- 28287** – Rythmol SR (propafenone hydrochloride) capsule extended release package insert. Research Triangle Park, NC: GlaxoSmithKline; 2018 Nov.
- 28293** – Thioridazine package insert. Morgantown, WV: Mylan Pharmaceuticals Inc.; 2016 Nov.
- 28301** – Mefloquine package insert. Princeton, NJ: Sandoz Inc.; 2013 Jul.
- 28307** – Haldol injection for immediate release (haloperidol) package insert. Titusville, NJ: Janssen Pharmaceuticals, Inc.; 2020 Feb.
- 28319** – Krantz MJ, Kutinsky IB, Robertson AD, et al. Dose-related effects of methadone on QT prolongation in a series of patients with torsade de pointes. *Pharmacotherapy* 2003;23:802-5.
- 28320** – Walker PW, Klein D, Kasza L. High dose methadone and ventricular arrhythmias: a report of three cases. *Pain* 2003;103:321-4.
- 28321** – Kornick CA, Kilborn MJ, Santiago-Palma J, et al. QTc interval prolongation associated with intravenous methadone. *Pain* 2003;105:499-506.
- 28322** – Gil M, Sala M, Anguera I, et al. QT prolongation and Torsades de Pointes in patients infected with human immunodeficiency virus and treated with methadone. *Am J Cardiol* 2003;92:995-7.
- 28341** – Kaletra (lopinavir; ritonavir) tablet and solution package insert. North Chicago, IL: AbbVie Inc; 2020 Mar.
- 28343** – Zoloft (sertraline) package insert. New York, NY: Pfizer; 2019 Apr.
- 28377** – Foscavir (foscarnet) package insert. Lake Forest, IL: Clinigen Healthcare, Ltd.; 2017 Feb.

28405 – Strattera (atomoxetine) package insert. Indianapolis, IN: Eli Lilly and Company; 2020 Feb.

28414 – Risperdal (risperidone tablets, oral solution, and orally disintegrating tablets) package insert. Titusville, NJ: Janssen Pharmaceuticals, Inc.; 2020 Jan.

28415 – Nora Goldschlager, Andrew E Epstein, Blair P Grubb, et al. Etiologic considerations in the patient with syncope and an apparently normal heart. *Arch Intern Med* 2003;163:151-62.

28416 – Hansten PD, Horn JR. Drug Interactions with Drugs that Increase QTc Intervals. In: *The Top 100 Drug Interactions - A Guide to Patient Management*. 2007 Edition. Freeland, WA: H&H Publications; 2007:144-8.

28419 – Owens RC Jr. Risk assessment for antimicrobial agent-induced QTc interval prolongation and torsades de pointes. *Pharmacotherapy* 2001;21:301-19.

28420 – Iannini PB. Cardiotoxicity of macrolides, ketolides and fluoroquinolones that prolong the QTc interval. *Expert Opin Drug Saf* 2002;1:121-8.

28421 – Levaquin (levofloxacin) tablet package insert. Titusville, NJ: Janssen Pharmaceutical, Inc.; 2019 Jul.

28423 – Avelox (moxifloxacin) package insert. Whitehouse Station, NJ: Merck and Co., Inc.; 2019 May.

28424 – Factive (gemifloxacin mesylate) package insert. Toronto, ON: Merus Labs International, Inc.; 2019 May.

28432 – Roden, DM. Drug-induced prolongation of the QT interval. *New Engl J Med* 2004;350:1013-22.

28442 – Sustiva (efavirenz) package insert. Princeton, NJ: Bristol-Myers Squibb Company; 2019 Oct.

28457 – Crouch MA, Limon L, Cassano AT. Clinical relevance and management of drug-related QT interval prolongation. *Pharmacotherapy* 2003;23:881-908.

28458 – Schmeling WT, Warltier DC, McDonald DJ, et al. Prolongation of the QT interval by enflurane, isoflurane, and halothane in humans. *Anesth Analg* 1991;72:137-44.

28532 – Proventil HFA (albuterol sulfate) Inhalational Aerosol package insert. Whitehouse Station, NJ: Merck Sharp & Dohme Corp., a subsidiary of MERCK & CO., INC.; 2017 Sept.

28592 – Zoladex (goserelin acetate 3.6 mg implant) package insert. Lake Forest, IL: TerSera Therapeutics LLC; 2019 Feb.

28611 – Prograf (tacrolimus) capsules, injection, and granules for oral suspension package insert. Astellas Pharma US, Inc.: Northbrook, IL; 2019 Jun.

28661 – Apokyn and Apokyn Pen (apomorphine) injection package insert. Louisville, KY: US WorldMeds LLC; 2019 May.

28674 – Diflucan oral tablet and suspension (fluconazole) package insert. New York, NY: Pfizer; 2019 Feb.

28737 – Inapsine (Droperidol) Injection package insert. Lake Forest, IL: Akorn, Inc.; 2011 Oct.

28752 – Thevenin J, Da Costa A, Roche F, et al. Flecainide induced ventricular tachycardia (torsades de pointes). *Pacing Clin Electrophysiol* 2003;26:1907-8.

28754 – Loeckinger A, Kleinsasser A, Maier S, et al. Sustained prolongation of the QTc interval after anesthesia with sevoflurane in infants during the first 6 months of life. *Anesthesiology* 2003;98:639-42.

28755 – Kleinsasser A, Loeckinger A, Lindner KH, et al. Reversing sevoflurane-associated Q-Tc prolongation by changing to propofol. *Anaesthesia* 2001;56:248-50.

28756 – Kuenszberg E, Loeckinger A, Kleinsasser A, et al. Sevoflurane progressively prolongs the QT interval in unpremedicated female adults. *Eur J Anaesthesiol* 2000;17:662-4.

28785 – Zyprexa (olanzapine, all formulations) package insert. Indianapolis, IN: Eli Lilly and Company; 2019 Oct.

28822 – Fareston (toremifene citrate) tablets package insert. Bedminster, NJ: Kyowa Kirin Inc.; 2017 May.

28834 – Cuprimine (penicillamine) package insert. Lawrenceville, NJ: Atom Pharma; 2010 Mar.

28855 – Zithromax (azithromycin 250 mg and 500 mg tablets and azithromycin oral suspension) package insert. New York, NY: Pfizer Inc.; 2019 Apr.

28879 – Pentamidine isethionate injection package insert. Manasquan, NJ: Seton Pharma, LLC; 2014 Jan.

28995 – Invirase (saquinavir) package insert. South San Francisco, CA: Genentech Inc.; 2019 Dec.

28996 – Septocaine (articaine; epinephrine injection) package insert. Louisville, CO: Septodont, Inc.; 2018 Nov.

29062 – Nesacaine (chloroprocaine hydrochloride) package insert. Lake Zurich, IL: Fresenius Kabi USA, LLC; 2018 Nov.

29064 – Citanest Plain Dental (prilocaine) injection package insert. York, PA: AstraZeneca; 2018 Nov.

29100 – Carbocaine (mepivacaine hydrochloride) package insert. Lake Forest, IL: Hospira, Inc.; 2018 Nov.

29113 – Sandostatin (octreotide) package insert. Stein, Switzerland: Novartis Pharma Stein AG Corporation; 2019 April.

29118 – Seroquel (quetiapine fumarate) package insert. Wilmington, DE: AstraZeneca Pharmaceuticals LP; 2020 Jan.

29758 – Aralen (chloroquine) package insert. Bridgewater, NJ: Sanofi-aventis U.S. LLC.; 2018 Oct.

29818 – Noroxin (norfloxacin) tablets package insert. Whitehouse Station, NJ: Merck and C., Inc.; 2016 Jul.

29948 – Maxaquin (lomefloxacin tablets) package insert. New York, NY: G.D. Searle LLC; 2006 May.

30106 – Imodium A-D Liquid and Caplets (loperamide HCL) consumer product labels. Fort Washington, PA: Johnson and Johnson Consumer Inc., McNeil Consumer Healthcare Division; 2019.

30163 – Agrylin (anagrelide) capsules package insert. Lexington, MA: Shire US Inc.; 2020 Feb.

30284 – McElnay JC, Mukhtar HA, D' Arcy PF, Temple DJ. In vitro experiments on chloroquine and pyrimethamine absorption in the presence of antacid constituents or kaolin. *J Trop Med Hyg* 1982;85:153-8.

30285 – McElnay JC, Mukhtar HA, D' Arcy PF, et al. The effect of magnesium trisilicate and kaolin on the in vivo absorption of chloroquine. *J Trop Med Hyg* 1982;85:159-63.

30369 – Vantas (histrelin implant) package insert. Chadds Ford, PA: Endo Pharmaceuticals Solutions Inc.; 2019 Feb.

30515 – Vesicare (solifenacin) package insert. Research Triangle Park, NC: GlaxoSmithKline; 2012 Jul.

30624 – Herrington AM, George KW, Moulds CC. Octreotide-induced bradycardia. *Pharmacotherapy* 1998;18:413-6.

30738 – Ofloxacin tablets package insert. Sacramento, CA: Nivagen Pharmaceuticals, Inc.; 2019 Feb.

31112 – Detrol (tolterodine tartrate) package insert. New York, NY: Pharmacia and Upjohn Co., division of Pfizer; 2016 Nov.

31217 – Penicillin G procaine package insert. New York, NY: Pfizer Inc; 2019 May.

31266 – Zofran (ondansetron) injection package insert. Research Triangle Park, NC: GlaxoSmithKline; 2017 Mar.

31353 – Synera (lidocaine; tetracaine) package insert. Sounderton, PA: Galen US Inc.; 2018 Nov.

31403 – Qualaquin (quinine sulfate) capsules package insert. Cranbury, NJ: Sun Pharmaceutical Industries, Inc.; 2019 Jun.

31723 – Kytril injection (granisetron) package insert. Nutley, NJ: Roche Pharmaceuticals; 2011 Nov.

31832 – Nexavar (sorafenib) package insert. Wayne, NJ; Bayer HealthCare Pharmaceuticals Inc.; 2020 March.

31938 – Ranexa (ranolazine extended-release tablets) package insert. Foster City, CA: Gilead Sciences, Inc. 2019 Oct.

32127 – Prozac (fluoxetine hydrochloride) package insert. Indianapolis, IN: Eli Lilly and Company; 2017 Mar.

32387 – Sprycel (dasatinib) package insert. Princeton, NJ: Bristol-Myers Squibb Company; 2014 April.

- 32722** – Charbit B, Albaladejo P, Funck-Brentano C, et al. Prolongation of QTc interval after postoperative nausea and vomiting treatment by droperidol or ondansetron. *Anesthesiology* 2005;102:1094-100.
- 32723** – Noxafil (posaconazole) package insert. Whitehouse Station, NJ: Merck & Co. Inc.; 2019 Mar.
- 32732** – Stollberger C, Huber JO, Finsterer J. Antipsychotic drugs and QT prolongation. *Int Clin Psychopharmacol* 2005;20:243-51.
- 32734** – Su KP, Lane HY, Chuang CL, et al. Olanzapine-induced QTc prolongation in a patient with wolff-parkinson-white syndrome. *Schizophrenia Research* 2004;66:191-2.
- 32745** – Dineen S, Withrow K, Voronovitch L, et al. QTc prolongation and high-dose olanzapine. *Psychosomatics* 2003;44:174-5.
- 32789** – Zolinza (vorinostat) capsules package insert. Whitehouse Station, NJ: Merck & Co., Inc.; 2018 Dec.
- 32827** – Tyzeka (telbivudine) package insert. East Hanover, NJ: Novartis Pharmaceuticals, Corp.; 2018 Dec.
- 33068** – Gajwani P, Pozuelo L, Tesar G, et al. QT interval prolongation associated with quetiapine (seroquel) overdose. *Psychosomatics* 2000;41:63-5.
- 33072** – Beelen AP, Yeo KTJ, Lewis LD. Asymptomatic QTc prolongation associated with quetiapine fumarate overdose in a patient being treated with risperidone. *Hum Exp Toxicol* 2001;20:215-9.
- 33074** – Furst BA, Champion KM, Pierre JM, et al. Possible association of QTc interval prolongation with co-administration of quetiapine and lovastatin. *Biol Psychiatry* 2002;51:264-5.
- 33136** – Dolophine (methadone) package insert. Eatontown, NJ: West-Ward Pharmaceuticals Corp.; 2019 Oct.
- 33192** – Tykerb (lapatinib) tablet package insert. Research Triangle Park, NC: GlaxoSmithKline; 2018 Dec.
- 33715** – Effexor XR (venlafaxine extended-release capsules) package insert. Philadelphia, PA: Wyeth Pharmaceuticals, Inc; 2017 Dec.
- 34335** – Giao PT, de Vries PJ. Pharmacokinetic interactions of antimalarial agents. *Clin Pharmacokinet* 2001;40:343-73.
- 34353** – Ducharme J, Farinotti R. Clinical pharmacokinetics and metabolism of chloroquine: focus on recent advancements. *Clin Pharmacokinet* 1996;31:257-74.
- 34389** – Xenazine (tetrabenazine) package insert. Deerfield, IL: Lundbeck, Inc.; 2017 Sep.
- 35401** – Coartem (artemether; lumefantrine) package insert. East Hanover, NJ: Novartis Pharmaceuticals Corporation; 2019 Aug.
- 36101** – Multaq (dronedarone) package insert. Bridgewater, NJ: Sanofi-aventis; 2014 Mar.

36146 – Fanapt (iloperidone) package insert. Rockville, MD: Vanda Pharmaceuticals, Inc.; 2017 Mar.

36250 – Sabril (vigabatrin) tablet/powder for oral solution package insert. Deerfield, IL: Lundbeck Inc.; 2020 Jan.

36343 – Saphris (asenapine) package insert. St. Louis, MO: Forest Pharmaceuticals, Inc.; 2017 Mar.

36615 – Vibativ (telavancin) package insert. Nashville, TN: Cumberland Pharmaceuticals Inc.; 2020 Feb.

37098 – Votrient (pazopanib) package insert. Research Triangle Park, NC: GlaxoSmithKline; 2011 Oct.

37292 – Istodax (romidepsin) package insert. Bedford, OH: Ben Venue Laboratories, Inc.; 2018 Nov.

38831 – Oleptro (trazodone hydrochloride) extended-release tablets package insert. Dublin, Ireland: Labopharm Europe Limited; 2014 Jul.

40233 – Sporanox (itraconazole) oral solution package insert. Titusville, NJ: Janssen Pharmaceuticals, Inc.; 2019 Mar.

40848 – Imovax Rabies (rabies vaccine) package insert. Swiftwater, PA: Sanofi-Pasteur Inc.; 2019 Oct.

40849 – RabAvert (rabies vaccine) package insert. Emeryville, CA: Novartis Vaccines and Diagnostics, Inc.; 2018 August.

40936 – Invega Sustenna (paliperidone palmitate injectable suspension) package insert. Titusville, NJ: Janssen Pharmaceuticals, Inc.; 2019 Jan.

40942 – Remeron and RemeronSolTabs (mirtazapine tablets and ODT tablets) package insert. Roseland, NJ: Organon USA, Inc.; 2020 Mar.

41124 – Staxyn (vardenafil orally disintegrating tablets) package insert. Research Triangle Park, NC: GlaxoSmithKline; 2017 Aug.

41231 – Foradil inhalation powder (formoterol fumarate) package insert. Kenilworth, NJ: Schering Corporation; 2012 Nov.

41235 – Butrans (buprenorphine transdermal system) package insert. Stamford, CT: Purdue Pharma L.P.; 2019 Oct.

41806 – Plaquenil (hydroxychloroquine) package insert. St. Michael, Barbados: Concordia Pharmaceuticals, Inc.; 2017 Jan.

41830 – Corvert (ibutilide) package insert. New York, NY: Pharmacia and Upjohn Company; 2016 Aug.

41958 – Eloxatin (oxaliplatin) package insert. Bridgewater, NJ: Sanofi-aventis U.S. LLC; 2020 April.

41984 – Primaquine phosphate package insert. Bridgewater, NJ: Sanofi-aventis; 2017 Jun.

42039 – Quelicin (succinylcholine) package insert. Lake Forest, IL: Hospira, Inc.; 2018 Jul.

42121 – Pradaxa (dabigatran) package insert. Ridgefield, CT: Boehringer Ingelheim Pharmaceuticals, Inc.; 2019 Nov.

42280 – Nuedexta (dextromethorphan hydrobromide; quinidine sulfate capsule) package insert. Aliso Viejo, CA: Avanir Pharmaceuticals, Inc.; 2019 Jun.

42449 – Halaven (eribulin mesylate) injection package insert. Woodcliffe Lake, NJ: Eisai Inc.; 2016 Oct.

42844 – FDA Drug Safety Communication: Abnormal heart rhythms associated with use of Anzemet (dolasetron mesylate). Retrieved December 17, 2010. Available on the World Wide Web at: <http://www.fda.gov/Drugs/DrugSafety/ucm237081.htm>.

42845 – Abilify (aripiprazole) tablets, discmelt orally-disintegrating tablets, oral solution, and intramuscular injection package insert. Tokyo, Japan: Otsuka America Pharmaceutical, Inc.; 2020 Feb.

43065 – Chlorpromazine package insert. Princeton, NJ: Sandoz Inc; 2019 Dec.

43258 – Eryped (erythromycin ethylsuccinate) package insert. Atlanta, GA: Arbor Pharmaceuticals, INC.; 2018 Apr.

43383 – Xylocaine (lidocaine injection), Xylocaine (lidocaine and epinephrine injection) package insert. Lake Zurich, IL: Fresenius Kabi; 2018 Nov.

43411 – Cipro (ciprofloxacin tablet; suspension) package insert. Wayne, NJ: Bayer HealthCare Pharmaceuticals Inc.; 2020 Mar.

43463 – Orap (pimozide) package insert. Sellersville, PA: Teva Pharmaceuticals USA; 2014 Mar.

43800 – Lupron Depot (leuprolide acetate for depot suspension) package insert. North Chicago, IL: AbbVie Inc; 2017 July.

43901 – Caprelsa (vandetanib) package insert. Wilmington, DE: AstraZeneca Pharmaceuticals LP; 2018 Oct.

43974 – Zithromax (azithromycin injection) package insert. New York, NY: Pfizer Inc.; 2019 Apr.

44026 – Advair HFA (fluticasone propionate; salmeterol inhalation aerosol) package insert. Research Triangle Park, NC: GlaxoSmithKline; 2019 Jan.

44058 – Prozac Weekly (fluoxetine hydrochloride delayed-release capsules) package insert. Indianapolis, IN: Lilly USA, LLC; 2017 Mar.

44059 – Sarafem (fluoxetine hydrochloride) package insert. Irvine, CA: Allergan USA, Inc; 2017 Jan.

44063 – Advair Diskus (fluticasone propionate; salmeterol inhalation powder) package insert. Research Triangle Park, NC: GlaxoSmithKline; 2019 Jan.

44376 – Edurant (rilpivirine) package insert. Titusville, NJ: Janssen Therapeutics; 2019 May.

44406 – Fosrenol (lanthanum carbonate chewable tablets and oral powder) package insert. Lexington, MA: Shire US Inc.; 2018 Nov.

44800 – Potiga (ezogabine) package insert. Research Triangle Park, NC: GlaxoSmithKline; 2016 May.

44979 – Arcapta Neohaler (indacaterol inhalation powder) package insert. East Hanover, NJ: Novartis Pharmaceuticals Corporation; 2019 May.

45335 – Zelboraf (vemurafenib) tablet package insert. South San Francisco, CA: Genentech USA, Inc.; 2017 Nov.

45411 – Trelstar (triptorelin pamoate for injectable suspension) package insert. Parsippany, NJ: Watson Pharma, Inc; 2018 Dec.

45458 – Xalkori (crizotinib) package insert. New York, NY: Pfizer Labs; 2019 June.

46655 – Ciprofloxacin injection (vials) package insert. Lake Forest, IL: Hospira, Inc. 2020 Jan.

46869 – Firmagon (degarelix) package insert. Parsippany, NJ: Ferring Pharmaceuticals Inc.; 2020 Dec.

47129 – Hydroxyzine hydrochloride injection package insert. Shirley, NY: American Regent, Inc.; 2016 Oct.

47221 – Propulsid (cisapride) package insert. Titusville, NJ; Janssen Pharmaceutica; 2006 Oct. NOTE: As of May 2000; Propulsid has only been available in the United States via an investigational limited access program to ensure proper patient screening and prescribing.

47357 – Quinidine gluconate extended release tablet package insert. Richmond, VA: Richmond Pharmaceuticals, Inc.; 2011 Jan.

47391 – Intron A (interferon alfa-2b, recombinant) package insert. Whitehouse Station, NJ: Merck & Co, Inc.; 2018 May.

48697 – Korlym (mifepristone) tablet package insert. Menlo Park, CA: Corcept Therapeutics Incorporated; 2017 May.

49951 – Ventolin HFA (albuterol sulfate) Inhalation Aerosol package insert. Research Triangle Park, NC: GlaxoSmithKline; 2014 Dec.

50507 – Luvox CR (fluvoxamine maleate extended-release capsules) package insert. Palo Alto, CA: Jazz Pharmaceuticals, Inc.; 2017 Jan.

51070 – Tambocor (flecainide) package insert. Scottsdale, AZ: Medicis, The Dermatology Company; 2011 Dec.

51289 – Inapsine (droperidol) package insert. Lake Forest, IL: Akorn, Inc.; 2011 Nov.

51740 – Maprotiline HCl tablet package insert. Morgantown WV: Mylan Pharmaceuticals Inc; 2014 Dec.

51793 – Xopenex (levalbuterol) inhalation solution package insert. Lake Forest, IL: Akorn, Inc.; 2019 Jan.

52330 – Naropin (ropivacaine) injection package insert. Lake Zurich, IL: Fresenius Kabi USA, LLC; 2018 Nov.

52331 – Marcaine (bupivacaine) injection package insert. Lake Forest, IL: Hospira, Inc.; 2018 Nov.

52603 – Iclusig (ponatinib) tablet package insert. Cambridge, MA: Takeda Pharmaceuticals Company Limited; 2020 Jan.

52746 – Sirturo (bedaquiline) tablet package insert. Titusville, NJ: Janssen Therapeutics; 2020 Jan.

53394 – Abilify Maintena (aripiprazole) extended-release intramuscular injection package insert. Rockville, MD:Otsuka America Pharmaceutical, Inc.; 2020 Feb.

55401 – Astagraf XL (tacrolimus extended-release capsules) package insert. Northbrook, IL: Astellas Pharma US, Inc.; 2018 Nov.

55578 – Owczuk R, Twardowski P, Dylczyk-Sommer A, et al. Influence of promethazine on cardiac repolarization: a double-blind, midazolam-controlled study. *Anaesthesia* 2009;64:609-614.

57094 – Zykadia (ceritinib) package insert. Indianapolis, IN: Novartis; 2019 March.

57377 – Altin C, Kanyilmaz S, Baysal S, et al. QT interval prolongation due to metronidazole administration. *Anadolu Kardiyol Derg* 2011;11:46-9.

57378 – Cohen O, Saar N, Swartzon M, et al. First report of metronidazole-induced QT interval prolongation. *Int J Antimicrob Agents* 2008;31:180-81.

57441 – Iribarren C, Round AD, Peng JA, et al. Validation of a population-based method to assess drug-induced alterations in QT interval: a self-controlled crossover study. *Pharmacoepidemiol Drug Saf* 2013;22:1222-32.

57803 – Cerdelga (eliglustat) capsules. Waterford, Ireland: Genzyme Ireland, Ltd.;2018 Sept.

58685 – Savaysa (edoxaban) package insert. Basking Ridge, NJ: Daiichi Sankyo, Inc.; 2019 Aug.

58766 – Tassigna (nilotinib) capsules package insert. East Hanover, NJ: Novartis Pharmaceuticals Corporation; 2019 Sept.

58782 – Lenvima (lenvatinib) package insert. Woodcliff Lake, NJ:Eisai Inc; 2020 Feb.

58821 – Farydak (panobinostat) capsules package insert. East Hanover, NJ: Novartis Pharmaceuticals Corporation; 2016 June.

59321 – CredibleMeds. QT drug lists. Available on the World Wide Web at <http://www.crediblemeds.org>.

59322 – Howes LG. Cardiovascular effects of drugs used to treat alzheimer's disease. *Drug Saf.* 2014;37:391–395.

59809 – Mamiya K, Sadanaga T, Sekita A, et al. Lithium concentration correlates with QTc in patients with psychosis. *J Electrocardiol* 2005;38:148-51.

- 59810** – van Noord C, Straus SM, Sturkenboom MC, et al. Psychotropic drugs associated with corrected QT interval prolongation. *J Clin Psychopharmacol* 2009;29:9-15.
- 59811** – Altinbas K, Guloksuz S, Caglar IM, et al. Electrocardiography changes in bipolar patients during long-term lithium monotherapy. *Gen Hosp Psychiatry* 2014;36:694-7.
- 60196** – Aristada (aripiprazole lauroxil) extended-release intramuscular suspension package insert. Waltham, MA: Alkermes, Inc.; 2017 Jun.
- 60270** – Belbuca (buprenorphine) buccal film package insert. BioDelivery Sciences International, Inc.: Raleigh, NC; 2019 Oct.
- 60281** – Cotellic (cobimetinib) tablets package insert. San Francisco, CA: Genentech USA, Inc; 2016 June.
- 60297** – Tagrisso (osimertinib) tablet package insert. Wilmington, DE: AstraZeneca Pharmaceuticals LP; 2018 Aug.
- 60372** – Mekinist (trametinib) tablets package insert. East Hanover, NJ: Novartis Pharmaceuticals Corporation; 2020 Apr.
- 60497** – Envarsus XR (tacrolimus) extended-release tablets. Cary, NC: Veloxis Pharmaceuticals, Inc.; 2018 Dec.
- 60612** – Aczone (dapsone gel 7.5%) package insert. Exton, PA: Almirall, LLC; 2019 Sept.
- 60686** – Fluconazole injection package insert. New York, NY: Pfizer; 2014 Jun.
- 60748** – Nuplazid (pimavanserin) package insert. San Diego, CA: Acadia; 2019 Sep.
- 60864** – US Food and Drug Administration (FDA). FDA Drug Safety Communication: FDA warns about serious heart problems with high doses of the antidiarrheal medicine loperamide (Imodium), including from abuse and misuse. Retrieved June 7, 2016. Available on the World Wide Web at: http://www.fda.gov/Drugs/DrugSafety/ucm504617.htm?source=govdelivery&utm_medium=email&utm_source=govdelivery
- 60957** – Avloclor (chloroquine phosphate) tablets. Summary of product characteristics. European package labeling. Avonbridge House, Bath Road, Chippenham, Wiltshire, SN15 2BB United Kingdom: Alliance Pharmaceuticals; 2016 Jun. Accessed: July 9 , 2016. Available at: <http://www.medicines.org.uk/emc/medicine/2272>
- 61195** – Levofloxacin injection package insert. North Brunswick, NJ: Claris Injectables Ltd.; 2018 Jun.
- 61470** – Vistaril (hydroxyzine pamoate) package insert. New York, NY: Pfizer Labs; 2016 Nov.
- 61759** – Ette EI, Brown-Awala EA, Essien EE. Chloroquine elimination in humans: effect of low-dose cimetidine. *J Clin Pharmacol* 1987;27:813-16.
- 61760** – Furst DE. Pharmacokinetics of hydroxychloroquine and chloroquine during treatment of rheumatic diseases. *Lupus* 1996;5(S1):S11-15.
- 61761** – Ali HM. Reduced ampicillin bioavailability following oral coadministration with chloroquine. *J Antimicrob Chemother* 1985;15:781-4.

61816 – Kisqali (ribociclib) tablets package insert. East Hanover, NJ: Novartis Pharmaceuticals Corporation; 2020 Jan.

61845 – Austedo (deutetrabenazine) tablets package insert. North Wales, PA: Teva Pharmaceuticals USA, Inc.; 2019 Jul.

61872 – Slovacek L, Ansorgova V, Macingova Z, et al. Tamoxifen-induced QT interval prolongation. *J Clin Pharm Ther.* 2008;33(4):453-5.

61906 – Rydapt (midostaurin) capsule package insert. East Hanover, NJ: Novartis Pharmaceuticals Corporation; 2020 Mar.

62037 – Bevyxxa (betrixaban) capsules package insert. South San Francisco, Ca: Portola Pharmaceuticals, Inc.; 2019 Jul.

62245 – Besponsa (inotuzumab ozogamicin) injection package insert. Philadelphia, PA: Wyeth Pharmaceuticals Inc; 2017 Aug.

62292 – Mylotarg (gemtuzumab ozogamicin) injection package insert. Philadelphia, PA: Pfizer Inc.; 2020 Feb.

62723 – Macrilen (macimorelin) package insert. Frankfurt am Main, Germany: Aeterna Zentaris GmbH; 2018 Jan.

63161 – Lucemyra (lofexidine) tablets package insert. Louisville, KY: US WorldMeds, LLC; 2018 May.

63317 – Braftovi (encorafenib) capsules package insert. Boulder, CO: Array BioPharma Inc.; 2020 April.

63589 – Soltamox (tamoxifen) oral solution package insert. Raleigh, NC: Midatech Pharma US Inc.; 2019 April.

63609 – Desyrel (trazodone) package insert. Locust Valley, NY: Pragma Pharmaceuticals, LLC.; 2017 June.

63729 – Levofloxacin premix injection package insert. Lake Forest, IL: Hospira; 2019 Jan.

63777 – Daurismo (glasdegib) tablets package insert. New York, NY: Pfizer Labs; 2020 Mar.

63787 – Xospata (gilteritinib) tablets package insert. Northbrook, IL: Astellas Pharma US, Inc.; 2019 May.

63821 – Tolsura (itraconazole) capsules package insert. Greenville, NC: Mayne Pharma; 2018 Dec.

63936 – Lamprene (clofazimine) package insert. East Hanover, NJ: Novartis Pharmaceuticals Corporation; 2019 Jan.

64031 – Mayzent (siponimod) tablets package insert. East Hanover, NJ: Novartis Pharmaceutical Corporation; 2019 Mar.

64391 – Beach SR, Kostis WJ, Celano CM, et al. Meta-analysis of selective serotonin reuptake inhibitor-associated QTc prolongation. *J Clin Psychiatry* 2014;75:e441-e449.

64392 – Glassman AH, O'Connor CM, Califf RM, et al. Sertraline treatment of major depression in patients with acute MI or unstable angina. *JAMA* 2002;288:701-709.

64394 – O'Connor CM, Jiang W, Kuchibhatla M, et al. Safety and efficacy of sertraline for depression in patients with heart failure: results of the SADHART-CHF (Sertraline Against Depression and Heart Disease in Chronic Heart Failure) trial. *J Am Coll Cardiol* 2010;56:692-699.

64396 – Beach SR, Celano CM, Sugrue AM, et al. QT prolongation, torsades de pointes, and psychotropic medications: a 5-year update. *Psychosomatics* 2018;59:105-122.

64562 – Wakix (pitolisant) tablets package insert. Plymouth Meeting, PA: Harmony Biosciences, LLC; 2019 Aug.

64567 – Rozlytrek (entrectinib) package insert. South San Francisco, CA: Genentech Inc.; 2019 Aug.

64576 – Xenleta (lefamulin) package insert. Dublin, Ireland: Nabriva Therapeutics US, Inc.; 2019 Aug.

64969 – Bynfezia Pen (octreotide acetate) injection package insert. Cranbury, NJ: Sun Pharmaceutical Industries, Inc; 2020 Jan.

65098 – Isturisa (osilodrostat) tablet package insert. Lebanon, NJ: Recordati Rare Disease, Inc.; 2020 Mar.

65157 – CredibleMeds. COVID-19 experimental therapies and TdP risk. Available on the World Wide Web at <https://crediblemeds.org/blog/covid-19-experimental-therapies-and-tdp-risk>. Accessed March 23, 2020.

65170 – Giudicessi JR, Noseworthy PA, Friedman PA, Ackerman MJ. Urgent guidance for navigating and circumventing the QTc prolonging and torsadogenic potential of possible pharmacotherapies for COVID-19 [published online ahead of print, March 25, 2020]. *Mayo Clin Proc* 2020;95.

65210 – University of Liverpool. COVID-19 Drug Interactions. Retrieved April 6, 2020. Available on the World Wide Web at <https://www.covid19-druginteractions.org/>.

65237 – Tiberghien F, Loo F. Ranking of P-glycoprotein substrates and inhibitors by a calcein-AM fluorometry screening assay. *Anti-Cancer Drugs* 1996;7:568-578.

65239 – Projean D, Baune B, Farinotti R, et al. In vitro metabolism of chloroquine: Identification of CYP2C8, CYP3A4, and CYP2D6 as the main isoforms catalyzing N-desethylchloroquine formation. *Drug Metab Dispos* 2003;31:748-754.


Monitoring Parameters


- blood glucose
- CBC
- ECG
- glucose-6-phosphate dehydrogenase (G6PD) activity
- ophthalmologic exam

IV Compatibility of Chloroquine with:

Legend




 = Compatible

 = Incompatible

 = Results uncertain, variable or dependent on conditions

ND = No Data Available

From Trissel's 2TM Clinical Pharmaceutics Database 

	Admixture	Syringe	Y-Site Administration	For Dilution
Normal saline- Sodium chloride 0.9%	ND	ND	ND	
Promethazine hydrochloride			ND	ND

Copyright 2004-2015 by Lawrence A. Trissel. All Rights Reserved. 

US Drug Names

- Aralen

Global Drug names

Argentina

- Nivaquine - (Sanofi-Aventis)

Australia

- Chlorquin - (Aspen)
- Nivaquine - (Rhone-Poulenc Rorer)

Austria

- Resochin - (Bayer)

Belgium

- Nivaquine - (Sanofi)

Brazil

- Clopirim - (Quimioterapica)
- Diclokin - (Kinder)
- Difosquin - (Vitamed)
- Palux - (Biolab Sanus)
- Quinacris - (Cristalia)

Canada

- Aralen - (Sanofi Synthelabo)

Czech Republic

- Delagil - (ICN)

Denmark

- Malarex - (Actavis)

Finland

- Heliopar - (Orion)

France

- Nivaquine - (Sanofi-Aventis)
- Nopalu - (Pharmacie Centrale des Armees)
- Savarine - (AstraZeneca)

Germany

- Arthrabas - (Tosse)
- Resochin - (Bayer)
- Weimerquin - (Biokanol)

Greece

- Avloclor - (IFET)
- Demoquine - (Demo)
- Savarine - (IFET (IΦET))

Hong Kong

- Chlorocin - (Deltapharm)
- Chlorquin - (Fisons)
- Syncoquin - (Synco)

Hungary

- Delagil - (PharmaSwiss)

India

- Bitaquine - (Bombay Tablet)
- Cadiquin - (Zydus)
- Chlorolex - (Lexica)
- Clokit - (Indoco)
- Cloquin - (Indoco)
- C-Quin - (Ikon)
- Emquin - (Merck)
- E-Vivax - (Themis Medicare)
- Idiquin - (Indian Drugs)
- Ingaquine - (Inga)
- Jagquin - (Jagsonpal)

- La-Quin - (Stadmed)
- Lariago - (Ipca)
- Larover - (Aglowmed)
- Malaquin - (PC India)
- Maliago - (Cipla)
- Maligon - (Unijules)
- Malswift - (Ind-Swift)
- Melubrin - (Ranbaxy)
- Neoquine - (Neon)
- Nivaquine-P - (Piramal)
- Paraquin - (Shreya)
- Resochin - (Bayer)

Indonesia

- Avloclor - (AstraZeneca)
- Malarex - (Actavis)
- Mexaquin - (Konimex)
- Resochin - (Bayer)
- Riboquin - (Dexa)

Ireland

- Avloclor - (AstraZeneca)
- Nivaquine - (Rhone-Poulenc Rorer)

Israel

- Aralen - (Sanofi Winthrop)
- Avloclor - (Zeneca)

Italy

- Dichinalex - (Recordati)

Mexico

- Aralen - (Sanofi-Aventis)
- Klorokin - (Zerboni)
- Maclorex - (Alpharma)
- Paluken - (Kener)

Netherlands

- Nivaquine - (Sanofi-Aventis)

New Zealand

- Chlorquin - (Healthcare Logistics)
- Nivaquine - (Aventis)

Philippines

- Aralen - (Sanofi Synthelabo)
- Chlorofoz - (Am-Europharma)

- Chloromax - (Oboi)
- Clorkin - (Doctors)

Poland

- Arechin - (Polfa Pabianice)

Portugal

- Resochina - (BayHealth)

Russian Federation

- Delagil - (ICN)

Singapore

- Avloclor - (AstraZeneca)
- Malarex - (Danone Dumex)

South Africa

- Anoclor - (Rolab)
- Daraclor - (Glaxo Wellcome)
- Daramal - (GSK)
- Daramal-Paludrine - (Zeneca)
- Mirquin - (Mirren)
- Nivaquine - (Winthrop)
- Plasmoquine - (Medchem)
- Promal - (Propan)

Spain

- Cidanchin - (Cidan)
- Resochin - (Kern)

Switzerland

- Chlorochin - (Streuli)
- Nivaquine - (Sanofi-Aventis)
- Pharmaquine - (Pharma Plus)
- Resochine - (Bayer)

Thailand

- Chewoquine - (Chew)
- Diroquine - (Atlantic)
- Genocin - (General Drugs)
- Malacin - (ANB)
- Maliaquine - (Sriprasit)
- Nitaquin - (Utopian)
- P-Roquine - (PP Lab)
- Sinmoquin - (SSP)

Turkey

- Kutlu - (Keymen)

Ukraine

- Delagil - (Meda)

United Kingdom

- Avloclor - (Alliance)
- Malarivon - (Wallace Mfg Chem.)
- Malaviron - (Wallace Mfg Chem.)
- Nivaquine - (Sanofi-Aventis)