

Proposed Treatment for COVID-19

Azithromycin ^[1]

Enviado por Frank Silva ^[2] el 28 marzo 2020 - 12:29pm



^[2]

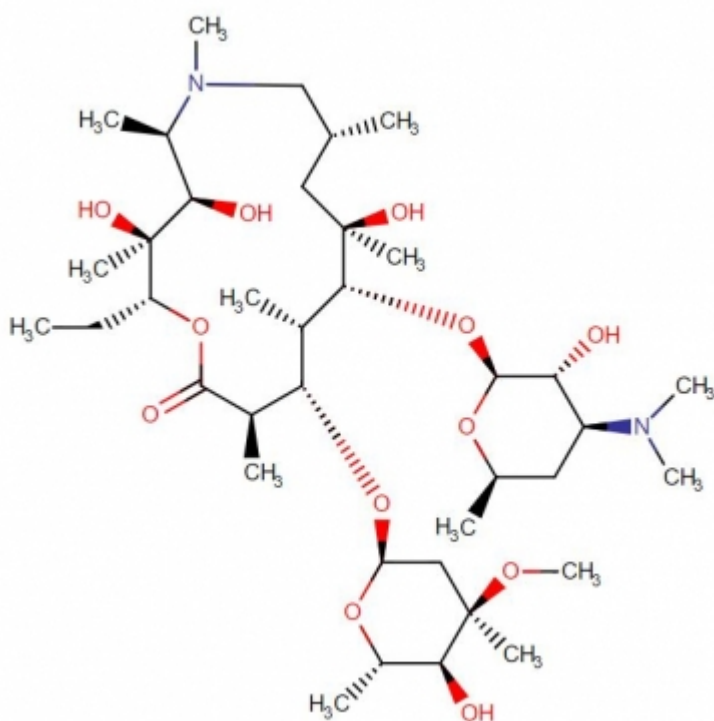


Photo by: Canadian Drug Bank

Molecular Formula: $C_{38}H_{72}N_2O_{12}$

Disclosure:

The following article is with the intent to inform the community about azithromycin and not to be used for medical purposes. Please seek medical attention if you think you have flu or flu-like symptoms before ingesting antibiotics.

Introduction:

Azithromycin is part of a family of antibiotics called macrolides. It is an azalide antibiotic.

The first macrolide was called Erythromycin discovered in 1949 at Eli Lilly by McGuire who isolated erythromycin, from the metabolic products of a strain of the actinomycete *Saccharopolyspora erthraea*, formerly known as *Streptomyces erythreus*, found in soil samples collected from the Philippines by the Filipino Scientist Abelardo Aguilar.

Azithromycin was discovered by a team of Croatian scientists at Pliva in the 1980s and marketed 8 years later in Central and Eastern Europe under the brand name **Sumamed**. In 1991, azithromycin gained FDA approval and was brought to the US market by Pfizer under the brand name of Zithromax.

Indication:

The indications for these antibiotics is for Gram-positive bacteria (cocci and bacilli).

For out-patient therapy of CAP (Community Acquired Pneumonia, pharyngitis or skin-structured infections, also used for gastrointestinal infection, and STDs.

Dosage:

The Loading dose is 500 mg, given on the first day and then 250 mg on day 2-5. This is a general statement, dosing varies according to medical criteria and on the disease, the patient is presenting.

Mechanism of Action:

Macrolides are bacteriostatic agents that inhibit protein synthesis by binding reversible to 50S ribosomal subunit of sensitive microorganisms, inhibit the translocation step, wherein a new synthesized peptidyl tRNA molecule moves from the receptor site on the ribosome to the peptidyl donor site.

Pharmacokinetics:

Distribution: The drug is widely distributed and rapidly absorbed and there is no blood-brain barrier distribution.

Elimination: The drug is non-renal.

Untoward Effects:

Cholestasis Hepatitis

-GI Toxicity

-Prolonged QT Syndrome

-Transient Auditory Impairment

Drug-Drug Interactions:

(Parent Drug) Erythromycin inhibits the cytochrome enzymes, in this case, the antibiotic inhibits the CYP 3A4, therefore patients taking the following medication should be aware of these effects:

1. Carbamazepine
2. Corticosteroids
3. Cyclosporine
4. Digoxin
5. Theophylline
6. Warfarin

Azithromycin has perhaps the most benign safety profile of the currently marketed macrolide agents. Indeed, only 0.7% of subjects receiving azithromycin discontinued therapy due to adverse events, compared with 2.6% of subjects receiving comparator drugs. (1)

Azithromycin is primarily eliminated unchanged and does not interact with the cytochrome P450 system.

Interestingly, there still are anecdotal reports of toxicity when azithromycin is used with lovastatin, warfarin, cyclosporine, disopyramide, and theophylline.

Drug-Vitamin Interaction

Vitamin B1 and B12: Although in short-term antibiotic therapy it is not a significant effect, in long-term antibiotic therapy, calcium supplementation may be needed (as well as folic acid, vitamin B12)

Clinical Trial and COVID-19

Clinical Trials that are recruiting participants are

Safety and Efficacy of Hydroxychloroquine Associated With Azithromycin in SARS-Cov-2 Virus (Coalition-I)

Description of Trial: COALITION I study aims to a compared standard of care,hydroxychloroquine plus azithromycin and hydroxychloroquine monotherapy for the treatment of hospitalized patients with COVID-19.

COALITION I will recruit 630 patients with infection by COVID-19 (210 per arm). The ordinal endpoint of status at 15 days will be the primary endpoint.

- I. Experimental Arm: Hydroxychloroquine after randomization, Hydroxychloroquine [400mg 2x/day, 12/12h] for **07 days**.

- II. Experimental Arm: Hydroxychloroquine + azithromycin After randomization, Hydroxychloroquine [400mg 2x/day, 12/12h] + azithromycin[500mg 1x/day] for **07 days**.
- III. No intervention: Control, Standard Treatment Protocol for 2019-nCoV infection.

Clinical Trial is set to end August 30, 2020

Safety and Efficacy of Hydroxychloroquine Associated with Azithromycin in SARS-CoV2 Virus (Alliance Covid-19 Brasil II)

This clinical trial will start recruiting 440 patients starting this next Monday, March 25, 2020. The responsible party is Hospital Israelita Albert Einstein and is a phase 3 clinical trial. Its description is will evaluate the effectiveness and safety of the use of hydroxychloroquine combined with azithromycin compared to hydroxychloroquine monotherapy in the clinical evolution by the ordinal scale of 7 points in adult patients hospitalized with pneumonia caused by infection by the SARS-CoV2 virus in Brazil.

1. Intervention Group: Hydroxychloroquine + azithromycin. After randomization, Hydroxychloroquine [400mg 2x/day, 12/12h] + azithromycin [500mg 1x/day] for **10 days**. Standard treatment is according to the treatment protocol for 2019-nCoV infection.
2. Active Control Group: Hydroxychloroquine. After randomization, Hydroxychloroquine [400mg 2x/day, 12/12h] for **10 days**. Standard treatment is according to the treatment protocol for 2019-nCoV infection.

Clinical Trial will end on August 30, 2020.

Proactive Prophylaxis With Azithromycin and Chloroquine in Hospitalized Patients With COVID-19 (ProPAC-COVID).

This clinical study will take place in Denmark it is a phase 4 clinical trial that will start recruitment of 226 participants in a randomized and the trial is set to start on April 1, 2020, and will end on March 31, 2020. The aim of the study is to investigate whether the treatment can shorten hospitalization, reduce the risk of non-invasive ventilation, admittance to Intensive Care Units and death.

The third clinical trial that involved Azithromycin:

Anti-Coronavirus Therapies to Prevent Progression of Coronavirus Disease 2019 (COVID-19) Trial (ACT COVID19)

This clinical trial is taking place in Canada by the Population Health Research Institute. Its description is that consists of two parallel trials evaluating azithromycin and chloroquine therapy (ACT) versus usual care in outpatients and inpatients who have tested positive for COVID-19. The trial is an open-label, parallel-group, randomized controlled trial with an adaptive design. Adaptive design features include adaptive intervention arms and adaptive sample size based on new and emerging data, it is to expected to recruit 1500 participants. It is set to start April 1, 2020, and end on December 31, 2020.

There is very promising clinical trial to start this week, but the majority of them have a timeline of 6 to 12 months. After clinical trials end, there is a time to collect and analyze correctly the data of the trial before its use.

Currently, Azithromycin is not labeled for use against COVID-19, so for people not diagnosed with COVID-19, this is not prophylaxis.

We must be careful with the information published over the internet, not to promote like with Plaquenil, the usage of medication not labeled for COVID-19 as if it were labeled for, and create a shortage of these medications for people who have conditions for which the medication is labeled.

Disclosure:

The following article is with the intent to inform the community about azithromycin and not to be used for medical purposes. Please seek medical attention if you think you have flu or flu-like symptoms before ingesting antibiotics.

References:

Amsden GW (1995) Macrolides vs azalides: a drug interaction update. *Ann Pharmacother* 29: 906-917

Bellosta S, Paoletti R, Corsini A. (2004) Safety of statins: focus on pharmacokinetics and drug interactions. *Circulation* 109 (23 Suppl1): III50-57

Chen, X., & Yu, B. (2020). First two months of the 2019 Coronavirus Disease (COVID-19) epidemic in China: Real-time surveillance and evaluation with a second derivative model. *Global Health Research and Policy*, 5(1), 7.

Dougherty, T. J., & Pucci, M. J. (2011). *Antibiotic Discovery and Development*. Springer Science & Business Media.

Granow EV, Tabor KJ, Kirchhoffer JB (2000) Potentially fatal interaction between azithromycin and disopyramide. *Pacing Clin Electrophysiol* 23: 1433-1435

Gautret P, Lagier JC, Parola P, et al. Hydroxychloroquine and azithromycin as a treatment of

COVID-19: results of an open-label non-randomized clinical trial [published online ahead of print, 2020 Mar 20]. *Int J Antimicrob Agents*. 2020;105949. doi:10.1016/j.ijantimicag.2020.105949

Hageman, J. (2020). The Coronavirus Disease 2019 (COVID-19). *Pediatric Annals*, 49(3), E99-E100.

Hopkins S (1991) Clinical toleration and safety of azithromycin *Am J Med* 91: 40S-45S

Hopkins S (1994) Clinical toleration and safety of azithromycin in adults and children. *Rev Contemp Pharmacother* 5:383-389

Peters DH, Friedel HA, McTavish D (1992) Azithromycin A review of its antimicrobial activity, pharmacokinetic properties and clinical efficacy. *Drugs* 44: 750-799.

Page RL 2nd, Ruscin JM, Fish F, Lapointe M (2001) Possible interaction between intravenous azithromycin and oral cyclosporine. *Pharmacotherapy* 21: 1436-1443.

Pollak PT, Slayter KL (1997) Reduced serum theophylline concentrations after discontinuation of azithromycin: evidence for an unusual interaction. *Pharmacotherapy* 17:827-829

Sarkar, B., Ullah, M., & Araf, Y. (2020). The Essential Facts of Wuhan Novel Coronavirus Outbreak in China and Epitope-based Vaccine Designing against COVID-19. *BioRxiv*, *BioRxiv*, Mar 6, 2020.

Shrader SP, Fermo JD, Dzikowski AL (2004) Azithromycin and warfarin interaction. *Pharmacotherapy* 24: 945-949

Wang D, Hu B, Hu C, et al. Clinical Characteristics of 138 Hospitalized Patients With 2019 Novel Coronavirus-Infected Pneumonia in Wuhan, China [published online ahead of print, 2020 Feb 7]. *JAMA*. 2020;e201585. doi:10.1001/jama.2020.1585

Yan-Rong Guo, Qing-Dong Cao, Zhong-Si Hong, Yuan-Yang Tan, Shou-Deng Chen, Hong-Jun Jin, . . . Yan Yan. (2020). The origin, transmission and clinical therapies on coronavirus disease 2019 (COVID-19) outbreak – an update on the status. *Military Medical Research*, 7(1), 1-10.

Tags:

- [COVID-19](#) ^[3]
- [Azythromycin](#) ^[4]

Copyright © 2006-Presente CienciaPR y CAPRI, excepto donde sea indicado lo contrario, todos los derechos reservados

[Privacidad](#) | [Términos](#) | [Sobre CienciaPR](#) | [Contáctenos](#)

Source URL: <https://www.cienciapr.org/es/blogs/members/proposed-treatment-covid-19-azithromycin>

Links

- [1] <https://www.cienciapr.org/es/blogs/members/proposed-treatment-covid-19-azithromycin>
- [2] <https://www.cienciapr.org/es/user/frank-valentin-silva>
- [3] <https://www.cienciapr.org/es/tags/covid-19>
- [4] <https://www.cienciapr.org/es/tags/azythromycin>