WHEN MEDICATION FOR MINOR PAIN ISN'T WORKING RECOMMEND COMBOGESIC®.

For mild to moderate acute pain. Superior efficacy. Comparable safety.¹



COMBOGESIC[®] (acetaminophen 325 mg + ibuprofen 97.5 mg) is indicated in adults over 18 years for the short-term management of mild to moderate acute pain and the reduction of fever.



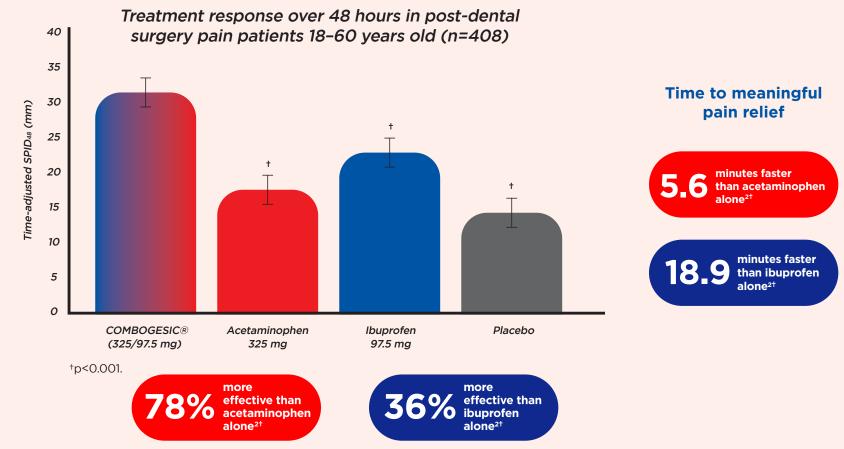
COMBOGESIC® A proven ratio

COMBOGESIC® offers a proven 325 mg to 97.5 mg ratio of acetaminophen to ibuprofen.

Sold in 46 countries

Combogesic^{*}

COMBOGESIC® showed a significantly superior treatment response compared to acetaminophen or ibuprofen alone ^{1,2,*}



SPID: Sum of Pain Intensity Differences.

*Prospective, multicentre, randomized, double-blind, placebo-controlled, Phase III trial of 408 adult volunteers aged 18 to 60 years experiencing moderate to severe pain after surgical removal of at least 2 impacted third molars, using 3 pills of a fixed-dose combination of acetaminophen 325 mg/ibuprofen 97.5 mg (acetaminophen 975 mg/ ibuprofen 292.5 mg each dose), compared with acetaminophen 975 mg or ibuprofen 292.5 mg alone 4 times a day (acetaminophen 3,900 mg or ibuprofen 1,170 mg per day).¹²

Combogesic[®]

COMBOGESIC® Simple, clear, flexible dosing



The maximum allowable daily dose for adults is 4,000 mg of acetaminophen and 1,200 mg of ibuprofen.

Getting to the maximum allowable daily dose:

Product	Tablets	<u>X / Day</u>	Dosing
Combogesic [®]	3	4 x	3,900 mg, 1,170 mg
325 mg acet.	2	6x	4,000 mg
200 mg ibu.	2	3x	1,200 mg

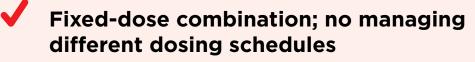
*3 tablets on subsequent doses on the advise of a physician if pain or fever does not respond to 2.

For oral administration and short-term use only. Do not take for fever for more than 3 days or pain for more than 5 days unless directed by a physician. Do not take COMBOGESIC[®] tablets with other acetaminophen- or ibuprofen-containing products. Take with food or milk if stomach upset occurs.

Combogesic[®]

Acet. = acetaminophen. Ibu. = ibuprofen.

COMBOGESIC® For more than just minor pain



Superior efficacy

For acute pain including headaches, back pain, strains and dental pain





Combogesic^{*}

Serious warnings and precautions

NSAIDs

Risk of cardiovascular (CV) adverse events:

- Use of some NSAIDs is associated with an increased incidence of cardiovascular adverse events, which can be fatal.
- Caution should be exercised in prescribing COMBOGES-IC[®] to any patient with ischemic heart disease (including but NOT limited to acute myocardial infarction, history of myocardial infarction and/or angina), cerebrovascular disease (including but NOT limited to stroke, cerebrovascular accident, transient ischemic attacks and/or amaurosis fugax) and/or congestive heart failure.
- Use of medicines such as COMBOGESIC[®] can promote sodium retention in a dose dependent manner, through a renal mechanism, which can result in increased blood pressure and/or exacerbation of congestive heart failure.

Risk of gastrointestinal (GI) adverse events:

- Use of medicines such as COMBOGESIC® is associated with an increased incidence of gastrointestinal adverse events.
- The chance of stomach bleeding is higher if the patient: $_{0}$ is aged 60 or older
 - ^o o takes a blood thinner or steroid drug
 - o takes with other drugs containing an NSAID like acetylsalicylic acid (ASA), ibuprofen, naproxen, or prescription anti-infl ammatory drugs
 - o has 3 or more alcoholic drinks every day while using this product.

Acetaminophen

LIVER WARNING:

- Acetaminophen has been associated with cases of acute liver failure, at times resulting in liver transplant and death.
- Most of the cases of liver injury are at doses that exceed 4,000 mg per day, and often involve more than one acetaminophen containing product.
- Severe or possibly fatal liver damage may occur if the patient takes:
 - ^o more than the recommended dose in 24 hours
 - ^o with other drugs containing acetaminophen
 - while drinking three (3) or more alcoholic drinks every day.
- Symptoms of liver damage may include yellowing of the skin/eyes.

Serious skin reactions:

• Acetamin gohen may cause serious skinreactions.

Other:

• COMBOGESIC[®] use during pregnancy should be avoided.

For more information:

Please consult the Product Monograph at

https://pdf.hres.ca/dpd_pm/00054759.PDF for

imporant information regarding contraindications and other warnings and precautions, adverse reactions, drug interactions and dosing. The Product Monograph is also available by calling BioSyent Pharma Inc. at 1-888-439-0013 or at **combogesic.ca**.

References: 1. COMBOGESIC® Product Monograph. BioSyent Pharma Inc. 2. Daniels SE, et al. Analgesic

efficacy of an acetaminophen/ibuprofen fixed-dose combination in modérate.to.cevera postoperative dental pain: a randomized, double-blind, parallel-group, placebo-controlled trial. Cli va78 78 ot 8;40(10):1765-76.

Combogesic[®]