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Name of Journal: *World Journal of Hepatology*

Manuscript NO: 58287

Manuscript Type: ORIGINAL ARTICLE

Basic Study

Obeticholic acid attenuates human immunodeficiency virus/alcohol metabolism-induced pro-fibrotic activation in liver cells

New-Aaron M *et al.* Alcohol, HIV infection and obeticholic acid

10

Moses New-Aaron, Murali Ganesan, Raghubendra Singh Dagur, Kusum K Kharbanda, Larisa Y Poluektova, Natalia A Osna

Abstract

BACKGROUND

The morbidity and mortality of human immunodeficiency virus (HIV)-infection is often associated with liver disease, which progresses slowly into severe liver dysfunction.



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Antifibrotic Therapies in the Liver

<https://www.ncbi.nlm.nih.gov/pmc/articles/PMC5743222>

Inhibition of the cannabinoid receptor 1 (CB1) reverses myofibroblast **activation** and **attenuates** experimental **liver** fibrosis. 109 This has passed the proof of principle state, and peripheral-acting CB1 antagonists that may circumvent adverse side effects on the central nervous system like depression are being developed. 110 In fibrotic NASH ...

Cited by: 71

Author: W. Z. Mehal, D. Schuppan, D. Schuppan

Publish Year: 2015

Obeticholic acid prevents carbon tetrachloride-induced ...

<https://www.sciencedirect.com/science/article/pii/S1567576919315474>

Dec 01, 2019 · FXR agonist **obeticholic acid** prevents carbon tetrachloride-induced **liver** fibrosis. • **Obeticholic acid attenuates** carbon tetrachloride-induced Smad3-mediated **activation** of hepatic stellate **cells**. • **Obeticholic acid** increases interaction between farnesoid X receptor and Smad3.

Cited by: 2

Author: Yuan-Yuan Fan, Wen Ding, Cheng Zhan...

Publish Year: 2019

Strategies and Endpoints of Antifibrotic Drug Trials

<https://www.ncbi.nlm.nih.gov/pmc/articles/PMC4515973>

There are also other potential sources, such as **mesothelial cells** near the liver capsule . Non-fibrogenic cells that initiate signals directing **MF function** may also be valuable cellular targets for therapy. For example, cholangiocytes O'Hara, 2013 #26084} secrete a variety of **pro-fibrotic signaling molecules**, and hepatocytes may **promote liver fibrosis** via release of apoptotic ...

Cited by: 30

Author: Natalie J. Torok, Jonathan A. Dranoff, D...

Publish Year: 2015

Frontiers | Targeting Myeloid-Derived Cells: New Frontiers ...

<https://www.frontiersin.org/articles/10.3389/fimmu.2019.00563> ▼

Mar 27, 2019 · The different mechanisms by which these cells can be targeted include **modulation**



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Obeticholic acid prevents carbon tetrachloride-induced ...

<https://www.sciencedirect.com/science/article/pii/S1567576919315474>

Dec 01, 2019 · 1. Introduction. Liver cirrhosis is currently ranked 11th among the most common causes of death in the world, and is an important reason for disability-adjusted life years and years of life loss, bringing a huge disease burden to the world [1,2].The cause of liver cirrhosis is due to sustained liver injury induced by different factors, such as alcohol abuse, nonalcoholic steatohepatitis (NASH ...

Cited by: 2 Author: Yuan-Yuan Fan, Wen Ding, Cheng Zhang, Li...
Publish Year: 2019

Obeticholic acid alleviate lipopolysaccharide-induced ...

<https://www.sciencedirect.com/science/article/pii/S1567576918310671>

Jan 01, 2019 · Obeticholic acid attenuates LPS-induced inflammation in the lungs. ... FXR is highly expressed in human and rodent animal lungs [17,18 ... W.J. You, B. Liu, H.D. JiangBile acids induce activation of alveolar epithelial cells and lung fibroblasts through farnesoid X receptor-dependent and independent pathways. Respiriology, 21 (6) (2016), pp ...

Cited by: 13 Author: Jun Fei, Lin Fu, Biao Hu, Yuan-Hua Chen, H...
Publish Year: 2019

Obeticholic acid for the treatment of non-alcoholic ...

<https://www.researchgate.net/publication/337796448...>

Dec 01, 2019 · The obeticholic acid is the first in a class of FXR ligands that has reached a clinical stage [40][41][42] and was originally approved in 2016 as a second-line treatment for primary biliary ...

Cells | Free Full-Text | Liver Fibrosis: Mechanistic ...

<https://www.mdpi.com/2073-4409/9/4/875/htm>

Liver fibrosis due to viral or metabolic chronic liver diseases is a major challenge of global health. Correlating with liver disease progression, fibrosis is a key factor for liver disease outcome and risk of hepatocellular carcinoma (HCC). Despite different mechanism of primary liver injury and disease-specific cell responses, the progression of fibrotic liver disease follows shared patterns ...

Cited by: 7 Author: Natascha Roehlen, Emilie Crouchet, Thoma...
Publish Year: 2020

[PDF] Betulinic acid alleviates endoplasmic reticulum stressâ ...

<https://bpspubs.onlinelibrary.wiley.com/doi/pdf/10.1111/bph.14570>

severe liver steatosis (Kong, Luyendyk, Tawfik, & Guo, 2009; Sinal et al., 2000). Obeticholic acid (OCA), a well-known FXR agonist, has been the most promising candidate to treat NAFLD (Musso, Cassader, & Gambino, 2016). Thus, screening for potent selective FXR activators has been the focus of research in the anti-NAFLD drug development

Cited by: 4 Author: Ming Gu, Ping Zhao, Shiyong Zhang, Shengji...
Publish Year: 2019

Farnesoid X Receptor Agonist as a new treatment option for ...

<https://www.researchgate.net/publication/323169673...>

The bile acid analogue obeticholic acid (OCA) is a selective farnesoid X receptor (FXR) agonist in development for treatment of several chronic liver diseases. FXR activation regulates lipoprotein ...

Betulinic acid alleviates endoplasmic reticulum stress ...

<https://bpspubs.onlinelibrary.wiley.com/doi/10.1111/bph.14570>

Jan 11, 2019 · Obeticholic acid (OCA), a well-known FXR agonist, has been the most promising candidate to treat NAFLD (Musso, Cassader, & Gambino, 2016). Thus, screening for potent selective FXR activators has been the focus of research in the anti-NAFLD drug development field, and more research into the mechanism by which FXR activation affects NAFLD is imperative.

A Bioreactor Technology for Modeling Fibrosis in Human and ...

<https://aasldpubs.onlinelibrary.wiley.com/doi/full/10.1002/hep.30651>

It has been reported that in human liver, αSMA + cells are present in the portal tracts, perivenular space, and liver lobule with discontinuous staining of qHSCs in adult human liver biopsies or surgical resections. 21 Consistent with this report, αSMA + cells were detected in central veins, portal tracts, and perisinusoidal spaces of normal ...

Cenicriviroc for the treatment of non-alcoholic ...

<https://www.researchgate.net/publication/323227193...>

CCR2 plays a critical role in recruiting monocytes with an inflammatory phenotype into the liver, while CCR5 attracts lymphocytes into the liver and is directly involved in fibrogenic activation ...

Therapeutic pro-fibrogenic signaling pathways in ...

<https://www.sciencedirect.com/science/article/pii/S0169409X1730073X>

Nov 01, 2017 · 2.1. Cells of innate immunity in regulating fibrosis 2.1.1. Monocytes and macrophages in optimal wound healing. Macrophages exhibit a major “regulatory” activity practically in any stage of repair and fibrosis , and this is due to their ability to display a remarkable plasticity and to adapt their behavior in response to environmental signals and cues, giving rise to different populations ...

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Dec 01, 2019 · 1. Introduction. **Liver** cirrhosis is currently ranked 11th among the most common causes of death in the world, and is an important reason for disability-adjusted life years and years of life loss, bringing a huge disease burden to the world [1,2]. The cause of **liver** cirrhosis is due to sustained **liver** injury induced by different factors, such as **alcohol** abuse, nonalcoholic steatohepatitis (NASH ...

Cited by: 2**Author:** Yuan-Yuan Fan, Wen Ding, Cheng Zhang...**Publish Year:** 2019

Mechanisms of bile acid mediated Inflammation in the Liver

<https://www.ncbi.nlm.nih.gov/pmc/articles/PMC5662014>

In the **liver**, as in other organs, an excessive **activation** of neutrophils induces additional tissue damage as demonstrated during hepatic ischemia-reperfusion, as well as in viral hepatitis, non-alcoholic fatty **liver** disease, alcoholic **liver** disease, **liver** fibrosis/cirrhosis and other causes of **liver** failure (66–68).

Cited by: 74**Author:** Man Li, Shi-Ying Cai, James L. Boyer**Publish Year:** 2017

Obeticholic acid alleviate lipopolysaccharide-induced ...

<https://www.sciencedirect.com/science/article/pii/S1567576918310671>

Jan 01, 2019 · **Obeticholic acid attenuates** LPS-induced inflammation in the lungs. ... FXR is highly expressed in **human** and rodent animal lungs [17,18 ... W.J. You, B. Liu, H.D. Jiang Bile acids induce **activation** of alveolar epithelial **cells** and lung fibroblasts through farnesoid X receptor-dependent and independent pathways. *Respirology*, 21 (6) (2016), pp ...

Cited by: 13**Author:** Jun Fei, Lin Fu, Biao Hu, Yuan-Hua Chen...**Publish Year:** 2019

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Cited by: 7**Author:** Natascha Roehlen, Emilie Crouchet, Tho...