

Document prepared by Nerve Center of TFORD, Venture Center, Pune Task Force on Repurposing of Drugs (TFORD) for COVID19

S&T Core Group on COVID19 constituted by PSA to Gol

Medicinal Herb Briefs: Glycyrrhiza glabra

Ref: TFORD/MHB/002 Date: 26 May 2020

About this document: This document summarizes information available on medicinal herb candidates for COVID19. One Medicinal Herb Brief document covers one candidate at a time.

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1. Summary Information on Glycyrrhiza glabra

Information About the	e Herb for Reported Indication(s)
Common Name	Liquorice, Gancao, Glycyrrhiza, sweet root, Yasti-madhu
Botanical Name (Family)	Glycyrrhizia glabra (Fabaceae)
Type of Plant/Source	Perineal Herb
of Herbal Ingredients	Source - Roots
	https://www.ema.europa.eu/en/documents/herbal-report/final-assessment-
	report-glycyrrhiza-glabra-l/glycyrrhiza-inflata-bat/glycyrrhiza-uralensis-fisch-
TION I C C	radix_en.pdf
TKDL Information	71 mentions for the Herb on TKDL
Indian Pharmacopeia Information	Herbal Drug Monograph available in Indian Pharmacopeia 2018
Reported	Anti-inflammatory, Expectorant, Anti-viral, Immunomodulator, Anti-ulcer,
Pharmacological	Hepatoprotective, Anti-oxidant, Anti-carcinogenic, Corticosteroid
Effects	https://www.ema.europa.eu/en/documents/herbal-report/final-assessment-
	report-glycyrrhiza-glabra-l/glycyrrhiza-inflata-bat/glycyrrhiza-uralensis-fisch-
Davisantad	radix en.pdf
Reported	Targets reported for Bioactive (for details please see Bioactive Section)
Therapeutic Target(s)	
Reported Mode of	Anti-inflammatory activity: By reducing nitric oxide and prostaglandin E2
Action	production
7100011	https://www.ncbi.nlm.nih.gov/pubmed/18075885
	Antiviral activity: Attributed to Glycyrrhizin which indirectly stimulates
	endogenous viral defense mechanisms
	Immunomodulator activity: Stimulation of IFN-gamma production by T
	cells
	https://www.ncbi.nlm.nih.gov/pubmed/9055991
Herb is reported to be beneficial for which conditions?	Treatment of Sore throat (demulcent), Cough, Bronchial Catarrh (expectorant), Allergic reactions, Arthritis, Rheumatis, Tuberculosis, Adrenocorticoid insufficiency
	 Prophylaxis and Treatment of Gastric Ulcers, Duodenal Ulcers, Dyspepsia Prevention of Liver Toxicity

		World Health Organization. 1999. 'Radix Glycyrrhizae.' WHO Monographs			
		on Selected Medicinal Plants, Vol. 1.			
Is the herb or its		Yes- Both			
extract form used	for	165- DO(11			
clinical purposes?					
Prescribed Dose		E 15 a per day of out as pourdered root, or day extracts equivalent to 200 600			
Prescribed Dose		5-15 g per day of cut or powdered root, or dry extracts equivalent to 200-600			
		mg of glycyrrhizin.			
		http://cms.herbalgram.org/expandedE/Licoriceroot.html#Administration(Ameri			
		can Botanical Council Database)			
Route of		Oral			
Administration					
Safety Profile (dos	se	Clinical studies show that short-term use (not more than 4 weeks) is safe.			
range in which it h	nas	However, chronic use can cause hypokalaemia, hypertension and, more			
been tested to be		rarely, cardiac rhythm disorders.			
safe in humans)		https://www.ema.europa.eu/en/documents/herbal-report/final-assessment-			
,		report-glycyrrhiza-glabra-l/glycyrrhiza-inflata-bat/glycyrrhiza-uralensis-fisch-			
		radix en.pdf			
Adverse events/S	ide	No adverse reactions have been associated with the herb when used within			
effects reported a		the			
the current	-	recommended dosage and treatment period.			
prescribed dose		Prolonged use (6 weeks) of excessive doses (50g/day) can lead to			
procention acce		pseudoaldosteronism, which includes potassium depletion, sodium retention,			
		oedema, hypertension, and weight gain. In rare cases, myoglobinuria			
		and myopathy can occur			
		World Health Organization. 1999. 'Radix Glycyrrhizae.' WHO Monographs on			
		Selected Medicinal Plants, Vol. 1.			
Donorted Drug He	orb				
Reported Drug-He interactions	erb	Because it increases potassium loss, Radix Glycyrrhizae (root) should not be			
interactions		administered for prolonged use with thiazide and loop diuretics or cardiac			
		glycosides. Because it reduces sodium and water excretion, the effectiveness			
		of drugs used in the treatment of hypertension may be reduced. Radix			
		Glycyrrhizae			
		should not be administered in conjunction with spironolactone or amiloride			
		World Health Organization. 1999. 'Radix Glycyrrhizae.' WHO Monographs on			
		Selected Medicinal Plants, Vol. 1.			
Link to Datasheet		https://www.ema.europa.eu/en/documents/herbal-report/final-assessment-			
		report-glycyrrhiza-glabra-l/glycyrrhiza-inflata-bat/glycyrrhiza-uralensis-fisch-			
		<u>radix_en.pdf</u>			
Regions where He	erb	Local Distribution: Himachal Pradesh, Jammu and Kashmir, Punjab			
is found		https://www.biodiversityofindia.org/index.php?title=Glycyrrhiza_glabra#tab=H			
		abitat and Geographical Spread			
Is the Herb or Ext	ract	Yes. Available as dried roots, crude powder and extract			
being sold in India	a?	· •			
Indian		Alkaloids Corporation			
Manufacturer(s)		https://www.pharmacompass.com/manufacturers-suppliers-exporters/licorice-			
		root			
International		AIPS Pharmaceutical (Japan), Ransom Naturals Ltd (UK), Finzelberg			
Manufacturer(s)		(Germany)			
manuacturer(3)		https://www.pharmacompass.com/manufacturers-suppliers-exporters/licorice-			
		root			
Cost of the Herb and		Licorice roots- Rs. 140-240/Kg, Licorice extract powder- Rs. 950/Kg, Licorice			
		· · · · · · · · · · · · · · · · · · ·			
Extract products in		extract- Rs. 450/50ml			
India	4 41	Major Dispetive for Deports displication (a)			
		Major Bioactive for Reported Indication(s)			
		MATION IS ONLY FOR THE MAJOR BIOACTIVE IN THE HERB			
What is the	Gly	cyrrhizic acid and its salts mainly Diammonium glycyrrhizinate			
Major Bioactive					
in the Herb?					
Is the Major	Yes				
Bioactive, in					
isolated form					

used for clinical	
purposes? Drug Bank ID	DB13751
Drug Barik ID	https://www.drugbank.ca/drugs/DB13751
Reported	Anti-inflammatory, Anti-viral, Immunomodulatory
Pharmacological	https://www.drugbank.ca/drugs/DB13751
effects	Tittps://www.drugbank.ca/drugs/DB13731
	Cartiagataraid 11 hata dahudraganaga igazuma 1 Tumar nagrasia factor
Reported	Corticosteroid 11-beta-dehydrogenase isozyme 1,Tumor necrosis factor,
Therapeutic	Caspase-3, Nuclear factor NF-kappa-B (Protein Group), Lipoprotein lipase https://www.drugbank.ca/drugs/DB13751
Target(s) Reported Mode	
of Action	 Anti-inflammatory activity: Generated via suppression of TNF alpha and Caspase 3. It also inhibits the translocation of NFkB into the nuclei and
OI ACION	conjugates free radicals. Some studies have shown a Glycyrrhizic-driven
	inhibition of CD4+ T cell proliferation via JNK, ERK and PI3K/AKT pathways.
	Antiviral activity: Inhibition of viral replication https://www.dr.uphople.co/dr.up/DR43754
Disastiva is	https://www.drugbank.ca/drugs/DB13751
Bioactive is	Chronic hepatitis in China and Japan (for over 20 years)
reported to be beneficial for	https://onlinelibrary.wiley.com/doi/pdf/10.1046/j.1365-2036.1998.00309.x https://www.sciencedirect.com/topics/neuroscience/glycyrrhizic-acid
which	
conditions?	https://www.ncbi.nlm.nih.gov/pmc/articles/PMC4052927/
Prescribed Dose	50mg to 100mg
r rescribed Dose	https://patents.google.com/patent/CN102188404A/en
Route of	Oral
Administration	Ordi
Safety Profile	Data not available
(dose range in	Data flot available
which it has	
been tested to	
be safe in	
humans)	
Adverse	In large amounts, glycyrrhizin salt can cause high blood pressure, water
events/Side	retention, & low potassium levels, which could lead to heart problems.
effects reported	Combined administration of glycyrrhizin salt with diuretics (water pills),
at the current	corticosteroids, or other medicines that decrease the body's potassium levels.
prescribed dose	https://www.ncbi.nlm.nih.gov/pmc/articles/PMC6836258/
Reported Drug-	94 drug-drug interactions reported
Bioactive	Inhibits CYP2C19 and CYP3A4
interactions	https://www.drugbank.ca/drugs/DB13751
Link to	Data not available
Datasheet for	
Bioactive	
Is the Bioactive	No
being sold in	
India?	
Indian	No
Manufacturer(s)	
International	CTTQ Pharma, Jiangsu Zhengda Pharmaceutical, Sino Biopharmaceutical,
Manufacturer(s)	Alchem International
	https://www.cttq.com/en/product/12203.htm
	http://www.jstsyy.com/product.asp
	https://www.marketscreener.com/SINO-BIOPHARMACEUTICAL-LI-
	6170858/news/Sino-Biopharmaceutical-VOLUNTARY-ANNOUNCEMENT-
	COMPLETION-OF-REGISTRATION-FOR-THE-CLINICAL-RESEAR-30063776/
Cook of the	http://www.alcheminternational.com/product/glycyrrhizinate-ammonium-mag/
Cost of the	Data not available
Bioactive in	
India	ut the Candidate for COVID-19
miorination Abo	ut the Candidate for COVID-19

Repurposing Claim	Prophylaxis & Treatment; New Indication (COVID-19) + New Dose (not confirmed)
Rationale for Repurposing for COVID19/MoA?	 Reported to be one of the Traditional Chinese Medicines suggested for use in COVID-19 pneumonia patients. https://www.sciencedirect.com/science/article/pii/S1043661820307556?via%3Dih ub A review article by Ayurvedic research groups suggest <i>G.glabra</i> could be considered for evaluation as a prophylactic therapy for COVID-19 https://www.ncbi.nlm.nih.gov/pmc/articles/PMC7177084/ Glycyrrhizic acid: Anti-viral activity reported for SARS-CoV (inhibits viral replication, adsorption, penetration) in a pre-clinical studies (in-vitro) https://www.ncbi.nlm.nih.gov/pubmed/12814717 https://www.ncbi.nlm.nih.gov/pmc/articles/PMC7128415/ According to a news article, Chinese National Medical Products Administration has approved the use of Lianhua-Qingwen capsule for the treatment of COVID-19. The capsule is a formulation of 11 herbs of which <i>G.glabra</i> is one of them. https://news.cgtn.com/news/2020-04-16/Beijing-approves-TCM-drugs-for-COVID-19-PJWgGfRAQg/index.html https://news.cgtn.com/news/2020-05-05/Expert-Lianhua-Qingwen-capsule-proven-effective-for-COVID-19-QeXiCOf18A/index.html
Proposed use as Primary or Adjuvant?	Primary
Pre-Clinical Data available for COVID-19	Data not available
Status of Clinical Trials	2 ongoing trials (1for Herb and 1 for Bioactive). Trial with Herb is in India
Trial Details	See details below

Trial ID/Link	Type of Trial	Form used	No. of patient s	Drug Combination/ Dose/ Stage of Disease	Primary and Secondary Measures	Has data from the trial been published?
ChiCTR20 00029768 (Therapy)	Randomized, open, controlled trial	Bioactive	30	Diammonium glycyrrhizinate enteric-coated capsules with vitamin C tablets Dose: Oral, 150mg, Tid Stage: Data not available	 Time to improvement of Virus infection symptoms Length of hospitalization /Healed and discharged Time to conversion of 2019-nCoV RNA result from RI sample Rate of preventing mild to moderate patients from shifting to severe Patients 	No

					 Time to Clinical recovery 7. Changes in lung inflammation absorption 	
CTRI/2020/ 05/025093 (Prophylaxi s)	Open label, Randomized, Controlled, Prospective, interventional, Community-based clinical study	Herb (extract)	1200	YASHTIMADHU TABLET Dose: 250 mg X 2 tablets bd for one month Stage: Data not available	Primary: Comparative assessment of occurrence of COVID-19 infection in healthy volunteers in community having at least 1 confirmed case already identified with control arm of Standard Prophylactic Care Secondary: 1. Comparative assessment of severity of Covid-19 infection in subjects taking YASHTIMADHU TABLET and those not taking it in following situations: a) Assessment of subjects not requiring hospitalization b) Severity of symptoms of hospitalized patients in wards c) Patients requiring ICU admission / Ventilator support 2. Global assessment of overall change as per the investigator (Efficacy assessment)	No

Key Data	Data not available
from	
Clinical	
Trials	
IP Status	Status/ Glycyrrhiza glabra
	Molecule
	Approved <u>225350</u>
	and Active Title: Anti-cough,anti-tussive and throat soothing synergistic herbal formulation
	applications Assignee: Council Of Scientific And Industrial Research (National Botanical

Pending applications 201811048303

Title: Ramban cough aushadi

Inventor: Maya Sharma Filing date: 20/12/2018 Status: Published: 15/02/2019

201811017566

Title: A formulation for increasing blood platelet counts

Inventor: HarparshantVir Singh Filing date: 10/05/2018 Status: Published: 01/06/2018

201621042902

Title: Herbal anti-asthmatic composition and process of preparation thereof

Inventor: Mohammad, MahafoozurrahmanDeen

Filing date:16/12/2016 Status: Published: 27/09/2019 201621006257

Title: Herbal composition for the treatment of herpes

Inventor: Mehra, Apurve Filing date:23/12/2016 Status: Published: 20/10/2017

201641001114

Title: Novel dietary supplementary immune system stimulator formulation and

method of preparation thereof

Inventor: Abdulsubahan Abdul Rahman

Filing date: 12/01/2016 Status: Published: 14/07/2017

1475/CHE/2015

Title: A composition comprising of Glycyrrhiza Glabra extract, and Vitex

Negundo extract and uses thereof

Assignee: ITC Life Sciences and Technology Centre

Filing date: 23/03/2015 Status: Published: 30/09/2016

3464/MUM/2014

Title: A novel synergistic herbal composition for enhancing immunity level in-vivo

and in-vitro: its process thereof

Inventor: Sadhana J. Rajput, Reshma A. Jain

Filing Date: 03/11/2017 Status: Published: 06/05/2016

808/MUM/2013

Title: Herbal composition for the treatment of herpes

Assignee: Mehra, Apurve Filing date: 15/03/2013 Status: Published 30/01/2015

3038/DEL/2012

Title: Herbal cough candy and process for the preparation of the same

Assignee: Amity University Filing date:28/09/2012 Status: Published 25/04/2014

1151/DEL/2012

Title: Immune booster herbal composition-internal use medicine

Assignee: Dhruva Kumar Sadh Filing date: 13/04/2012 Status: Published 18/05/2012

908/DEL/2012

Title: Asthma care herbal composition - internal use medicine

Assignee: Dhruva Kumar Sadh

Filing Date: 27/03/2012 Status: Publication 04/05/2012

3286/CHE/2011

Title: A process for preparation of synergestic herbal preparation of extracts of

PicrorhizaKurroa and Glycyrrhiza Glabra having medicinal values

Inventor: Anil Kumar M Filing date: 23/09/2011 Status: Published: 14/06/2013

2716/DEL/2011

Title: A novel ayurvedic formulation for the treatment of cough and other

associated symptoms

assignee: akums drugs & pharmaceuticals limited

Filing date:20/09/2011 Status: Published: 22/03/2013

1335/DEL/2011

Title: Ayurvedic/ herbal dry syrup formulation for treatment of respiratory

allergies

Assignee: BACFO Pharmaceuticals (India) Ltd

Filing date: 06/05/2011 Status: Published 9/11/2012

2946/MUM/2009

Title: A process for the preparation of herbal extracts of some Indian medicinal plants species of traditional ayurveda, use thereof for anti-mycobacterial activity

Assignee: CSRD, People's Group

Filing date: 21/12/2009 Status: Published: 10/02/2012

2930/MUM/2009

Title: Herbal formulation for respiratory tract problems, cough curiosa and

asthma in unique dosage form Assignee: Vasu Healthcare Pvt. Ltd.

Filing date: 18/12/2009 Status: Published: 30/08/2013

1632/MUM/2009

Title: Herbal tablet, powder and paste for combustion to treat respiratory tract infections and disorders along blood energiser work by way of its vapour

inhalation

Inventor: Bharat Manharlal Parikh

Filing date: 17/07/2009 Status: Published: 14/10/2011

1858/DEL/2006
Title: A herbal formulation effective against cough and process for preparation

thereof

Assignee: The Director, Central Council For Research In Unani Medicine

(CCRUM)

Filing date: 18/08/2006 Status: Published: 04/04/2008

1313/DEL/2006

Title: A preparation for the treatment of running and stuffy nose, productive or non-productive cough with or without fever and to a process for the preparation

thereof

Assignee: Central Council For Research Of Ayurveda And Siddha

Filing date: 31/05/2006 Status: Published:07/12/2007 2939/DELNP/2004 Title: A herbal soft drink

Assignee: Council Of Scientific And Industrial Research (National Botanical

Research Institute, Lucknow) Filing Date: 29/09/2004 Status: Published 27/04/2007

2929/MUM/2009

Title: A hepatoprotective herbal composition

Assignee:RatnamShiv Prakash And Pundarikakshudu, Kilambi

Filing Date: 18/12/2009 Status: Published 26/11/2010

3347/DEL/2015

Title: A novel antacid herbal oral powder formulation

Inventor: Narinderpal Singh Aulakh

Expired or <u>220700</u>

Lapsed application or status not known

Title: A bio-enhancer and bioavailability facilitator composition

Assignee: Council Of Scientific And Industrial Research (Central Institute Of

Medicanal And Aromatic Plants, Lucknow)

Filing Date: 31/08/2000

Grant date: 25/07/2008

Status: Patent Has Ceased on 02/09/2013

3338/KOLNP/2006

Title: Herbal composition

Assignee:Lavender Hill Projects Pty Ltd

Filing Date: 13/11/2006

Status: Application Appears To Have Been Abandoned

2419/DEL/2004

Title: A pharmaceutical composition useful for the treatment of asthma

Assignee:Council Of Scientific & Industrial Research (Institute of Genomics and

Integrative Biology, New Delhi) Filing Date: 02/12/2004

Status: Application Appears To Have Been Abandoned

236432 (No link on WIPO site)

Title: A process for isolating and purifying Glycyrrhizic Acid (*Glycyrrhiza Glabra*) roots effective against leishmaniasis and composition containing the same

Assignee:Bose Institute Filing Date: 23/08/2005

Status: Patent Has Ceased On 24/08/2015

1622/DELNP/2010

Title: Adhering troches with topically active ingredients for treatment of throat,

esophagus, and stomach Assignee:Orahealth Corporation Filing Date: 09/03/2010

Legal Status: Application Appears To Have Been Abandoned

633/MUM/2010

Title: Herbal composition for oral administration Assignee: J.B. Chemicals And Pharmaceuticals Ltd.

Filing Date: 10/03/2010

Status: Application Has Been Abandoned

2554/DEL/2011

Title: Herbal energy drink in different flavours

Assignee: Amity University Filing Date: 06/09/2011

Status: Application Appears to Have Been Abandoned

3784/CHE/2011

Title: An immune booster Assignee:ArvindVarchaswi Filing Date: 04/11/2011

Status: Application Appears To Have Been Abandoned

136/KOL/2013

Title: Synergistic herbal composition for the treatment of cough and the process

of preparing thereof

Assignee:Pawan Kumar Pareek

Filing Date: 06/02/2013

Status: Application Appears To Have Been Abandoned

217095

Title: A synergistic composition useful for management of asthma Assignee: Dabur Pharma Limited And Dabur Research Foundation

Filing Date: 04/06/2001

Status: Patent Has Ceased On 04/06/2010

2522/DEL/2008

Title: A novel nutraceutical formulation for vitality and virility

Assignee: Amity University Filing Date: 06/11/2008

Status: Application Has Been Abandoned

1149/DEL/2012

Title: Liver corrective herbal composition internal use medicine

Inventor: Dhruva Kumar Sadh Filing Date: 13/04/2012

Status: Application Appears To Have Been Abandoned

Other Key Reference s

- https://pubchem.ncbi.nlm.nih.gov/compound/Diammonium-glycyrrhizinate
- https://www.nature.com/articles/news030609-16
- https://www.thelancet.com/journals/lancet/article/PIIS0140-6736(03)13615-X/fulltext

2. Background information

About TFORD-COVID19

The Principal Scientific Advisor to the GoI, Dr K VijayRaghavan, has constituted a S&T Core Group on COVID19. Under the aegis of the S&T Core Group on COVID19, a Task Force has been constituted focused on Repurposing of Drugs for COVID19 (in short "TFORD-COVID19"). The Task Force is being coordinated by Dr V Premnath, Head, NCL Innovations at CSIR-NCL and Director, Venture Center and Dr Anurag Agarwal, Director, CSIR-IGIB. The Nerve Center for the Coordination is located be at Venture Center, Pune (located in the campus of CSIR-NCL).

Credits

Editors: Dr. Priya Nagaraj & Dr Smita Kale; Contributors: Dr. Priya Nagaraj, Dr. Vidula Walimbe, Dr. Smita Kale, Dr. Kirtee Wani, Dr. Tejas Shah, Dr. Swati Joshi, Dr. Manisha Premnath, Dr. Premnath V; Information also contributed by Dr Gopakumar Nair, GNAS and GnanLex.

About Advisory Group

The Nerve Center at TFORD-COVID19 has constituted an inter-disciplinary Advisory Group. This Advisory Group reviews the information compiled by the Nerve Center, provides suggestions on data, information sources, organization of data etc. while also providing inputs to refine the analysis and create a structured information base to support decision-making. The Advisory Group also provides expert input and opinions on certain selected points where experience-based inputs are needed. The members of the Advisory Group for each Discussion Paper are listed at https://nclinnovations.org/covid19/teams/.

Disclaimer

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