

Supplementary material, Appendix 1

Search Strategy:

PubMed

("Aloe Vera" OR "Aloe emodin" OR "Withania somnifera" OR "Ashwagandha" OR "Atractylis gummifera" OR "Dysosma" OR "Butterbur" OR "Petasites hybridus" OR "Rhamnus purshiana" OR "Cascara-Sagrada" OR "Cascara" OR "Larrea tridentata" OR "Chaparral" OR "Symphytum officinale" OR "Comfrey" OR "Dictamnus dasycarpus" OR "Eucalyptus globulus" OR "Equisetum Arvense" OR "Field Horsetail" OR "Flavocoxid" OR "Malabar tamarind" OR "Garcinia Cambogia" OR "Teucrium chamaedrys" OR "Germander" OR "Chelidonium majus" OR "Greater celandine" OR "Green Tea" OR "Camellia sinensis" OR "Gynura segetum" OR "Herbalife" OR "Polygonum multiflorum" OR "He-Shou-Wu" OR "Fo-ti" OR "Shou Wu Pian" OR "Hydroxycut" OR "Callilepis laureola" OR "Impila" OR "Jin Bu Huan" OR "Lycopodium serratum" OR "Kava kava" OR "Piper methysticum" OR "Kratom" OR "Mitragyna speciosa" OR "Ephedra sinica" OR "Ma Huang" OR "Malva sylvestris" OR "Malva" OR "Azadirachta indica" OR "Margosa oil" OR "Margosa" OR "Mistletoe" OR "Viscum album" OR "Move Free" OR "Noni" OR "Morinda citrifolia" OR "Mentha pulegium" OR "Pennyroyal" OR "Psoralea corylifolia" OR "Red Yeast Rice" OR "Monascus purpureus" OR "Roselle" OR "Hibiscus sabdariffa" OR "Cassia angustifolia" OR "Senna" OR "angustifolia C." OR "Skullcap" OR "Scutellaria baicalensis" OR "Huang Qin" OR "Scutellaria lateriflora" OR "Hypericum perforatum" OR "Hypericum" OR "Hyperico" OR "St. John's Wort" OR "Turmeric" OR "Curcuma longa" OR "Senecio" OR "Crotalaria" OR "Heliotropium" OR "Sacaca" OR "Croton Cajucara Benth" OR "Isabgol" OR "Plantago ovata") AND ("liver toxicity" OR "hepatotoxicity" OR "liver injury" OR "liver failure" OR "hepatitis") NOT ("Rat" OR "Rats" OR "Mice" OR "Wombats" OR "Sheep")

BIREME

("Aloe Vera" OR "Aloe emodin" OR "Withania somnifera" OR "Ashwagandha" OR "Atractylis gummifera" OR "Dysosma" OR "Butterbur" OR "Petasites hybridus" OR "Rhamnus purshiana" OR "Cascara-Sagrada" OR "Cascara" OR "Larrea tridentata" OR "Chaparral" OR "Symphytum officinale" OR "Comfrey" OR "Dictamnus dasycarpus"

OR "Eucalyptus globulus" OR "Equisetum Arvense" OR "Field Horsetail" OR
 "Flavocoxid" OR "Malabar tamarind" OR "Garcinia Cambogia" OR "Teucrium
 chamaedrys" OR "Germander" OR "Chelidonium majus" OR "Greater celandine" OR
 "Green Tea" OR "Camellia sinensis" OR "Gynura segetum" OR "Herbalife" OR
 "Polygonum multiflorum" OR "He-Shou-Wu" OR "Fo-ti" OR "Shou Wu Pian" OR
 "Hydroxycut" OR "Callilepis laureola" OR "Impila" OR "Jin Bu Huan" OR "Lycopodium
 serratum" OR "Kava kava" OR "Piper methysticum" OR "Kratom" OR "Mitragyna
 speciosa" OR "Ephedra sinica" OR "Ma Huang" OR "Malva sylvestris" OR "Malva" OR
 "Azadirachta indica" OR "Margosa oil" OR "Margosa" OR "Mistletoe" OR "Viscum
 album" OR "Move Free" OR "Noni" OR "Morinda citrifolia" OR "Mentha pulegium" OR
 "Pennyroyal" OR "Psoralea corylifolia" OR "Red Yeast Rice" OR "Monascus purpureus"
 OR "Roselle" OR "Hibiscus sabdariffa" OR "Cassia angustifolia" OR "Senna" OR
 "angustifolia C." OR "Skullcap" OR "Scutellaria baicalensis" OR "Huang Qin" OR
 "Scutellaria lateriflora" OR "Hypericum perforatum" OR "Hypericum" OR "Hyperico"
 OR "St. John's Wort" OR "Turmeric" OR "Curcuma longa" OR "Senecio" OR
 "Crotalaria" OR "Heliotropium" OR "Sacaca" OR "Croton Cajucara Benth" OR
 "Isabgol" OR "Plantago ovata") AND ("liver toxicity" OR "hepatotoxicity" OR "liver
 injury" OR "liver failure" OR "hepatitis") NOT ("Rat" OR "Rats" OR "Mice" OR
 "Wombats" OR "Sheep")

EMBASE

('aloe vera'/exp OR 'aloe vera' OR 'aloe emodin'/exp OR 'aloe emodin' OR 'withania
 somnifera'/exp OR 'withania somnifera' OR 'ashwagandha'/exp OR 'ashwagandha'
 OR 'atractylis gummifera'/exp OR 'atractylis gummifera' OR 'dysosma' OR
 'butterbur'/exp OR 'butterbur' OR 'petasites hybridus'/exp OR 'petasites hybridus' OR
 'rhamnus purshiana'/exp OR 'rhamnus purshiana' OR 'cascara-sagrada'/exp OR
 'cascara-sagrada' OR 'cascara'/exp OR 'cascara' OR 'larrea tridentata'/exp OR 'larrea
 tridentata' OR 'chaparral'/exp OR 'chaparral' OR 'symphytum officinale'/exp OR
 'symphytum officinale' OR 'comfrey'/exp OR 'comfrey' OR 'dictamnus
 dasycarpus'/exp OR 'dictamnus dasycarpus' OR 'eucalyptus globulus'/exp OR
 'eucalyptus globulus' OR 'equisetum arvense'/exp OR 'equisetum arvense' OR 'field
 horsetail' OR 'flavocoxid'/exp OR 'flavocoxid' OR 'malabar tamarind' OR 'garcinia
 cambogia'/exp OR 'garcinia cambogia' OR 'teucrium chamaedrys'/exp OR 'teucrium
 chamaedrys' OR 'germander'/exp OR 'germander' OR 'chelidonium majus'/exp OR

'chelidonium majus' OR 'greater celandine' OR 'green tea'/exp OR 'green tea' OR
 'camellia sinensis'/exp OR 'camellia sinensis' OR 'gynura segetum'/exp OR 'gynura
 segetum' OR 'herbalife'/exp OR 'herbalife' OR 'polygonum multiflorum'/exp OR
 'polygonum multiflorum' OR 'he-shou-wu'/exp OR 'he-shou-wu' OR 'fo-ti' OR 'shou
 wu pian' OR 'hydroxycut'/exp OR 'hydroxycut' OR 'callilepis laureola'/exp OR
 'callilepis laureola' OR 'impila' OR 'jin bu huan'/exp OR 'jin bu huan' OR 'lycopodium
 serratum'/exp OR 'lycopodium serratum' OR 'kava kava'/exp OR 'kava kava' OR
 'piper methysticum'/exp OR 'piper methysticum' OR 'kratom'/exp OR 'kratom' OR
 'mitragyna speciosa'/exp OR 'mitragyna speciosa' OR 'ephedra sinica'/exp OR
 'ephedra sinica' OR 'ma huang'/exp OR 'ma huang' OR 'malva sylvestris'/exp OR
 'malva sylvestris' OR 'malva'/exp OR 'malva' OR 'azadirachta indica'/exp OR
 'azadirachta indica' OR 'margosa oil'/exp OR 'margosa oil' OR 'margosa' OR
 'mistletoe'/exp OR 'mistletoe' OR 'viscum album'/exp OR 'viscum album' OR 'move
 free' OR 'noni'/exp OR 'noni' OR 'morinda citrifolia'/exp OR 'morinda citrifolia' OR
 'mentha pulegium'/exp OR 'mentha pulegium' OR 'pennyroyal'/exp OR 'pennyroyal'
 OR 'psoralea corylifolia'/exp OR 'psoralea corylifolia' OR 'red yeast rice'/exp OR 'red
 yeast rice' OR 'monascus purpureus'/exp OR 'monascus purpureus' OR 'roselle'/exp
 OR 'roselle' OR 'hibiscus sabdariffa'/exp OR 'hibiscus sabdariffa' OR 'cassia
 angustifolia'/exp OR 'cassia angustifolia' OR 'senna'/exp OR 'senna' OR 'angustifolia
 c.' OR 'skullcap'/exp OR 'skullcap' OR 'scutellaria baicalensis'/exp OR 'scutellaria
 baicalensis' OR 'huang qin'/exp OR 'huang qin' OR 'scutellaria lateriflora'/exp OR
 'scutellaria lateriflora' OR 'hypericum perforatum'/exp OR 'hypericum perforatum' OR
 'hypericum'/exp OR 'hypericum' OR 'hiperico' OR 'st. john wort' OR 'turmeric'/exp
 OR 'turmeric' OR 'curcuma longa'/exp OR 'curcuma longa' OR 'senecio'/exp OR
 'senecio' OR 'crotalaria'/exp OR 'crotalaria' OR 'heliotropium'/exp OR 'heliotropium'
 OR 'sacaca' OR 'croton cajucara benth' OR 'isabgol' OR 'plantago ovata'/exp OR
 'plantago ovata') AND ('liver toxicity'/exp OR 'liver toxicity' OR 'hepatotoxicity'/exp
 OR 'hepatotoxicity' OR 'liver injury'/exp OR 'liver injury' OR 'liver failure'/exp OR
 'liver failure' OR 'hepatitis'/exp OR 'hepatitis' OR 'hepatic injury'/exp OR 'hepatic
 injury' OR 'hepatic failure'/exp OR 'hepatic failure') NOT ('rat'/exp OR 'rat' OR
 'rats'/exp OR 'rats' OR 'mice'/exp OR 'mice' OR 'wombats' OR 'sheep'/exp OR 'sheep')
 AND ('article'/it OR 'article in press'/it OR 'chapter'/it OR 'conference abstract'/it OR
 'conference paper'/it OR 'conference review'/it OR 'editorial'/it OR 'erratum'/it OR
 'letter'/it OR 'note'/it OR 'short survey'/it)

Lilacs

("Aloe Vera" OR "Aloe emodin" OR "Withania somnifera" OR "Ashwagandha" OR "Atractylis gummifera" OR "Dysosma" OR "Butterbur" OR "Petasites hybridus" OR "Rhamnus purshiana" OR "Cascara-Sagrada" OR "Cascara" OR "Larrea tridentata" OR "Chaparral" OR "Symphytum officinale" OR "Comfrey" OR "Dictamnus dasycarpus" OR "Eucalyptus globulus" OR "Equisetum Arvense" OR "Field Horsetail" OR "Flavocoxid" OR "Malabar tamarind" OR "Garcinia Cambogia" OR "Teucrium chamaedrys" OR "Germander" OR "Chelidonium majus" OR "Greater celandine" OR "Green Tea" OR "Camellia sinensis" OR "Gynura segetum" OR "Herbalife" OR "Polygonum multiflorum" OR "He-Shou-Wu" OR "Fo-ti" OR "Shou Wu Pian" OR "Hydroxycut" OR "Callilepis laureola" OR "Impila" OR "Jin Bu Huan" OR "Lycopodium serratum" OR "Kava kava" OR "Piper methysticum" OR "Kratom" OR "Mitragyna speciosa" OR "Ephedra sinica" OR "Ma Huang" OR "Malva sylvestris" OR "Malva" OR "Azadirachta indica" OR "Margosa oil" OR "Margosa" OR "Mistletoe" OR "Viscum album" OR "Move Free" OR "Noni" OR "Morinda citrifolia" OR "Mentha pulegium" OR "Pennyroyal" OR "Psoralea corylifolia" OR "Red Yeast Rice" OR "Monascus purpureus" OR "Roselle" OR "Hibiscus sabdariffa" OR "Cassia angustifolia" OR "Senna" OR "angustifolia C." OR "Skullcap" OR "Scutellaria baicalensis" OR "Huang Qin" OR "Scutellaria lateriflora" OR "Hypericum perforatum" OR "Hypericum" OR "Hyperico" OR "St. John's Wort" OR "Turmeric" OR "Curcuma longa" OR "Senecio" OR "Crotalaria" OR "Heliotropium" OR "Sacaca" OR "Croton Cajucara Benth" OR "Isabgol" OR "Plantago ovata") AND ("liver toxicity" OR "hepatotoxicity" OR "liver injury" OR "liver failure" OR "hepatitis") NOT ("Rat" OR "Rats" OR "Mice" OR "Wombats" OR "Sheep")

Cochrane Library

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Open Grey

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SCIELO

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SCOPUS

TITLE-ABS-KEY("Aloe Vera" OR "Aloe emodin" OR "Withania somnifera" OR "Ashwagandha" OR "Atractylis gummifera" OR "Dysosma" OR "Butterbur" OR "Petasites hybridus" OR "Rhamnus purshiana" OR "Cascara-Sagrada" OR "Cascara" OR "Larrea tridentata" OR "Chaparral" OR "Symphytum officinale" OR "Comfrey" OR "Dictamnus dasycarpus" OR "Eucalyptus globulus" OR "Equisetum Arvense" OR "Field Horsetail" OR "Flavocoxid" OR "Malabar tamarind" OR "Garcinia Cambogia" OR "Teucrium chamaedrys" OR "Germander" OR "Chelidonium majus" OR "Greater celandine" OR "Green Tea" OR "Camellia sinensis" OR "Gynura segetum" OR "Herbalife" OR "Polygonum multiflorum" OR "He-Shou-Wu" OR "Fo-ti" OR "Shou Wu Pian" OR "Hydroxycut" OR "Callilepis laureola" OR "Impila" OR "Jin Bu Huan" OR "Lycopodium serratum" OR "Kava kava" OR "Piper methysticum" OR "Kratom" OR "Mitragyna speciosa" OR "Ephedra sinica" OR "Ma Huang" OR "Malva sylvestris" OR "Malva" OR "Azadirachta indica" OR "Margosa oil" OR "Margosa" OR "Mistletoe" OR "Viscum album" OR "Move Free" OR "Noni" OR "Morinda citrifolia" OR "Mentha pulegium" OR "Pennyroyal" OR "Psoralea corylifolia" OR "Red Yeast Rice" OR "Monascus purpureus" OR "Roselle" OR "Hibiscus sabdariffa" OR "Cassia angustifolia" OR "Senna" OR "angustifolia C." OR "Skullcap" OR "Scutellaria baicalensis" OR "Huang Qin" OR "Scutellaria lateriflora" OR "Hypericum perforatum" OR "Hypericum" OR "Hyperico" OR "St. John's Wort" OR "Turmeric" OR "Curcuma longa" OR "Senecio" OR "Crotalaria" OR "Heliotropium" OR "Sacaca" OR "Croton Cajucara Benth" OR "Isabgol" OR "Plantago ovata") AND TITLE-ABS-KEY("liver toxicity" OR "hepatotoxicity" OR "liver injury" OR "liver failure" OR "hepatitis") AND TITLE-ABS-KEY("Rat" OR "Rats" OR "Mice" OR "Wombats" OR "Sheep")

Web of Science

TOPIC:(("Aloe Vera" OR "Aloe emodin" OR "Withania somnifera" OR "Ashwagandha" OR "Atractylis gummifera" OR "Dysosma" OR "Butterbur" OR "Petasites hybridus" OR "Rhamnus purshiana" OR "Cascara-Sagrada" OR "Cascara" OR "Larrea tridentata" OR "Chaparral" OR "Symphytum officinale" OR "Comfrey" OR "Dictamnus dasycarpus" OR "Eucalyptus globulus" OR "Equisetum Arvense" OR "Field Horsetail" OR "Flavocoxid" OR "Malabar tamarind" OR "Garcinia Cambogia" OR "Teucrium chamaedrys" OR

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Supplementary Table 1 Baseline features of herb-induced liver injury (HILI) with more than 1% of prevalence

Variable	Patients n = 775 (100%)
Mean age (yr)	44.59
Sex (female)	512 (66)
Race	
White	71(9.1)
Black	5 (.6)
Other	40 (5.1)
Application	
Weight Loss	138 (17.8)
Psychiatry disorders	50 (6.4)
Pain control	34 (4.3)
Physical improvement	33 (4.2)
Health improvement	24 (3)

Sleep disorders	22 (2.8)
Symptoms	
Jaundice	371 (47.8)
Abdominal Pain	173 (22.3)
Nausea	141 (18.1)
Fatigue	131 (16.9)
Urine alteration	114 (14.7)
Coluria	99 (12.7)
Hematuria	4 (.5)
Vomiting	87 (11.2)
Hepatomegaly	84 (10.8)
Pruritus	73 (9.4)
Ascites	54 (6.9)
Stools alteration	45 (5.8)
Clay colored	44 (5.6)
Melena	1 (.1)
Anorexia	40 (5.1)
Encephalopathy	31 (4)
Malaise	28 (3.6)
Asthenia	27 (3.4)
Diarrhea	23 (2.9)
Weakness	23 (2.9)
Hyporexia	18 (2.3)
Myalgia	14 (1.8)
Fever	13 (1.6)
Lethargy	11 (1.4)
Antibodies	
ANA	45 (5.8)
ASMA	23 (2.9)
AMA	4 (.5)
Anti-dsDNA	2 (.2)
Actin	1 (.1)
Anti-LKM	1 (.1)
AMA-M2	1 (.1)

cANCA	1 (.1)
pANCA	1 (.1)
Rheumatoid Factor	1 (.1)
AST (U/L)	1331
ALT (U/L)	1490
ALP (U/L)	279
GGT (U/L)	264
Total Bilirubin (mg/dL)	13.17
Direct Bilirubin (mg/dL)	8.19
HILI pattern	
Hepatocellular	584 (75.3)
Cholestatic	70 (9)
Mix	68 (8.7)
Sinusoidal obstruction syndrome	41 (5.2)
Ductopenia	3 (.3)
Granulomatous Hepatitis	2 (.2)
Steatosis	2 (.2)
Nontippable	2 (.2)
Other	3 (.3)
Use duration (days)	130
Latency (days)	127
Stop and manifestation (days)	1
Normalization liver function (days)	80
Comorbidities	
Hypertension	36 (4.6)
Obesity	36 (4.6)
Dyslipidemia	33 (4.2)
Alcohol use	24 (3)
Hypothyroidism	23 (2.9)
Osteoarthritis	12 (1.5)
Anxiety	9 (1.1)
Asthma	9 (1.1)
Depression	7 (.9)
None	304 (39.2)

Other medications	
Vitamins and Minerals	53 (6.38)
Antihypertensive	47 (6)
NSAIDs	39 (5)
Estrogen/Progesterone	34 (4.3)
Statins	20 (2.5)
Paracetamol	18 (2.3)
Diuretics	17 (2.1)
Antibiotics	15 (1.9)
Diabetes mellitus drugs	12 (1.5)
Corticoid	12 (1.5)
Benzodiazepine	12 (1.5)
Proton Pump Inhibitor	10 (1.2)
SSRI	10 (1.2)
None	452 (58.3)
Temporal relation	22 (15.2)
Biopsy	318 (41)
Viral Hepatitis	
Hep A	2 (.2)
Hep B	3 (.3)
Hep C	7 (.9)
CMV	2 (.2)
HSV	2 (.2)
Alcoholic liver disease	2 (.2)
Bile duct obstruction	1 (.1)
Other causes	
Pregnancy	1 (.1)
Acute Hypotension	1 (.1)
Rechallenge	70 (9)
Previous Report Hepatotoxicity	775 (100)
Causality	
Definitive	22 (2.8)
Highly probable	8 (1)
Probable	122 (15.7)

Possible	49 (6.3)
Improbable	2 (0.2)
Excluded	7 (0.9)
Maria and Victorino (M & V)	8.95 (SD 3.84)
Treatment	
Supportive care	571 (73.6)
Liver transplantation	67 (7.7)
Corticoid	38 (4.3)
Acetylcysteine	21 (2.5)
Glutathione	21 (2.5)
Ursodeoxycholic acid	21 (2.5)
Adenosylmethionine	18 (2)
Glycyrrhizin	18 (2)
Polyene Phosphatidylcholine	18 (2)
Vitamin K	14 (2)
Antibiotics	10 (1.2)
Cholestyramine	3 (.3)
Outcome	
Recovery	679 (87.6)
Chronicity	11 (1.4)
Death	49 (6.3)

Supplementary Table 2 Herbs details features that induced-liver injury

Generic Or Trade Names	Latin name	N	Countries†	Indications†	Symptoms†	HILI Patterns†
He-Shou-Wu; Fo-Ti; Shou Wu Pian.	<i>Polygonum multifloru m</i>	91	China (54.9), South Korea (28.5)	Hair Supplement (26.6), Insomnia (8.8), Improve Coordination (8.8)	Jaundice (80), Choloria (17.7), Fatigue (15.5), Hepatomegaly (13.3)	Hepatocellular (72.5), Cholestatic (18.6), Mixed (8.7)
Green Tea Extract	<i>Camellia sinensis</i>	90	USA (32.5), Spain (19.1), Japan (14.6)	Weight Loss (71.4), Physical Performance (6.3), Health Improvement (4.7)	Jaundice (61.4), Fatigue (30.4), Nausea (27.5), Abdominal Pain (26)	Hepatocellular (78.8), Cholestatic (8.8), Mixed (11.1)
Kava Kava	<i>Piper methysticu m</i>	62	Germany (65), USA (21.6), Spain (3.3)	Hair Supplement (26.6), Insomnia (8.8), Coordination (8.8)	Jaundice (86.8), Nausea (47.8), Fatigue (34.7)	Hepatocellular (77.4), Cholestatic (9.6), Mixed (8)
Greater Celandine	<i>Chelidonium majus</i>	48	Germany (81.2), Italy (8.3)	Pain (38.1), Dyspesia (11.9)	Jaundice (34.2), Choloria (31.4), Pruritus (28.5), Nausea (27.7)	Hepatocellular (72.9), Cholestatic (6.2), Mixed (20.8)
Germander	<i>Teucrium Chamaedrys</i>	35	France (28.5), Greece (22.8), USA (14.2)	Dyslipidemia (31), Weight Loss (27.5),	Jaundice (77.1), Nausea (17.1), Choloria (17.1), Abdominal Pain (17.1)	Hepatocellular (94.2), Mixed (5.7)

				Diabetes Mellitus (13.7)		
Skullcap	<i>Scutellaria spp.</i>	35	USA (45.7), Australia (14.2), Scotland (11.4)	Health Improvement (25), Osteoarthritis (20.8), Arthralgias (12.5)	Jaundice (68), Nausea (15.3), Choluria (20)	Hepatocellular (74.2), Cholestatic (5.7), Mixed (14.2)
Kratom	<i>Mitragyna speciosa</i>	33	USA (75), Canada (6.2), Sweden (6.2)	Pain (38.8)	Jaundice (70), Choluria (53.3), Abdominal Pain (43.5), Nausea (23.3), Fatigue (23.3)	Hepatocellular (45.4), Cholestatic (27.2), Mixed (21.2)
Tusanqi Or Jusanqi	<i>Gynura segetum</i>	29	China (75.8), Hong Kong (20.6), New Zealand (3.4)	Cooking (80), Traumatic Injuries (20)	Ascites (100), Hepatomegaly (86.9), Jaundice (26)	Sinusoidal Obstruction Syndrome (100)
Garcinia Cambogia; Malabar Tamarind	<i>Garcinia gummi-gutta</i>	29	USA (39.2), Italy (35.7)	Weight Loss (100)	Abdominal Pain (56), Jaundice (44), Nausea (40), Vomiting (24), Fatigue (16)	Hepatocellular (86.2), Cholestatic (10.3), Mixed (3.4)
Ma Huang	<i>Ephedra sinica</i>	27	USA (68), South Korea (12), United Kingdom (8)	Weight Loss (57.8), Physical Performance (15.7), Insomnia (15.7)	Abdominal Pain (72.7), Nausea (63.6), Jaundice (54.5), Fatigue (45.4), Hepatomegaly (36.3)	Hepatocellular (96), Mixed (4)
Chaparral	<i>Larrea tridentata</i>	26	USA (80.7), Australia (7.6), Canada (7.6)	Health Improvement (47.3)	Jaundice (84), Abdominal Pain (40), Fatigue (36), Nausea (28), Pruritus (24)	Hepatocellular (84.6), Steatosis (15.3)

Senna; Sene.	<i>Senna spp.</i>	25	India (24), Italy (16), Yemen (16)	Constipation (42.8), Accidental Ingestion (38.1)	Jaundice (43.4), Abdominal Pain (34.7), Encephalopathy (20), Asthenia (20), Choluria (13)	Hepatocellular (84), Cholestatic (4), Mixed (12)
Aloe Vera	<i>Aloe vera</i>	22	South Korea (18.1), Spain (9), USA (9.)	Weight Loss (50), Constipation (14.2)	Abdominal Pain (50), Jaundice (40.9), Fatigue (31.8), Nausea (27.2)	Hepatocellular (86.3), Cholestatic (4.5), Mixed (9)
Jin Bu Huan	<i>Lycopodium serratum</i>	19	USA (73.6), Canada (10.5), Italy (10.7)	Insomnia (72.22), Pain Management (16.67)	Fatigue (52.6), Hepatomegaly (42.1), Pruritus (36.8), Jaundice (31.5)	Hepatocellular (84.2), Cholestatic (5.2), Mixed (10.5)
Valerian	<i>Valeriana officinalis</i>	16	Germany (31.2), Scotland (25), USA (18.7)	Health Improvement (40), Insomnia (40), Anti-Stress (20)	Jaundice (75), Choluria (25), Abdominal Pain (18.7), Clay Stools (18.7), Fatigue (18.7)	Hepatocellular (56.2), Cholestatic (25), Mixed (18.7)
Red Yeast Rice; Red Koji Rice.	<i>Monascus purpureus</i>	16	Italy (75), USA (18.7), Belgium (6.2)	Dyslipidemia (85.7), Dyspepsia (14.2)	Jaundice (37.5), Fatigue (25), Clay Stools (25)	Hepatocellular (87.5), Mixed (6.2), Other (6.2)
Black Cohosh	<i>Actaea racemosa</i>	15	Germany (26.6), Australia (20), Belgium (13.3), United Kingdom (13.3)	Menses symptoms (50)	Jaundice (46.1), Nausea (30.7), Abdominal pain (23)	Hepatocellular (80), Cholestatic (13.3), Mixed (6.6)
Noni	<i>Morinda citrifolia</i>	14	Austria (28.5), Spain (21.4), USA (21.4)	Health improvement (28.5), improves immunity (28.5)	Jaundice (61.5), Abdominal pain (38.4), Nausea (28.5), Hepatomegaly (23), Fatigue (21.4), Vomiting (21.4)	Hepatocellular (100)

Senecio	<i>Senecio spp.</i>	14	USA (28.5), Brazil (21.4), Spain (21.4), Lesotho (14.4)	Mysticism (20)	Ascites (71.4), Jaundice (57.4), Hepatomegaly (42.8), Abdominal distension (21.4), Vomiting (21.4)	Sinusoidal Obstruction Syndrome (64.2), Hepatocellular (35.7)
Turmeric	<i>Curcuma longa</i>	13	USA (38.4), Australia (23)	Health Improvement (20), Dyslipidemia (20), Arthralgias (20), Osteoarthritis (20), Stroke prevention (20)	Jaundice (66.6), Choluria (41.6), Abdominal Pain (25), Pruritus (16.6)	Hepatocellular (100)
Dictamnus	<i>Dictamnus</i>	12	United Kingdom (91.6), Netherlands (8.3)	Eczema (54.5), Psoriasis (27.2)	Jaundice (66.6), Choluria (33.3), Nausea (33.3), Anorexia (25)	Hepatocellular (91.6), Cholestatic (8.3)
St. John's Wort	<i>Hypericum perforatum</i>	12	Germany (58.3)	Depression (62.5)	Jaundice (77.7), Abdominal Pain (33.3), Pruritus (33.3), Choluria (22.2), Fatigue (22.22), Hepatomegaly (22.2)	Hepatocellular (66.6), Cholestatic (25), Mixed (8.3)
Black Catechu	<i>Senegalia catechu</i>	11	USA (81.8), Colombia (9), Germany (9)	Osteoarthritis (40), Arthralgias (30)	Jaundice (80), Choluria (40), Pruritus (40), Nausea (33.3), Vomiting (33.3)	Hepatocellular (72.7), Cholestatic (9), Mixed (18.1)
Ginseng	<i>Panax spp.</i>	10	USA (55.5), United Kingdom (22.2)	Weight Loss (42.8)	Jaundice (60), Ascites (40), Fatigue (30), Nausea (30)	Hepatocellular (70), Sinusoidal Obstruction Syndrome (30)

Pennyroyal	<i>Mentha pulegium</i>	9	USA (77.7), Germany (11), United Kingdom (11)	Abortive (60), Accidental ingestion (20), Vitiligo (20)	Abdominal pain (66.6), Nausea (44.4), Vomiting (44.4), Rash (33.3), Lethargy (33.3), Fatigue (22.2), Jaundice (22.2), Hepatomegaly (22.2)	Hepatocellular (77.7), Steatosis (11.1), Other (11.1)
Cascara-Sagrada; Cascara Buckthorn.	<i>Rhamnus purshiana</i>	9	USA (55.5), Spain (22.2)	Constipation (80), Heath improvement (20)	Jaundice (75), Anorexia (37.5), Choloria (25), Fatigue (25), Pruritus (25)	Hepatocellular (77.7), Sinusoidal Obstruction Syndrome (22.2)
Ashwagandha	<i>Withania somnifera</i>	9	Island (42.8), Canada (28.5), Australia (14.2), Denmark (14.8)	Health improvement (42.8), Anti-anxiety (28.5)	Jaundice (87.5), Pruritus (62.5), Fatigue (50), Abdominal pain (37.5), Nausea (37.5), Choloria (12.5)	Hepatocellular (44.4), Mixed (44.4), Cholestatic (11.1)
Cardo-Do-Visco	<i>Atractylis gummifera</i>	8	France (62.5), Greece (12.5), Tunisia (12.5), Morocco (12.5)	Culinary (37.5), Burn (25), Mysticism (12.5), Weight Loss (12.5), Oxyuriasis (12.5)	Vomiting (87.5), Jaundice (25), Nausea (25), Abdominal pain (12.5)	Hepatocellular (37.5), Other (62.5)
Comfrey	<i>Symphytum Spp.</i>	8	USA (37.5), Germany (25), United Kingdom (25)	Culinary (42.8), Health improvement (28.5)	Ascites (50), Weight loss (37.5), Jaundice (25), Anorexia (25), Hepatomegaly (25)	Sinusoidal Obstruction Syndrome (75), Hepatocellular (12.5), Mixed (12.5)
Babchi	<i>Psoralea corylifolia</i>	8	Hong Kong (37.5)	Vitiligo (80), Irritable Bowel Syndrome (20)	Jaundice (57.1), Vomiting (42.8), Choloria (28.5), Nausea (28.5), Lethargy (28.5), Weakness (28.5)	Hepatocellular (62.5), Cholestatic (25), Mixed (12.5)

Field Horsetail	<i>Equisetum arvense</i>	7	Italy (42.4)	Weight Loss (66.6), Pain improvement (16.6), Health Improvement (16.6)	Jaundice (66.6), Pruritus (50), Fatigue (33.3)	Hepatocellular (71.4), Cholestatic (28.5)
Bitter Orange	<i>Citrus aurantium</i>	7	Italy (50)	Weight (83.3)	Jaundice (85.7), Nausea (66.6), Abdominal Pain (50)	Hepatocellular (85.7), Cholestatic (14.2)
Coneflowers; Equinácea.	<i>Echinacea spp.</i>	6	Australia (16.6), France (16.6), Germany (16.6), Israel (16.6), New Zeland (16.6), USA (16.6)	Health Improvement (50), Pain management (25), Osteoarthritis (25)	Jaundice (83.3), Hepatomegaly (50), Clay stools (33.3), Hematuria (33.3), Lethargy (33.3)	Hepatocellular (100)
Heliotrope	<i>Heliotropium spp.</i>	5	India (60), Germany (40)	Epilepsy (50), Culinary (25), Vitiligo (25)	Ascites (60), Jaundice (60), Abdominal Pain (40), Hepatomegaly (40), Choloria (20), Fatigue (20)	Steatosis (80), Hepatocellular (20)
Saw Palmetto	<i>Serenoa repens</i>	5	USA (60), Germany (20), Italy (20)	Benign prostatic hyperplasia (75), Pain (25)	Jaundice (75), Abdominal Pain (50), Pruritus (50)	Hepatocellular (100)
Ba Jiao Lian	<i>Dysosma pleianthus</i>	4	Taiwan (100)	Dysmenorrhea (33.3), Salivary Gland Tumor (33.3), Pain (33.3)	Diarrhea (100), Vomiting (75), Abdominal pain (100), Nausea (50)	Hepatocellular (100)
Rattlepods	<i>Crotalaria spp.</i>	3	China (33.3), Ecuador (33.3), USA (33.3)	Culinary (50), Pain (50)	Ascites (100), Hepatomegaly (100), Abdominal Pain (66.6),	Sinusoidal Obstruction Syndrome (100)

					Jaundice (66.6), Asterixis (33.3), Nausea (33.3), Vomiting (33.3)	
Impila	<i>Callilepis laureola</i>	3	South Africa (100)	NA	Ascites (100), Hepatomagaly (100), Abdominal pain (66.6), Jaundice (66.6), Asterixis (33.3), Nausea (33.3)	Sinusoidal Obstruction Syndrome (100)
Mistletoe	<i>Viscum spp.</i>	3	Germany (33.3), South Korea (33.3), Germany (33.3)	Anti-stress (33.3), Chronic Prostatitis Syndrome (33.3), Health Improvement (33.3)	Jaundice (66.6), Choluria (33.3), Nausea (33.3), Lethargy (33.3)	Hepatocellular (33.3)
Yerba Mate	<i>Ilex paraguarie nsis</i>	3	Brazil (33.3), Italy (33.3), United Kingdom (33.3)	Weight Loss (66.6)	Abdominal pain (66.6), Hepatomegaly (66.6), Jaundice (33.3), Pruritus (33.3), Ascites (33.3)	Cholestatic (66.6), Sinusoidal Obstruction Syndrome (33.3)
Kombucha Tea	<i>Saccharom yces cerevisiae</i>	3	USA (66.6), Israel (33.3)	Pain (100)	Abdominal Pain (66.6), Choluria (66.6), Jaundice (66.6), Hepatomegaly (66.6)	Hepatocellular (66.6), Cholestatic (33.3)
Soy Isoflavones	<i>Glycine max</i>	3	Spain (66.6), France (33.3)	NA	Asthenia (33.3), Jaundice (33.3)	Mixed (100)
Indian Pennywort	<i>Centella asiatica</i>	3	Argentina (100)	Weight Loss (100)	Abdominal Pain (100), Asthenia (100), Jaundice (100), Hepatomegaly (100), Choluria (66.6), Clay Stools (33.3)	Steatosis (100)

Red Bush Tea; Rooibos.	<i>Aspalathus linearis</i>	3	USA (66.6), Finland (33.3)	Health Improvement (100)	Abdominal Pain (33.3), Choluria (33.3), Jaundice (33.3), Nausea (33.3), Malaise (33.3), Pruritus (33.3), Vomiting (33.3)	Hepatocellular (100)
Milk Thistle; Cardo- Mariano.	<i>Silybum marianum</i>	3	USA (33.3), Ireland (33.3), Netherlands (33.3)	Detoxification (100)	Jaundice (66.6), Choluria (33.3), Clay Stools (33.3), Constipation (33.3), Encephalopathy (33.30, Nausea (33.3), Lethargy (33.3), Pruritus (33.3)	Mixed (66.6), Hepatocellular (33.3)
Peppermint	<i>Mentha pip- erita</i>	2	Germany (33.3), Spain (33.3)	NA	Fatigue (100), Jaundice (100), Asthenia (50), Choluria (50)	Hepatocellular (100)
Coltsfoot	<i>Tussilago farfara</i>	2	Austria (50), France (50)	Osteoarthritis (100)	Abdominal Pain (50), Ascites (50), Hepatomegaly (50), Vomiting (50), Diarrhea (50)	Hepatocellular (50), Sinusoidal Obstruction Syndrome (50)
Horse Chestnut	<i>Aesculus hippocasta- num</i>	2	Spain (50), Japan (50)	NA	Jaundice (100), Hepatomegaly (50), Pruritus (50)	Hepatocellular (50), Cholestatic (50)
Tinospora Crispa	<i>Tinospora crispa</i>	2	France (100)	Diabetes Mellitus (50), Pain (50)	Jaundice (100), Asthenia (100), Abdominal pain (50), Choluria (50), Clay Stools (50), Hepatomegaly (50)	Hepatocellular (100)
Malva	<i>Malva spp.</i>	2	Turkey (100)	Culinary (50), Gastritis (50)	Dyspepsia (50), Nausea (50), Lethargy (50), Headaches (50)	Hepatocellular (50), Mixed (50)

Margosa Oil	<i>Azadirachta indica</i>	2	Canada (50), Malaysia (50)	Multiple Sclerosis (50)	Fatigue (50), Pruritus (50), Vomiting (50), Encephalopathy (50)	Hepatocellular (50), Cholestatic (50)
Oleander	<i>Nerium oleander</i>	1	Turkey	Metastatic Synovial Sarcoma	Abdominal Pain, Ascites, Choloria	Hepatocellular
Maya Nut	<i>Brosimum alicastrum</i>	1	Turkey	HIV	Asterixis, Clay Stools, Choloria, Encephalopathy, Jaundice, Nausea, Malaise	Hepatocellular
Southern Blue Gum	<i>Eucalyptus globulus</i>	1	Germany	NA	NA	Hepatocellular
African Geranium	<i>Pelargonium sidoides</i>	1	Germany	NA	Fatigue	Hepatocellular
Lesser Celandine; Pilewort	<i>Ficaria verna</i>	1	Turkey	NA	Jaundice	Hepatocellular
Kudzu Root Extract	<i>Pueraria spp.</i>	1	South Korea	Health improvement	Jaundice, Choloria	Hepatocellular
Maitake	<i>Grifola frondosa</i>	1	USA	NA	Hypothermia, Coma	Mixed
Adenostyles Alliariae	<i>Adenostyles alliariae</i>	1	Austria	NA	Abdominal Pain, Ascites, Hepatomegaly, Nausea, Vomiting	Sinusoidal Obstruction Syndrome
Mexican Poppy; Cardosanto.	<i>Argemone mexicana</i>	1	Mexico	NA	Abdominal Pain, Fatigue, Jaundice, Hepatomegaly, Nausea, Vomiting	Cholestatic
Willow Bark Tea	<i>Salix spp.</i>	1	Canada	Upper airway infection	Hepatomegaly	Hepatocellular
Sanyak	<i>Dioscorea rhizome</i>	1	South Korea	NA	NA	Hepatocellular

Green Coffee	<i>Coffea canephora</i>	1	Slovenia	NA	Chest Pain, Jaundice, Nausea, Pruritus, Vomiting	Hepatocellular
Chamomile	<i>Matricaria recutita</i>	1	Australia	NA	Ascites	Sinusoidal Obstruction Syndrome
Chaga Tea	<i>Inonotus obliquus</i>	1	USA	Anticoagulati on	NA	Mixed
Yang Ti Cao; Serralhinha.	<i>Emilia sonchifolia</i>	1	Hong Kong	Health Improvement	Abdominal Distension, Hepatomegaly	Other
Java Tea	<i>Orthosipho n aristatus</i>	1	Spain	Weight Loss	Fatigue, Jaundice, Pruritus	Hepatocellular
Black Nightshade	<i>Solanum nigrum</i>	1	India	Digestion	Abdominal Pain, Hepatomegaly, Jaundice	Hepatocellular
Mango	<i>Mangifera genus</i>	1	Yemen	Pain	Abdominal Pain, Anorexia, Fatigue	Hepatocellular
Isabgol; Psyllium.	<i>Plantago psyllium</i>	1	Italy	Constipation	Anorexia, Fatigue, Hepatomegaly, Jaundice	Hepatocellular

† () percentage values; NA: not available; USA: United States of America