

Welcome



USP Adulterants Database

Beyond the Looking-Glass

Anton Bzhelyansky Scientific Liaison Dietary Supplements and Herbal Medicines United States Pharmacopeial Convention (USP)

GInAS 2017 Meeting October 11, 2017



Rockville, Maryland USA

Overview



- Introduction to USP
- USP Adulterants Database
- Database Demo (time permitting)
- Challenges of Cheminformatics (bonus)



Introduction to USP



Working with Partners

From the standards we create to the partnerships and conversations we foster, our scientists, advocates and network of experts are critical to constructing and reinforcing the foundation for quality in health and healthcare.

Building a Healthier World



United States Pharmacopeial Convention



- Private, not-for-profit organization (since 1820)
- Establishes and disseminates public written
 standards for quality, purity, identity, strength, and
 labeling of medicines, dietary supplements and foods
- Provides physical Reference Standards in support of tests in the USP–NF, DSC, HMC and FCC
- Educates manufacturers, practitioners, and others seeking information on quality and USP standards





USP Adulterants Database*

*working title, subject to change



Adulterants Database Precursors



- ▶ USP Workshops on DS and Food Adulteration (2011 2015)
- USP Expert Panel on Adulteration (July 2013 Present)
- USP General Chapter <2251> (Official: August 1, 2016)
- Dietary Supplements Stakeholder Forums (2011, 2013, 2015)
- USP Dietary Supplements Strategic Initiative (2014)
- ► Taiwan Conference on Adulteration of DS (2015, 2017)

Adulterants Database



- Relational database developed in Drupal
- Five primary entities:
 - Instance of Adulteration (Report)
 - Adulterant
 - Product
 - Primary Company
 - Secondary Company (dba)
- Numerous Fields for Each Entity



Adulterants Database – Primary Relationships

One-to-one



One-to-many



One-to-one



[Recall]

One-to-many



[Mfg, Dist, Retailer]

One-to-many



Adulterants Database – Snapshot









1200 Reports

Reference Source Type

- Agency Report (1109)
- Journal Article (73)
- Website (39)
- Application Note (4)
- Media Broadcast (1)
- Media Electronic (1)

2001 Compounds

Adulterant Functions

- stimulant (260)
- androgen/anabolic (191)
- opioid analgesic (144)
- narcotic (126)
- erectogenic (118)
- · synthetic cannabinoid (94)
- · antiinflammatory (86)
- antihypertensive (84)
- cathinone (83)
- analgesic (72)

Show more

1050 Products

1 Intended Use(s)

- · Sexual Enhancement (458)
- Weight Loss (386)
- Sports Performance Enhancement (156)
- Antirheumatic (25)
- Other (21)
- Psychoactive Substance (3)
- Sleep Aid (2)
- Antidiabetic (1)

700 Companies

♠ Company

- r thomas marketing llc (26)
- · bodybuilding.com, Ilc (24)
- novacare, Ilc (24)
- · hi-tech pharmaceuticals, inc. (18)
- a & h focal, inc. (15)
- atlas operations, inc. (13)
- organic herbal supply, inc. (13)
- usplabs, llc (11)
- bee extremely amazed llc (10)
- · shh trading pte. Itd (10)
- mynicknaxs, Ilc. (9)
- universal abc beauty supply international, inc. (8)
- · vmg global inc. (8)

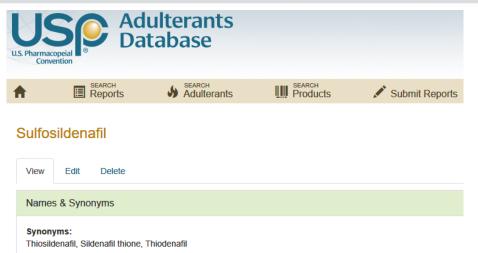
Adulterants Database – Data Sources



Primary data streams:

- 1. Product-focused "Agency reports":
 - a. FDA (USA)
 - b. Health Canada
 - c. TGA (Australia)
 - d. HSA (Singapore)
 - e. National agencies of Hong Kong, South Korea, Japan, Taiwan, Czech Republic, etc.
- Compound-focused databases: SWGDRUG (US DEA), NIST (USA), Project <u>RESPONSE</u> (Europe), <u>Forendex</u>, etc.
- 3. Peer-Reviewed Literature original and review articles. Typically, target a specific adulterant (e.g., structure elucidation), a class of adulterants, or describe a particular analytical methodology.
- 4. Media reports industry press, online publications, conventional print media, TV and radio broadcasts. Limited in number and used to provide additional context.
- 5. Specialized documents court proceedings, scientific reports, application notes, etc.

Adulterant Record – Disambiguation



Affected foreign products

Chao Jimengnan SuperPowerful Man tablets

Product

Maxidus capsules

Fu Yuan Chun capsules

Qing Tian Zhu tablets

AS Number:	479073-79-5₺
NII Code:	HRL3FWK7XV _®
Chi Key:	LJUBVCQVNMLSTQ-UHFFFAOYSA-N ்
EGG Code:	Search KEGG Databases
ubChem ID:	10228242광
TC Code:	Search ATC ₪

15 mg of product per tablet).







Undeclared Ingredient

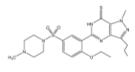
sulfosildenafil

tadalafil

tadalafil

sildenafil

sulfohydroxyhomosildenafil



2007). These products were composed of 2 or 10 sand-colored tablets (13-

Adulterant Record – Features

Phenolphthalein



-aa

Delete

Names & Synonyms

Chemical Name (IUPAC): 3,3-bis(4-Hydroxyphenyl)-2-benzofuran-1-one

A06AB04 ₽

Synonyms:

3,3-bis(4-Hydroxyphenyl)-1(3H)-isobenzofuranone, 3,3-bis(4-Hydroxyphenyl)-2-benzofuran-1(3H)-one, 3,3-Bis(4-hydroxyphenyl)phthalide, 3,3-bis(p-Hydroxyphenyl)phthalide, alpha-di(p-Hydroxyphenyl)phthalide

Other Names:

ATC Code:

Agoral, Alophen, Chocolax, Colax, Doxan, Doxidan, Espotabs, Euchessina, Evac-q-kwik, Evac-q-tabs, Evac-u-gen, Evac-v-lax, Ex-lax, Feen-a-mint, Femilax, Koprol, Laxin, Laxogen, LILO, Medilax, Phenolax, Phthalimetten, Phthalin, Prulet, Purga, Purgen, Purgophen, Spulmako-lax, Trilax

Identification Codes			
CAS Number:	77-09-8ଜ		
UNII Code:	6QK969R2IF₽		
InChi Key:	KJFMBFZCATUALV-UHFFFAOYSA-N®		
KEGG Code:	D05456⊮		
PubChem ID:	4764∉ੋ		





DB May be searched by:

- Any of the synonyms
- Any of the acronyms, trade, commercial or 'street' names
- Any of the chemical identifiers: CAS, UNII, InChI Key, KEGG Code, PubChem ID
- nearly every iteration of the name
- DEA codes and schedules

Adulterant Record - Features (cont.)

Tramadol



Delete

Delete

Names & Synonyms

Chemical Name (IUPAC): (1RS,2RS)-2-(Dimethylaminomethyl)-1-(3-methoxyphenyl)cyclohexanol Synonyms:

(+)-Tramadol, 2-[(Dimethylamino)methyl]-1-(3-methoxyphenyl)cyclohexanol, DEA No. 9752, DEA Schedule IV

Other Names

Acema, Actidol, Acty, Acugesic, Adamon, Admadol, Adolonta, Altadol, Amadol, Amadol, Ammitran, An-Tian, Ana-Q, Anadol, Analtram, Anangor, Anatram, Andalpha, Arodol, Arrestadol, Astradol, Atdol, Avdol, Axytram, Bei Pin, Bestodol, Bing Ning, Biodalgic, Biodil, Biodol, Biomadol, Biotram, Biotrama, Biotrany, Boldol, Bolodol, Bramadol, By-madiol, Cadol, Calmador, Calmador, Camadol, Cambidol, Cambidol, Centrasic, Ceparidin, CG-MAC, Cincro Plus, Citra, Clog, Combitram, Contradoc, Contram, Contramal, Conzip, Cormadol, Corsadol, Cortram, Cosdol, Crispin, Cromatodol, Cruzac, Cuntrol, D.M.Dol, Da Ma Err, Damadol, Damed, Damol, Darol, Didol, Doctramado, Dolan, Dolana, Dolbest, Dolbis, Dolfi, Dolfre, Dolga, Dolgesik, Dolma, Dolma, Dolmax, Dolmar, Dolocap, Dolocet, Dolodol, Doloned, Dolomed, Doloria, Doloran, Dolotram, Dolotram, Dolotramin, Dolotramine, Dolpar, Dolpar, Dolpar, Dolpar, Dolsta, Dolsta, Dolta, Dolta, Dolta, Dolta, Dolta, Dolta, Dolta, Dolta, Dolotram, Do Domadol, Dorless, Drobit, Dromadol, Durodor, Durotram, E-Dol, Ecodolor, Eltram, Esgipyrin, Etigesic, Eufindol, Fada Tramadol, Febrex, Feng Tong Ding, Forgesic, Formador, Fortradol, Fraxidol, FS, Gelotradol, Gemadol, GenRX Tramadol, Getpar, Glimadol, Gudil, Haldotram, Haledol, Hetradol, Hua Jie Wei, Hua Qu, Hyperdol, Idol, Imadol, Indolpara, Ingesic Forte, Inodol, Iodol, Ivydol, Ixprim, Jetra, Jpdol, KAlma, Kamadol, Katrasic, Kdol-P, Kedol, Kevtram, Kontram, Lanalget, Le Shi Pu Kang, Leedol, Lexidol, Lucidol, Lucidol, Merdol, Mabron, Madol, Madola, Mandolgin, Manol, Mapdol-P, Marodol, Medol, Meradol, Meridol, Metazac, Metracop, Microdol, Milador, Minidol, Mipro, Mobiya, Monoalgic, Monocrixo, Muaction, Nettram, Neutram, Nictram, Noax, Nobligan, Nomal, Nonalges, Nopidol, Notil, Novadol, Nufapotram, Nycodol, OC-Dol, Odel, Omodel, Ondol, OPI-OT, Opidol, Opiqesic, Orasic, Oratram, Orchidol, Orozumadol, Osdol, Osmadol, Ospidol, Oxxalgan, Ozitram, Pacmadol, Painadol, Paindol, Paine, Painlax, Paratel-P, Patradol, Patral, Paxilfar, Paxmax, Pengesic, Penover, Pinorec, Plazadol, Poltram, Postadol, Predalgic, Prontalgin, Prontofort, Protradon, PTR, Qi Zhi, Qimaite, Qu Feng, Qu Ming, Qu Teng, Qu Tong Kang, Qutong, Racetram, Rajdol, Ralgen, Ralivia, Ramadol, Ramax, Redimol, Relidol, Ridil, Rofy, Rotamol, Rui Li Ping, Rybix, Ryzolt, Sayadol, Sefmal, Seminac, Sensitram, Servodol, Siatram, Sigmadol, Simatral, Simudol, Sintradon, Slovadol, Siatram, Sigmadol, Sigma Souladol, Soztram, Splint Forte, Sridol, Stemadol, Strom, Supridol, Sylador, T Dol, T-Long, Tacil, Tanadol, Tadol, Tai Mei Ding, Takadol, Takol, Talnex, Tamadol, Tamolan, Tamriv, Tamvrin, Taridol, Taxidol, Taz, TDL, TDX, Tecsadol, Temadol, Tendia, Teramadol, Theradol, Ti Ma Er, Tial, Timarol, Timasen, Tinlol, Tioner, Tiparol, Tlusic, TM-Plus, TMD, Tofdol, Tol-A, Tolma, Tong Ting, Topalgic, Toptra, Trabar, Trabilin, Trace, Traceta, Tracin-P, Tracine, Tractadol, Tradef, Tradol, Tradolan, Tradolgesic, Tradolor, Tradonal, Tradorec, Tradosik, Tradyl, Traflash, Tragesic, Tragesik, Trail, Trak, Tral-ac, Tralenil, Tralgiol, Tralgiol, Tralgic, Tralodie, Tram-Proxyvon, Trama, Tramabene, Tramabeta, Tramacalm, Tramacalm, Tramacaet, Tramacet, Tramacip, Tramacon, Tramaconti, Tramacotil Uno, Tramacure, Tramacure, Tramada, Tramader, Tramadex, Tramadin, Tramadin, Tramado, Tramadol, Tramadolo, Tramadolo, Tramadolo, Tramadon, Trama Tramaforte, Tramag, Tramagem, Tramagesic, Tramagetic, Tramagit, Tramahexal, -Tramake, Tramakind, Tramaklosidol, Tramal, Tramalan, Tramalek, Tramalex, Tramalgic, Tramalgin, Tramalin, Tramamed, Tramamerck, Tramanil, Tramapine, Tramasol, T Tramatyrol, Tramazac, Trambax, Trambo, Rramcod, Tramcontin, Tramdop, Tramed, Tramedo, Tramedo, Tramedon, T Trami, Tramico, Tramisol, Tramiolin, Tramiet, Tramned, Tramnom, Tramo, Tramoda, Tramodin, Tramoflex, Tramolin, Tramolin, Tramospas, Tramrot, Tramp, Tramplas, Tramquel, Tramrod, Tramsars, Tramtor, Tramundal, Tramundin, Tramy, Tranal, Tranat, Trandol, Trandy, Transc, Tranzen, Trapain, Trapain, Trapain, Trapain, Trapain, Trapain, Trapain, Trapain, Transcen, Transce Trasic, Trasik, Travex, Travictol, Trawel, Traxdol, Trazac, Trazodec, TRD, Treat, Tremolo, Tremtec, Trexol, Tridural, Trodon, Trofel, Trol, Troma, Tromar, Tromy,

Tropidol Trosic Trugesic Truges Truge Truge Truge Truge Trugedol Ultragon Ultragon Ultragon Ultragon Ultragon Unidol Unitragon Unidol Unitragon Unitragon Unidol Unidol Unitragon Unidol Unitragon Unidol Unidol Unitragon Unidol Unidol





- (could be a lot of) Synonyms
- Chemical Identifiers

dentification Codes		
CAS Number:	27203-92-5 ਵੀ	
UNII Code:	39J1LGJ30J ള	
InChi Key:	TVYLLZQTGLZFBW-ZBFHGGJFSA-N₽	
KEGG Code:	D08623₽	
PubChem ID:	33741 ਦੀ	
ATC Code:	N02AX02₫	

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Adulterant Record – Features (cont.)

Other Descriptors

Adulterant Functions: analgesic, narcotic

Adulterant Source: Synthetic Chemical Formula : C16H25NO2

Molecular Weight: 263.381

References

Monograph(s):

Monograph: Attachment: USP - Tramadol Hydrochloride

usp_-_tramadol_hydrochloride.pdf

Monograph: Attachment:

Monograph: USP - Tramadol Hydrochloride Tablets
Attachment: Jusp - tramadol hydrochloride tablets.pdf

....

USP - Tramadol Hydrochloride Extended-Release Tablets

| usp - tramadol hydrochloride extended-release tablets.pdf

USP - Tramadol Hydrochloride Compounded Oral Suspension

usp - tramadol hydrochloride compounded oral suspension.pdf

Monograph: Attachment:

USP - Tramadol Hydrochloride Compounded Oral Suspension, Veterinary

ig usp - tramadol hydrochloride compounded oral suspension veterinary.pdf

Monograph: Attachment:

Reference Standard(s):

Reference Standard:

USP Tramadol Hydrochloride RS

1672600

Item Number:

Other Reference(s):

SWGDRUG &, WHO - Critical Review (2014) &, Wikipedia &, DrugBank &, m/z Cloud &, Designer Drugs Online &, DEA Schedule &, Tramadol—A True Natural

Product? &, Occurrence of the Synthetic Analgesic Tramadol in an African Medicinal Plant &, Occurrence of the Synthetic Analgesic Tramadol (Supporting Information) &





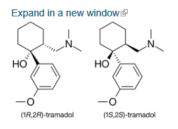
- Other attributes: functions, formula, molecular weight
- References to compendial sources whenever appropriate
- Sources of Reference Materials whenever available
- References to other sources, databases, online resources, research literature

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@ 2017 LISP

Adulterant Record – Features (cont.)

Additional Information



Search for Reports containing:Tramadol

Attachment(s): 7 6 1 tramadol update.pdf

occurrence_of_the_synthetic_analgesic_tramadol_in_an_african_medicinal_plant_2013.pdf

doccurrence_of_the_synthetic_analgesic_tramadol_in_an_african_medicinal_plant_2013_-_supporting_info.pdf

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• Technically, the database is equipped to accept any file, this is just underutilized for now. [e.g., NMR FIDs]





- Additional Information:
 Chemical structure;
 essentially, any kind of a graphics file, including chromatograms, spectra
- Search for Reports containing Tramadol
- Attachments relevant and informative, handpicked: toxicology, pharmacology, analysis

Product Record – Features





Spirodex

View

Edit

Delete

About

Intended Uses: Weight Loss, Sports Performance Enhancement

Product Ingredients:

Caffeine. SPIROBLEND (A proprietary blend consisting of Oxytropis *falcate* extract [whole plant] standardized for biogenic amine content, Camelia *sinensis*, hordenine, Geranium *maculatum* extract [leaves and stems] standardized for 4-methylhexan-2-amine content. Paullinia *cupana* [whole plant] extract standardized for purine content). **Other Ingredients**: Dicalcium Phosphate, Stearic Acid, Magnesium Stearate, Croscarmellose Sodium, Silicon Dioxide, Hydroxypropyl Methylcellulose (E5 and E15), Polyethylene Glycol, Water, Titanium Dioxide (color), FD&C Yellow No. 5.

Product Website: Gaspari Nutrition, Spirodex, Mood Enhancing Stimulant Compound, 425 mg, 60 Tablets &

Company Information

Distributor

Gaspari Nutrition, Inc.

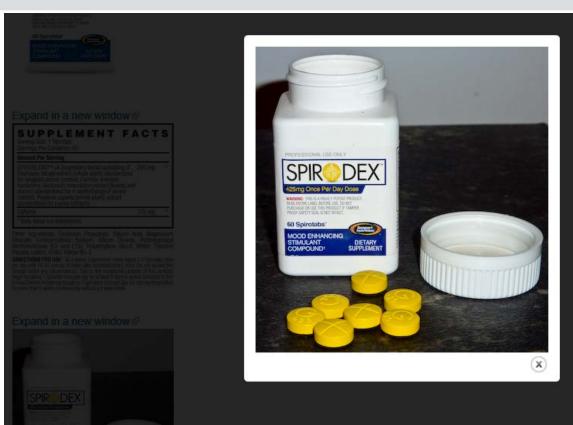
Manufacturer

Hi-Tech Pharmaceuticals, Inc.

Product record includes:

- Intended use(s)
- List of declared ingredients as appears on the package
- Link to the product website, if still active
- Information on the companies affiliated with the product: Manufacturer, Distributor, Retailer (this information is not readily available and requires research)

Product Record - Features (cont.)







Product record also includes:

- Images of the product packaging
- Supplement Facts panel, preferably with legible list of ingredients
- Images of the individual dosage units, specifically highlighting any features
- Link to the database records referencing the product
- Magnification and full-sized images

Report – Features

Workflow History

Report 25276

View





Reference Source Information

Reference Source Type: Agency Report
Specific Subject or Title of Reference Source:
Gadget Island, Inc. Dba Gear Isle Issues Voluntary Nationwide Recall of Dietary Supplements Due to Undeclared Active Pharmaceutical Ingredients
Reporting Agency: United States - Food and Drug Administration (FDA)
Reference Source URL: Reference Source Information Online Reference Source Date: Sep 20, 2017

Adulterated Product Information

Delete

Adulterants and products

Product: Grande X
Adulterant(s):

Sildenafil, Tadalafil, Desmethylcarbodenafil

Report record includes:

- Report number (could be searched on)
- Reference type (agency report, article, app note, etc.)
- Title of the reference
- Link to the original URL
- Date the original notice, or journal article, was published
- Product found to be adulterated
- The list of adulterants

Report – Features (cont.)

Recall Information

Recall Firm: Gadget Island, Inc.

UPC Code: 640793555440 ₽ Container Type: Blister card

Containers per Package: 30
Doses/Servings per Container: 1

Dosage/Serving Size: 1 Capsule

Unit Dose Form/Type: Hard-shell capsule

Unit Dose Color: gold

California Department of Public Health &

Additional Information

Attachments: 2017-09-20_-_fda_-_gadget_island.pdf

2017-09-20 - fda - cdph- gadget_island.pdf

2017-10-04_-_fda_-_recall_-_gadget_island.pdf

Administrative Information

Workflow: Approved





Report record includes:

- Company conducting the recall
- Specific features of the recalled product: UPC code, description of the packaging configuration and the unit dose features
- Concurrent recall notices issued by other agencies
- Copies of the relevant recall-related notices as pdfs

Company Record – Features

Hi-Tech Pharmaceuticals, Inc.

Delete





Doing Business As:

Applied Lifescience Research (ALR) Industries, Inc.

Advanced Performance Supplements (APS) Nutrition

LG Sciences

FormuTech Nutrition

Innovative Labs

iForce Nutrition

Sports One

Nittany Pharmaceuticals, Inc.

Prime Nutrition

Tribravus Enterprises, LLC.

Company Address and Contact Information

Address

Address Type: Street Address: Mailing Address 6015 B Unity Drive

City:

Norcross GA

State/Province: Zip/Postal Code:

30071

Country:

United States of America (the)

Contact Information

Phone(s):

Phone Type: Toll-free

Phone Number: <u>+1 (888) 855-7919</u>

Company record includes:

- List of 'brands', 'labels', aliases (dba) under which the products appear
- Address(es) verified by specialized websites (BBB, Bloomberg, Corporation Wiki, Bizapedia, Manta, etc.)
- Contact Information (phone numbers, email)

Company Record – Features (cont.)





Other Information

URL(s):

Hi-Tech Pharmaceuticals @

About Hi-Tech Pharmaceuticals &

Hi-Tech Pharmaceuticals Wins Crucial Court Motions Against Harvard Doctor, Dr. Pieter Cohen, in a Slander Lawsuit Seeking \$200 M @

Hi-Tech Pharmaceuticals acquires co-ownership of Prime Nutrition ₪

The Truth Behind Hi-Tech Pharmaceuticals with Jared Wheat (Video) №

Feature on Jared Wheat, CEO and President of Hi-Tech Pharma@

Press Release: Hi-Tech Pharmaceutical Sues Pieter A. Cohen for \$50 Million in Compensatory Damages &

Press Release: Hi-Tech Pharmaceuticals Acquires Top Secret Nutrition Adding to its Premier Line Up of Sports Nutrition Brands &

Notes/Comments:

In 2015, Hi-Tech Pharmaceuticals acquired a handful of inventive brands, including iForce Nutrition, Nittany Pharmaceuticals, Advanced Pharmaceuticals and Nutritionals. FormuTech Nutrition, LG Sciences, and Innovative Laboratories.

Jared Robert Wheat - CEO, President, and Chairman

Stephen D. Smith & - Executive VP

John Brandon Schopp - Director of Contract Manufacturing

Timothy Fulmer & - Corporate Counsel/Legal

Attachment:

red-yeast-rice-indictment-dmaa-no-sell-order-20171004.pdf

red-yeast-rice-indictment-2-2017-0928.pdf

red-yeast-rice-indictment-20170928.pdf

Company record includes:

- Links to business and relevant public information about the company
- Trademarks registered by the company and company owners
- Company principals and their relevant business endeavors, including other business ventures
- Documents relevant to the company's business (here: court documents)

Challenges to Overcome



- Significant data fragmentation.
- A lot of intentionally misleading and frequently unverifiable information: lists of nonexistent ingredients, ridiculous claimed amounts, made-up company addresses, 'formulation' changes are frequent, and that may include the adulterants.
- The fleeting nature of web data: recalled products and entire sites typically disappear within hours of the published notifications. Need to resort to Google Cache and Wayback Machine to get useful descriptive information.
- Adulterants and product names may appear in different languages:
 - the translated product names may be unrecognizable to the original language carriers ('Old Chinese')
 - Adulterants may persist for a long time in one language without being translated into others. Ergo, the same adulterant
 may be 'discovered' numerous times and persist for a long time as 'parallel' entities with duplicative chemical identifiers
 - Difficulties with relying on 'machine translation' Thebacon (dihydrocodeinone enol acetate)
- Counterfeit adulterated products
- Challenges in cheminformatics roughly 10% of the records in the database required reconciliation: names, CAS registry numbers, InChI Keys may be for other stereoisomers, or hydrates, or salts

Proposed Solutions / Improvements



- Rapid exchange of new adulterant information. Work with FDA SRS to issue UNII; make chemical characterization data available via a common platform as soon as practicable, coordinate confirmatory analysis by other labs if the sample amount permits this
- Make product information available: full package, closeup of unit dosage forms, etc.
- Expanded analysis details to the extent possible, including chromatograms and spectra, experimental conditions including equipment specifics, settings, parameters, etc.
- Make content of peer-reviewed journal publications available, e.g., through Creative Commons mechanisms. Share data.

Acknowledgements



- Lynn Haas
- Michael Lynch
- Gabriel Giancaspro
- George Wang
- Dennis Gorecki
- Nam-Cheol Kim



In memoriam





Mark Roman, PhD 1968 - 2014

USP Biannual Adulteration Workshop February 27-28, 2018

Rockville, Maryland



Questions



Stay Connected

(301) 230-6303 | ANB@usp.org





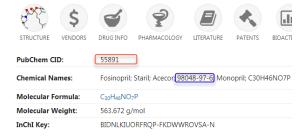
Challenges of Cheminformatics*

*bonus

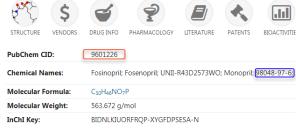
Fosinopril



Fosinopril



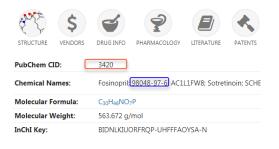
Fosinopril



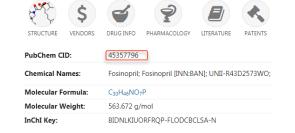
Fosinopril



Fosinopril



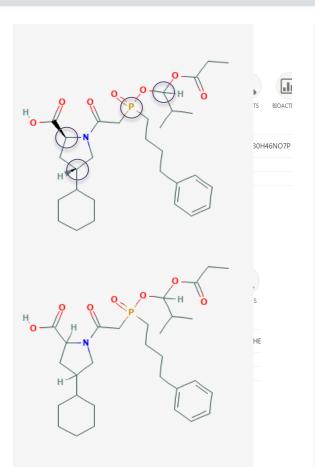
Fosinopril

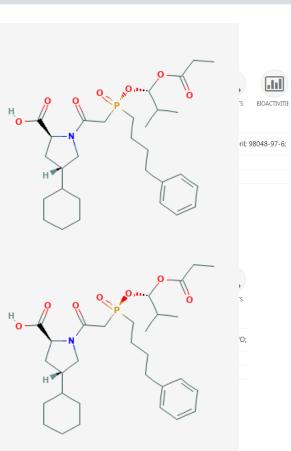


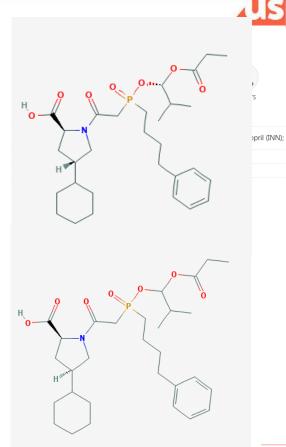
Fosinopril



Fosinopril







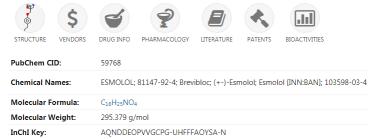


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Esmolol



ESMOLOL

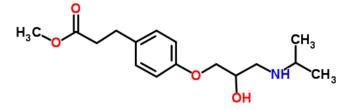






Substance Name: Esmolol [INN:BAN] RN: 81147-92-4 UNII: MDY902UXSR

InChlKey: AQNDDEOPVVGCPG-UHFFFAOYSA-N



Listed CAS may be that FDA UNII i MDY902UXSR Other Registry Numbers

i 84057-94-3

Related Registry Number

Registry Numbers **CAS Registry Number**

i 81147-92-4

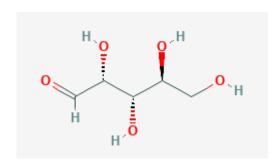
i 103598-03-4

181161-17-3 (Hydrochloride)

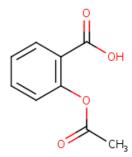
of a hydrate or a salt

Arabinose and Aspirin





3.3 Other Identifiers	X 2
3.3.1 CAS	× c
147-81-9	▶ from ChemIDplus
5328-37-0	▶ from ChemlDplus, DrugBank, EPA DSStox
20235-19-2	· Irom chemiopus, brugount, El A baston



TOXNET > ChemIDplus > Substance



Start New Query

Substance Name: Aspirin [USP:BAN:JAN]

RN: 50-78-2

UNII: R16CO5Y76E

InChlKey: BSYNRYMUTXBXSQ-UHFFFAOYSA-N

Registry Numbers

CAS Registry Number

i 50-78-2

FDA UNII

i R16CO5Y76E

Other Registry Numbers

i 11126-35-5

i 11126-37-7

ACHIRAL

▶ from European Chemicals Agency - ECHA

i 2349-94-2 i 26914-13-6 i 98201-60-6

報道発表資料 2017年08月22日 福祉保健局

医薬品成分を含有する製品の発見について

都では、いわゆる健康食品による健康被害発生の未然防止のため、都内で販売等される製品の調査及び成分検査を行っています。

今般、下記5製品の成分検査を行ったところ、医薬品成分である「イブプロフェン」、「シルデナフィル」及び「ジメチルジチオデナフィル」が検出され、

いわゆる健康食品において医薬品成分を含むものは医薬品とみなされ、厚生労働大臣の承認を受けずに製造販売することは、「医薬品、医療機器等の品 質、有効性及び安全性の確保等に関する法律」(以下、「医薬品医療機器等法」という。)で禁止されています。

なお、これまでに当該製品による健康被害発生の報告は受けていません。

製品概要

品名	形状	検出成分
スーパーイヌトウキ	カプセル	イブプロフェン
藏秘强肾王	丸剤	シルデナフィル
Dark Horse 黑馬	カプセル	シルデナフィル
ROCK SOLID	カプセル	ジメチルジチオデナフィル
PRO enhance	カプセル	ジメチルジチオデナフィル

ROCK SOLID







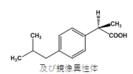
PRO enhance



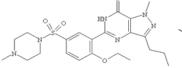




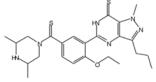




【イブプロフェン】



【シルデナフィル】



【ジメチルジチオデナフィル】

報道発表資料

Welfare and health department, 22 August 2017

On the discovery of products containing pharmaceutical ingredients

In the capital, in order to prevent the occurrence of health damage caused by so-called health foods, we conduct surveys of products sold in the city and inspection of ingredients.

As a result of conducting the ingredient tests of the following 5 products, the pharmaceutical ingredients "ibuprofen", "sildenafil" and "dimethyldithiodenafil" were detected.

Those containing pharmaceutical ingredients in so-called health foods are regarded as pharmaceuticals, and manufacturing and selling without the approval of the Minister of Health, Labor and Welfare is a "law on securing quality, efficacy and safety of pharmaceuticals, medical devices, etc." (Hereinafter referred to as the "Medical Equipment and Medical Equipment Act. etc.").

In addition, we have not received any report on the occurrence of health damage by the product so far.

Product Summary

Item name	shape	Detectable component
Super Inu	capsule	Ibuprofen
Secret kidney king	Pill	Sildenafil
Dark Horse uma	capsule	Sildenafil
ROCK SOLID	capsule	Dimethyldithiodenafil
PRO enhance	capsule	Dimethyldithiodenafil

ROCK SOLID







PRO enhance



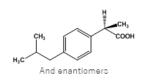




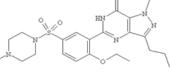


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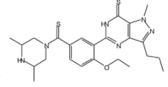
	Item na		ROCK SOLID
		name	Food containing Ezo Bogi extract
	Display content	Raw material name	Orind Make Incorporated
(Four)			Kanagawa prefecture Yokohama-shi Aoba-ku Eida-cho 1151-40 102
			Ezo Houkogi, Seiyushi Willi, Jujube, Cayenne, Nettle, pumpkin seed, Oats
	S	hape	capsule
Detectable component		le component	61 milligram detection of dimethyldithiodenafil in 1 capsule
	Item name	Item name	PRO enhance
		name	Processed food containing concentrated ginseng
	Display content	Seller	Orind Make Incorporated
(Five)	Seller	Kanagawa prefecture Yokohama-shi Aoba-ku Eida cho 1151-40-102	
		Raw material name	Ginseng, Fort, Gotzkola, Saw Palmetto, Menthol
	s	hape	capsule
	Detectable component		61 milligram detection of "dimethyldithiodenafil" in 1 capsule



[lbuprofen]



[Sildenafil]



[Dimethyldithiodenafi



dimethyldithiodenafil



All

deos

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About 34 results (0.83 seconds)

Did you mean: dimethyl dithio denafil

Lambada = RUSH ', LADY PAPILLON 'कार - !!! 나 here 안 안전 처.

https: //m.mfds.go.kr/renewal/.../menuForward.jsp? ... ▼ Translate this page
Jan 19, 2016 - 무허가 의 식당 인 'Lambada = RUSH', 'LADY PAPILLON' 에서 **Dimethyldithiodenafil** 이 검 출 에 [제품 정보] * 제품 1 - 제품 에: Lambada = RUSH

Partial revision of "Criteria on Scope of Pharmaceuticals"

https://search.e-gov.go.jp/servlet/PcmFileDownload? seqNo ... - Translate this page ATP. Adenosine-5'-triphosphate. Dimethyl dithiodenafil dimethyldithiodenafil . Name. Other name etc . Remarks etc .. Remarks. Olive bear. Aobana / Tsukiksa / Gyokubuna / Commelina.

[PDF] Medicine Genesis 1012 No. 1 .pdf (133KB)

www.pref.tokushima.jp/docs/2016042200481 /.../ 20161012001.pd ... ▼ Translate this page Oct 12, 2016 - Acid. Attachment 2 of 3. Others (chemical substances etc.) Table Hydrobromide Dextromethorphan Next, add as follows. Dimethyldithiocarbamate de. Nafiru. Dimethyldithiodenafil .

[PDF] January 2017 issue No. 267 - Hiroshima Prefecture Pharmaceutical A... www.hiroyaku.or.jp/pdf/journal/No 267.pdf ▼ Translate this page

Jan 1, 2017 - Attachment 2 of 3. Following addition of dehydrotromethorphan hydrobromide of other (chemical substances etc.) as follows: Dimethyldithiocarbamate Dinah fill. **Dimethyldithiodenafil**.

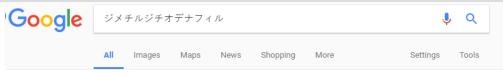
[PDF] About revision of criteria on the scope of pharmaceuticals - Ministry o...

https://kouseikyoku.mhlw.go.jp/.../20161012iyakuhinnohani.pdf ▼ Translate this page
Oct 12, 2016 - Acid. Attachment 2 of 3. Others (chemical substances etc.) Table Hydrobromide
Dextromethorphan Next, add as follows. Dimethyldithiocarbamate de. Nafiru. Dimethyldithiodenafil

제공 기관스 검색 X - 스마트 컨슈머 - 소비자 종합 정보망@

smartconsumer.go.kr/user/is /.../ MoreViewSearchBbsList2.do? ... Capt ... ▼ Translate this page 국발 사유 무허 가 의 식당 인 'lambada = rush' 'lady papillon' 에서 dimethyldithiodenafil 이 검출 를 · 미, 아 세트 아미 펜 펜 성명 버유 중성 에서 유 사실 국발 2015-07-14.

Products with this adulterant have been documented in Japan and Korea since at least late 2015, i.e., for almost two years. Meanwhile, there appears to be no mention of this compound in the Englishlanguage literature.



Welfare on December 18, 2013), hydroxythio Homosildenafil) was detected.

About 462 results (0.35 seconds)

Attention to unapproved drugs including sildenafil (ingredients similar to s...

hfnet.nih.go.jp/contents/detail3132.html ▼ Translate this page

Dec 18, 2015 - Analysis of the product sold on the internet revealed that sildenafil-like ingredient (

dimethyldithiodenafil (designated as a pharmaceutical ingredient by the Ministry of Health, Labor and

[PDF] About finding products containing pharmaceutical ingredients - Minist...

www.mhlw.go.jp/file/04...11126000.../tokyo-press-kidora.pdf ▼ Translate this page
Dec 18, 2015 - dimethyldithiodenafil "was detected (designated by the Ministry of Health, Labor and
Welfare as a pharmaceutical ingredient on December 18, 2015). "Hydroxythiohomosildenafil", which has
been conventionally designated as a pharmaceutical ingredient, was also detected from one product
among them.

Please pay attention to health foods etc where pharmaceutical ingredient...

www.pref.okayama.jp/page/detail-86420.html ▼ Translate this page
(3) Shouty MORE. Dimethyldithiocarbamate Dinah fill . Dimethyldithiocarbamate Dinah fill . Dimethyldithio Dinah fill , hydroxy thio homo sildenafil. H27.12.18, Tokyo [PDF file / 313KB].勃王KING Star. Hydroxy thio homo sildenafil. Chiodenafiru. H 27. 12.18 ...

What is homo thiodenafil - chemical substance dictionary Weblio dictionary

www.weblio.jp> learning> chemical substance dictionary ▼ Translate this page
Weblio dictionary> academic discourse> chemical substance dictionary> meaning and explanation of
homo thio denafir 1 - methyl - 3 - propyl - 5 - [2 - ethoxy - 5 - (4- ethylpiperazinosulfonyl) phenyl] - 1 H pyrazolo [4,3-d] pyrimidin-7 (6H) -thione , 1- methyl -3-propyl-5 - [2-ethoxy-5 - (4-ethylpiperazino ...

Detection of pharmaceutical ingredients from health food | - Mario Clinic

marioclinic.com/info/ Detection of pharmaceutical ingredients from health food / ▼ Translate this page Aug 25, 2017 - which is the main component of Viagra has a "sildenafil" or similar chemical structure " dimethyl dithio Dinah fill is due to the product has been manufactured and sold which contains".

Recently in the United States as "Desmethyl · Carbodia Phil", still sildenafil ...

Note that the number of relevant searches on the Japanese spelling exceeds that in English more than tenfold. We may conclude that this adulterant is fairly wellknown in Japan.

ジメチルジチオデナフィル (Let's be Chemists!)



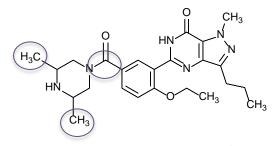
$$\begin{array}{c|c} & & & \\ &$$

H₃C O O HN N N CH₃

ジメチルジチオデナフィル

Sildenafil

Dimethylsildenafil



Thiodenafil

Carbodenafil

Dimethylcarbodenafil

ジメチルジチオデナフィル (Let's be Chemists!)



ジメチルジチオデナフィル

Dithiodimethylcarbodenafil

Searched:

- -SciFinder (CAS Chemical Abstracts)
- -structure and sub-structure searches in available databases (check out MolView!)
- -nearly every iteration of the name



ジメチルジチオデナフィル (Let's be Chemists!)



Dithiodimethylcarbodenafil

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Food Additives & Contaminants: Part A> Volume 34, 2017 - Issue 7

Enter keywords, authors, DOI etc.

2 results (0.84 seconds)

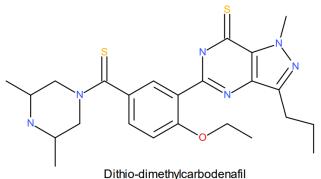
Did you mean: Dithio Dimethyl Carbodenafil

Phosphodiesterase-5 inhibitors and their analogues as adulterants of ... www.tandfonline.com/doi/full/10.1080/19440049.2017.1336674?src=recsys

Jun 29, 2017 ... dioxohongdenafil (14), dithiodesmethylcarbodenafil (15), dithiodimethylcarbodenafil (16), hydroxychlorodenafil (20), hydroxyhomosildenafil ...

Phosphodiesterase-5 inhibitors and their analogues as adulterants of ... www.tandfonline.com/doi/pdf/10.1080/19440049.2017.1336674?needAccess=true

May 27, 2017 - 15. Dithiodesmethylcarbodenafil. C23H30N6OS2. 9.933. 471.1996. 343.07, 371.10, 414.14. 16. Dithiodimethylcarbodenafil. C24H32N6OS2.



Original Articles 53

CrossRef citations

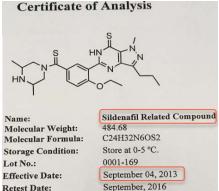
Altmetric

Phosphodiesterase-5 inhibitors and their analogues as adulterants of herbal and food products: analysis of the Malaysian market, 2014–16

Nur Baizura Bujang, Chin Fei Chee 📵, Choon Han Heh 📵, Noorsaadah Abd. Rahman 📵 & Michael I. C. Buckle 2 (1) Pages 1101-1109 | Received 07 Feb 2017, Accepted 27 May 2017, Accepted author version posted online: 05 Jun 2017,

Retest Date:





43

Summary



- Compound ジメチルジチオデナフィル, known to the Japanese and Korean chemists for about two years, and referred to as dimethyldithiodenafil, has not been seen in the English-language research and enforcement sources. It was not registered with CAS, and was not found in any of the surveyed online databases.
- Same compound, with no assigned chemical identifiers, has been made commercially available by CacheSyn, Inc. the research chemicals supplier for at least four years (since 2013), as 'Sildenafil Related Compound', without a name, CAS or any other registration.
- Researchers in Malaysia used the commercial standard from CacheSyn that they termed systematically correctly dithiodimethylcarbodenafil
- To a typical layperson surveying literature, or even a relatively well-informed laboratory scientist, it is difficult to impossible to connect the following without extensive digging:

ジメチルジチオデナフィル = dimethyldithiodenafil = dithiodimethylcarbodenafil = 'Sildenafil Related Compound'

Conclusion



- Correct chemical nomenclature is the cornerstone of successful chemical analysis, indispensable in conducting meaningful analytical work and disseminating its outcomes within the scientific community;
- USP Database scientists invest significant effort in maintaining the chemical nomenclature accurate. USP is collaborating with the US FDA Substance Registration System chemists to attain maximum accuracy in chemical records collected in the USP database, as well as in other online resources such as FDA SRS, PubChem, ChemIDplus, GInAS (Global Ingredient Archival System) and Wikipedia;
- ▶ Upon verification of the chemical information for the newly discovered chemical entities, SRS scientists issue a unique ingredient identifier (UNII) a non-proprietary, free, unique, unambiguous, non-semantic, alphanumeric identifier based on a substance's molecular structure and/or descriptive information. UNII codes help remove nomenclatural ambiguity frequently associated with CAS Registry Numbers;
- USP strives to make the Adulterants Database resource useful to practicing chemists and envisions its role as a convenient platform for exchanging essential analytical data about adulterating compounds, both established and new.

Thank You

