

DOPE: DEVASTATION OF POTENTIATION EFFICACY OF HOMEOSTASIS

¹*Dr. Kishor Dholwani, ²Kushal Nandi, ²Amrita Chakraborty, ²Dr. Dhruvo Jyoti Sen and ³Dr. Dhananjay Saha

¹Laxminarayandev College of Pharmacy, Narmada Nagar, Beside Swaminarayan School, Bholav, Bharuch, Gujarat, India.

²Department of Pharmaceutical Chemistry, School of Pharmacy, Techno India University, Salt Lake City, Sector-V, EM-4, Kolkata-700091, West Bengal, India.

³Deputy Director, Directorate of Technical Education, Bikash Bhavan, Salt Lake City, Kolkata-700091, West Bengal, India.

*Corresponding Author: Dr. Kishor Dholwani

Laxminarayandev College of Pharmacy, Narmada Nagar, Beside Swaminarayan School, Bholav, Bharuch, Gujarat, India.

Article Received on 04/08/2021

Article Revised on 25/08/2021

Article Accepted on 15/09/2021

ABSTRACT

DOPE: Drugs Oppress People Every-day is performance enhancing substances, also known as performance enhancing drugs (PED), are substances that are used to improve any form of activity performance in humans. All doping drugs are four quadrant molecules having four rings structure moieties. This is called as cyclopentanoperhydrophenanthrene ring which has four rings fused in nature [A, B, C, D] rings; which is steroid. All are anabolic in nature. A well-known example involves doping in sport, where banned physical performance enhancing drugs are used by athletes and bodybuilders. Athletic performance enhancing substances are sometimes referred to as **Dope Drugs**.

KEYWORDS: Steroids, Anabolic agents, Dope, PED, WADA.

INTRODUCTION

The father of anabolic steroids in the United States was John Ziegler (1917–1983), a physician for the U.S. weightlifting team in the mid-20th century.



Figure 1: Father of doping John Ziegler.

Anabolic Androgenic Steroids (AAS): These are the substances that have both anabolic and androgenic properties. 'Anabolic' means 'tissue building' and 'androgenic' means 'masculinizing'. The anabolic properties may affect accelerated growth of muscles and bones while the androgenic properties may affect

development of male reproductive system and secondary male sexual characteristics such as hairiness and deep voice. The anabolic androgenic steroids can be derived both endogenously (natural) as well as exogenously (synthetic).^[1]

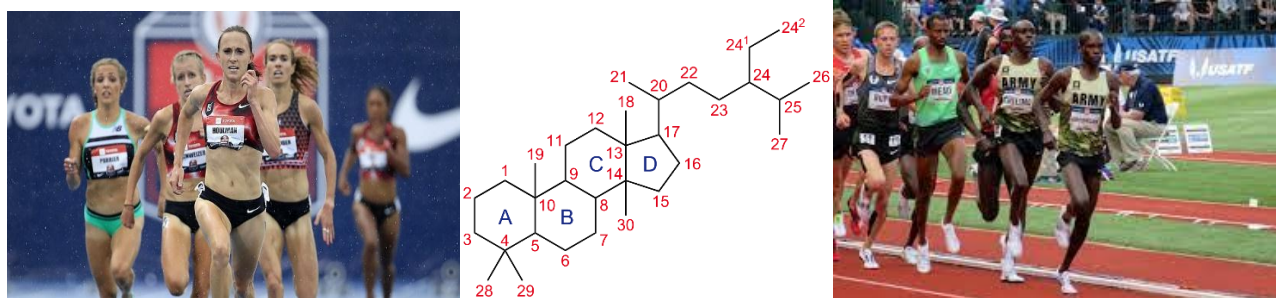


Figure 2: Doping in race.

After administration of anabolic androgenic steroids, the formation of protein is promoted in genital organ, skin, skeleton and muscles. Athletes may be tempted to use anabolic androgenic steroids to improve their physical and physiological capacity to train and compete at highest level by reducing associated fatigues and recovery duration. In an impression to increase muscular power and strength these substances are sometimes taken

by athletes involved in weightlifting, throwing and other sports involving strength parameters.^[2]

Side Effects of Anabolic Androgenic Steroids

The side effects associated with anabolic androgenic steroids are extremely serious and are divided into general, male specific and female specific.

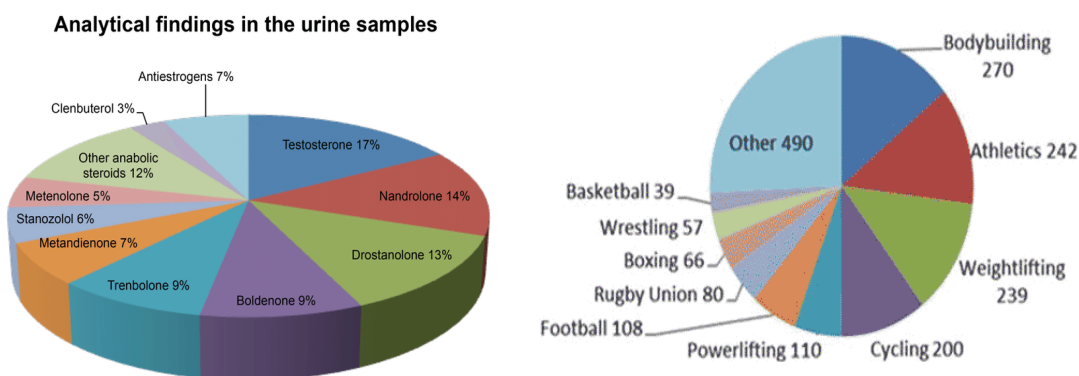


Figure 3: Dope test.

General Side Effects

- Greasy skin and acne
- Infertility
- Hypertension
- Liver and kidney dysfunction
- Aggressive behaviour
- Tumour

- Diminished male hormone production
- Diminished sperm production
- Impotence
- Alopecia
- Prostate cancer

Male specific Effects

- Breast development [gynecomastia]
- Testicular atrophy

Female specific Effects

- Male pattern hair growth and baldness
- Menstruation disturbances
- Decreased size of breast
- Deeper voice (hoarseness)



Figure 4: WADA & Dope test lab.

Other anabolic agents: Other anabolic agents are substances which pharmacologically are not related to anabolic androgenic steroids, but may have the similar anabolic effect. This class of substances has been added in the WADA [World Anti-Doping Agency] list of prohibited substances and methods because of clenbuterol and zeranol abuse in sports.^[3]

Side Effects of other Anabolic Agents

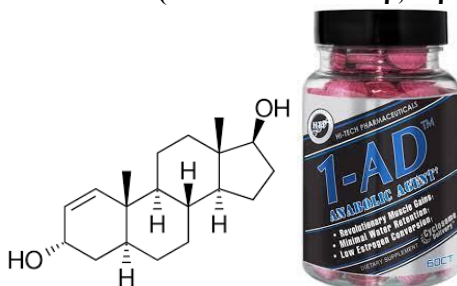
- Trembling
- Restlessness, aggressive behavior
- Anxiety
- Arrhythmias
- Muscle cramps



Figure 5: Dope steroids to athletes.

Anabolic androgenic steroids (AAS)

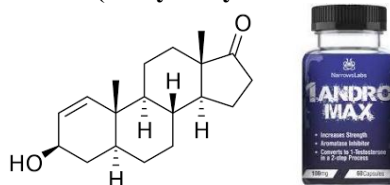
1-Androstenediol (5 α -androst-1-ene-3 β , 17 β -diol)



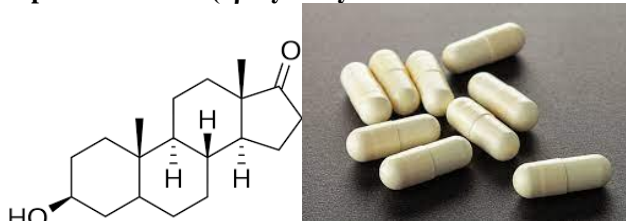
1-Androstenedione (5 α -androst-1-ene-3, 17-dione)



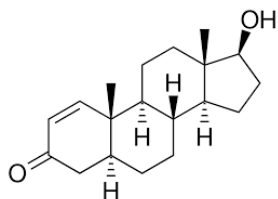
1-Androsterone (3 α -hydroxy-5 α -androst-1-ene-17-one)



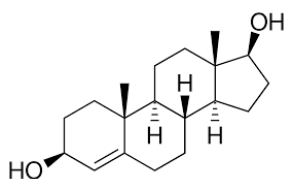
1-Epiandrosterone (3 β -hydroxy-5 α -androst-1-ene-17-one)



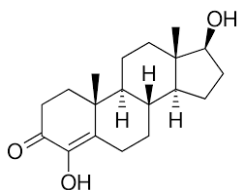
1-Testosterone (17β-hydroxy-5α-androst-1-en-3-one)



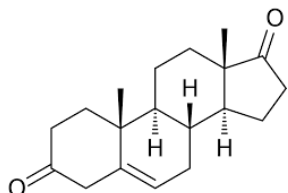
4-Androstenediol (androst-4-ene-3β,17β- diol)



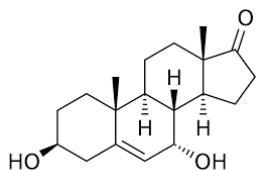
4-Hydroxytestosterone (4,17β-dihydroxyandrost-4-en-3-one)



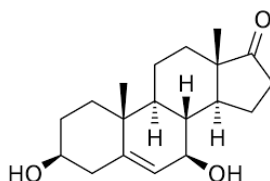
5-Androstenedione (androst-5-ene-3,17- dione)



7α-hydroxy-DHEA (3β,7α-Dihydroxyandrost-5-ene-17-one)



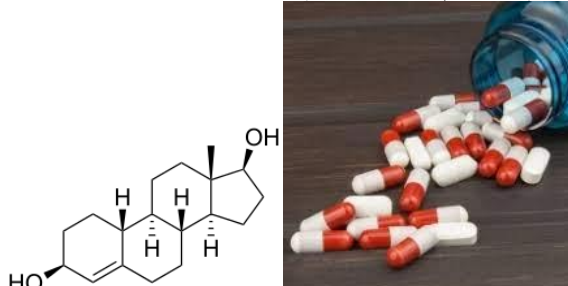
7β-hydroxy-DHEA (3β,7β-dihydroxyandrost-5-ene-17-one)



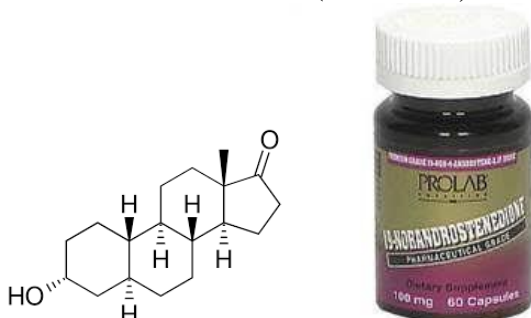
7-Keto-DHEA ((3 β)-3-Hydroxyandrost-5-ene-7,17-dione)



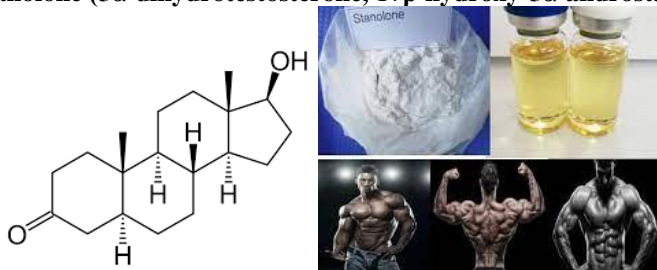
19-Norandrostenediol (estr-4-ene-3,17-diol)



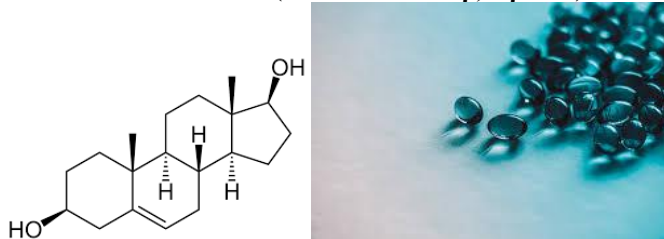
19-Norandrostenedione (estr-4-ene-3,17- dione)



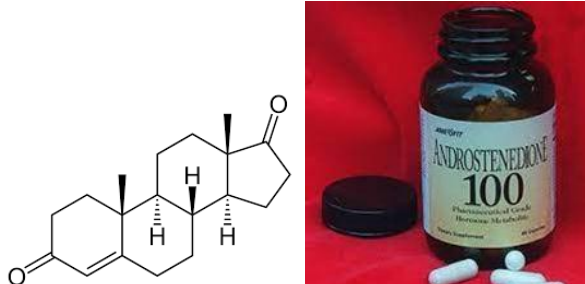
Androstanolone (5 α -dihydrotestosterone, 17 β -hydroxy-5 α -androstan-3-one)



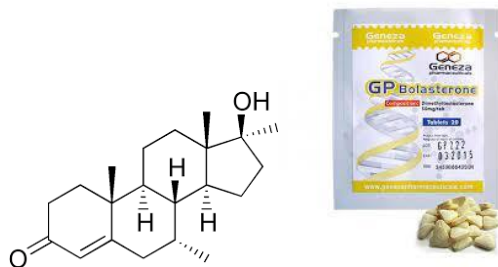
Androstenediol (androst-5-ene-3 β ,17 β -diol)



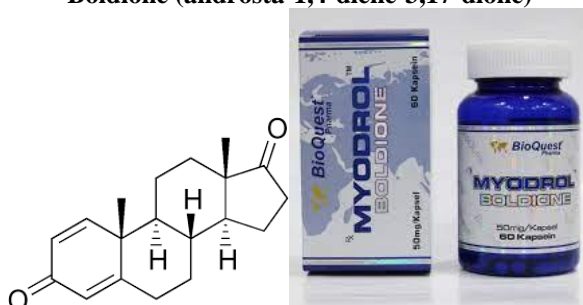
Androstenedione (androst-4-ene-3,17- dione)



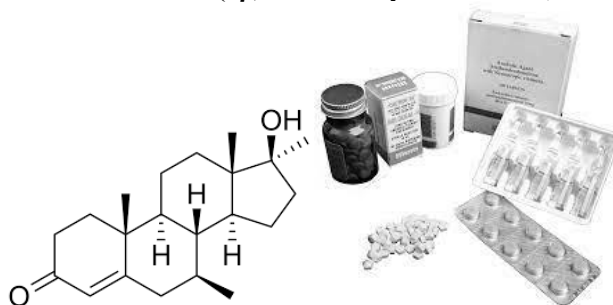
Bolasterone (7 α ,17 α -dimethyltestosterone)



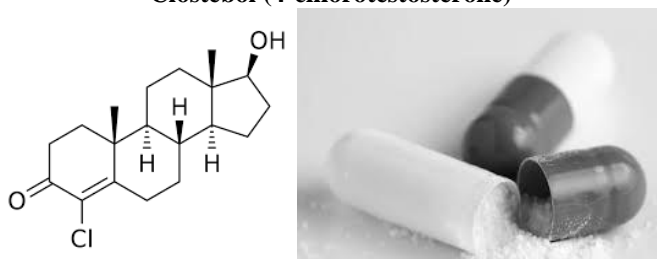
Boldione (androsta-1,4-diene-3,17-dione)



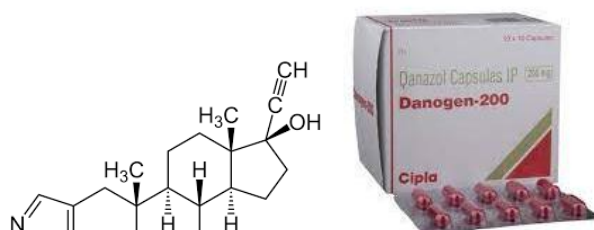
Calusterone (7 β ,17 α -dimethyltestosterone)



Clostebol (4-chlorotestosterone)



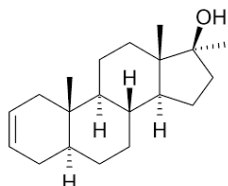
Danazol ([1,2]oxazolo[4',5':2,3]pregna-4-en-20-yn-17 α -ol)



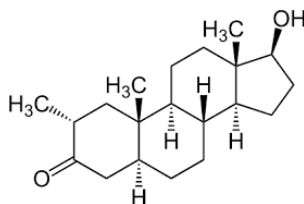
Dehydrochloromethyltestosterone (4-chloro-17 β -hydroxy-17 α -methylandrosta-1,4-dien-3-one)



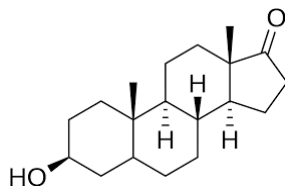
Desoxymethyltestosterone (17 α -methyl-5 α - androst-2-en-17 β -ol and 17 α -methyl-5 α - androst-3-en-17 β -ol)



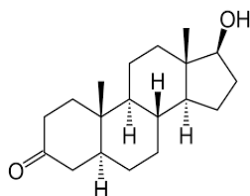
Drostanolone (2 α -Methyl-4,5 α -dihydrotestosterone)



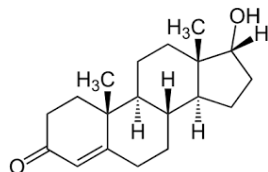
Epiandrosterone (3 β -hydroxy-5 α -androst-17-one)



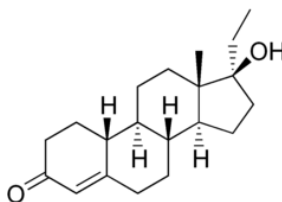
Epi-dihydrotestosterone (17 β -hydroxy-5 β - androst-3-one)

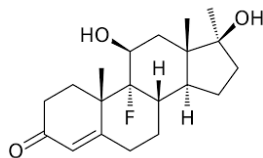
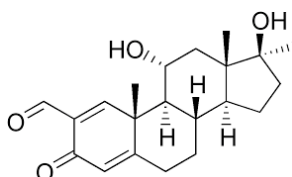
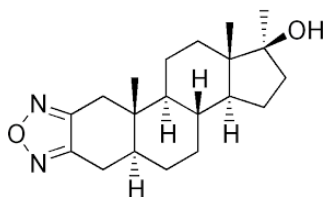
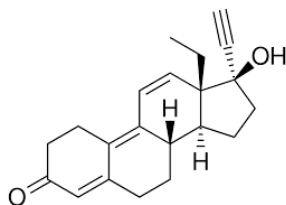
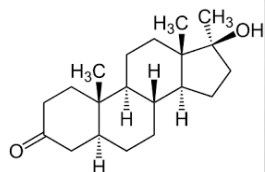
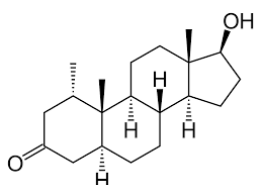


Epitestosterone (androst-4-en-17 α -ol-3-one)

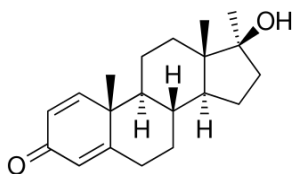


Ethylestrenol (19-norpregna-4-en-17 α -ol)

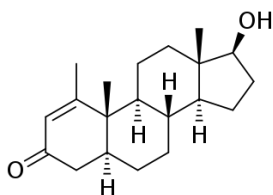


Fluoxymesterone (9 α -Fluoro-11 β -hydroxy-17 α -methyltestosterone)**Formebolone (2-formyl-11 α -hydroxy-17 α -methyl- δ 1-testosterone)****Furazabol (17 α -methyl [1,2,5] oxadiazolo[3',4':2,3]-5 α -androstan-17 β -ol)****Gestrinone (17 α -Ethynyl-18-methyl- δ ^{9,11}-19-nortestosterone)****Mestanolone (17 α -Methyl-4,5 α -dihydrotestosterone)****Mesterolone (1 α -Methyl-5 α -androstan-17 β -ol-3-one)**

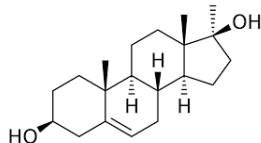
Metandienone (17 β -hydroxy-17 α - methylandrosta-1,4-dien-3-one)



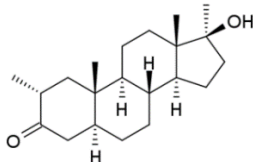
Metenolone (1-Methyl- δ^1 -4,5 α -dihydrotestosterone)



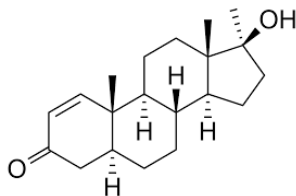
Methandriol (17 α -methylandrosta-5-ene-3 β ,17 β -diol)



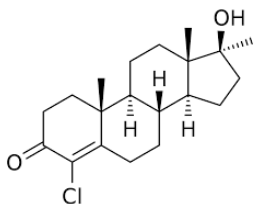
Methasterone (17 β -hydroxy-2 α ,17 α - dimethyl-5 α -androst-3-one)



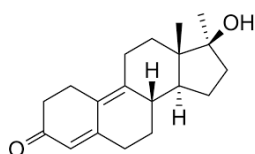
Methyl-1-testosterone (17 β -hydroxy-17 α - methyl-5 α -androst-1-en-3-one)



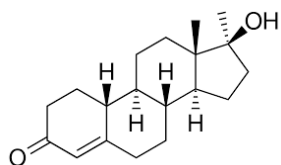
Methylclostebol (4-chloro-17 α -methyltestosterone)



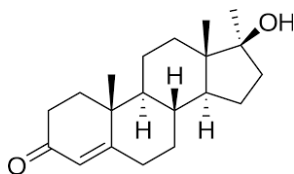
Methyldienolone (17β-hydroxy-17α-methylestra-4,9-dien-3-one)



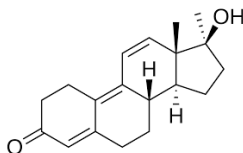
Methylnortestosterone (17β-hydroxy-17α-methylestr-4-en-3-one)



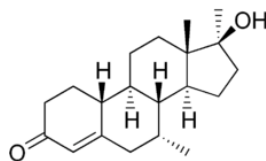
Methyltestosterone (17α-Methyltestosterone)



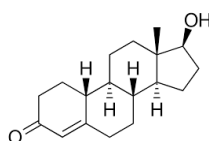
Metribolone (methyltrienolone, 17β-hydroxy-17α-methylestra-4,9,11-trien-3-one)



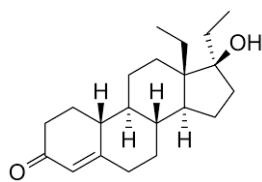
Mibolerone (7α,17α-dimethyl-19-nortestosterone)



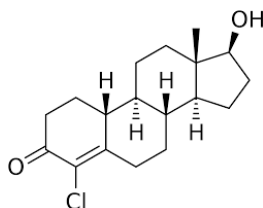
Nandrolone (19-nortestosterone)



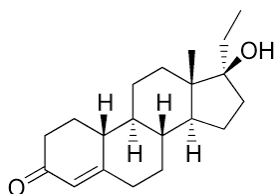
Norboleone (17 α -Ethyl-18-methyl-19-nortestosterone)



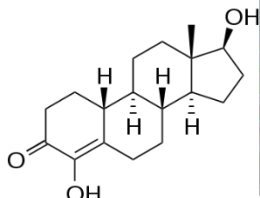
Norclostebol (4-chloro-17 β -ol-estr-4-en-3-one)



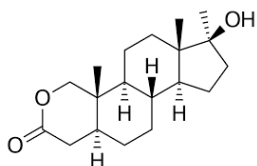
Norethandrolone (17 α -Ethyl-19-nortestosterone)



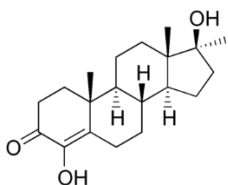
Oxabolone (4-Hydroxy-19-nortestosterone)



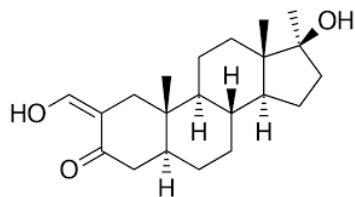
Oxandrolone (17 α -Methyl-2-oxa-5 α -androstan-17 β -ol-3-one)



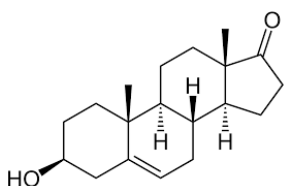
Oxymesterone (4-Hydroxy-17 α -methyltestosterone)



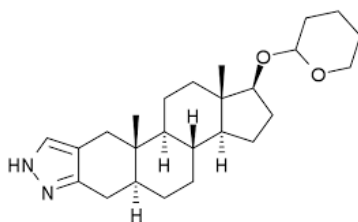
Oxymetholone (2-Hydroxymethylene-17 α -methyl-4,5 α -dihydrotestosterone)



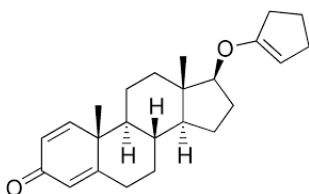
Prasterone (dehydroepiandrosterone, DHEA, 3 β -hydroxyandrost-5-en-17-one)



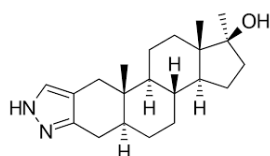
Prostanzol (17 β -[(tetrahydropyran-2-yl) oxy]-1'H-pyrazolo[3,4:2,3]-5 α -androstane)



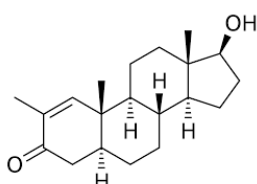
Quinbolone (1-Dehydrotestosterone 17 β -cyclopent-1-enyl ether)

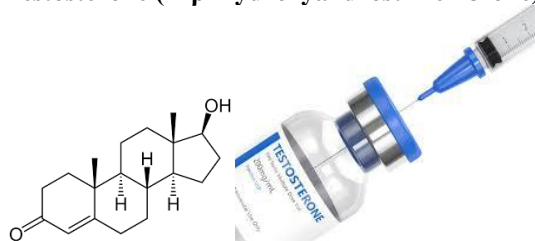
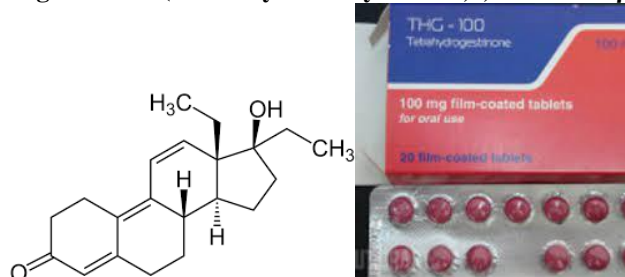
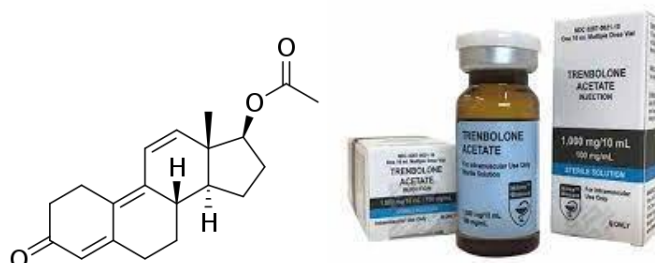


Stanozolol (17 α -Methyl-2'H-5 α -androst-2-eno[3,2-c]pyrazol-17 β -ol)



Stenbolone (2-Methyl-5 α -androst-1-en-17 β -ol-3-one)



Testosterone (17 β -Hydroxyandrost-4-en-3-one)**Tetrahydrogestrinone (17 α -Ethyl-18-ethylestra-4,9,11-trien-17 β -ol-3-one)****Trenbolone acetate (17 β -hydroxyestr-4,9,11-trien-3-one acetate)**

and other substances with a similar chemical structure or similar biological effect(s).

Other anabolic agents

Including, but not limited to

Clenbuterol, selective androgen receptor modulators [SARMs, e.g. andarine, LGD-4033 (ligandrol), enobosarm (ostarine) and RAD140], tibolone, zeranol and zilpaterol.^[4-10]

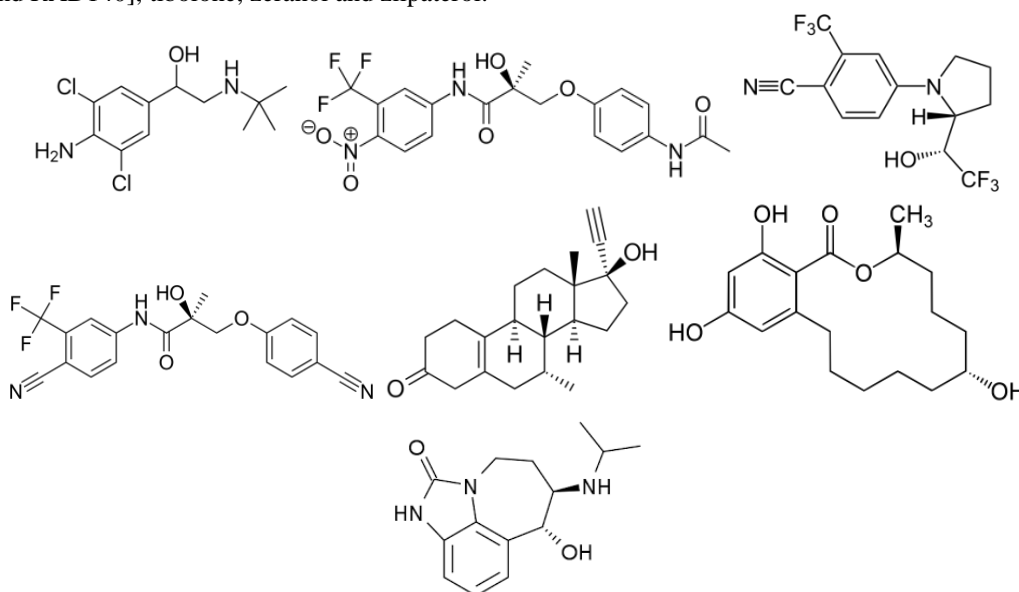


Figure 6: SARMs.

The following substances, and other substances with similar chemical structure or similar biological effect(s), are prohibited.

Erythropoietin [EPO] and agents affecting erythropoiesis

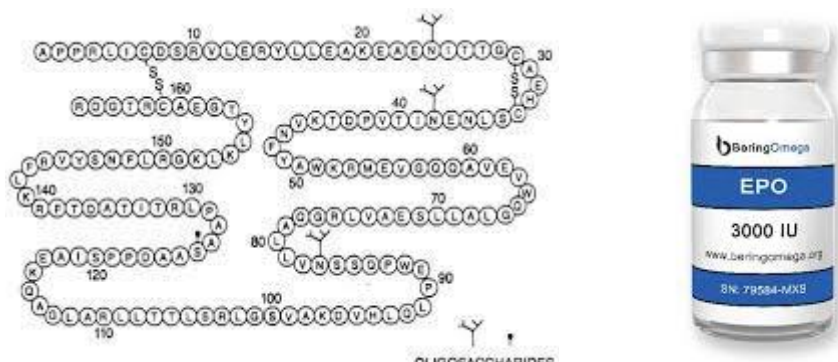


Figure 7: EPO.

Including, but not limited to

Erythropoietin receptor agonists, e.g. darbepoietins (dEPO); erythropoietins (EPO)

EPO-based constructs [e.g. EPO-Fc, methoxy polyethylene glycol-epoetin beta (CERA)]; EPO-mimetic agents and their constructs (e.g. CNTO-530, peginesatide).

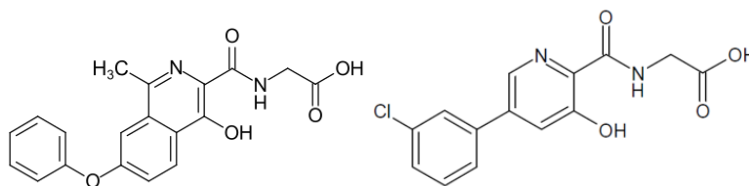
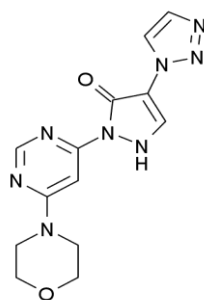
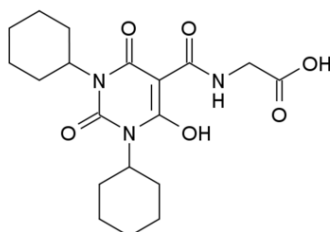
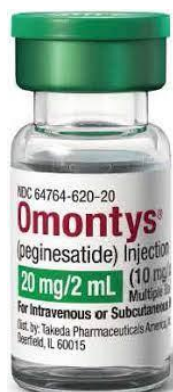
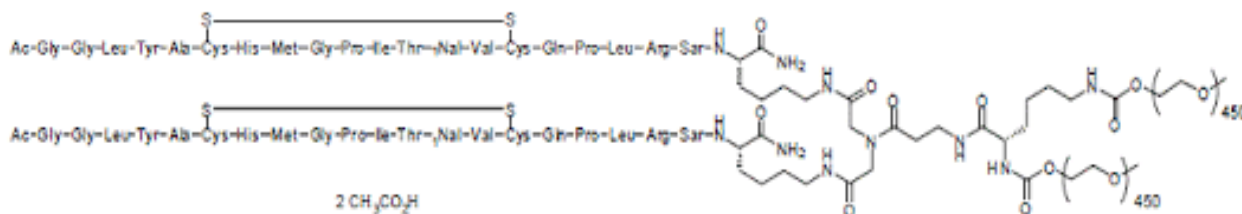


Figure 8: EPO agonists.

Hypoxia-inducible factor (HIF) activating agents, e.g. cobalt; daprodustat (GSK1278863); IOX2; molidustat (BAY 85-3934); roxadustat (FG-4592); vadadustat (AKB-6548); xenon.

GATA inhibitors, e.g. K-11706.

Transforming growth factor beta (TGF- β) signalling inhibitors, e.g. luspatercept; sotatercept.

Innate repair receptor agonists, e.g. asialo EPO; carbamylated EPO (CEPO).

Peptide hormone and their releasing factors: Chorionic gonadotrophin (CG) and luteinizing hormone (LH) and their releasing factors in males, e.g. buserelin, deslorelin, gonadorelin, goserelin, leuporelin, nafarelin and triptorelin.

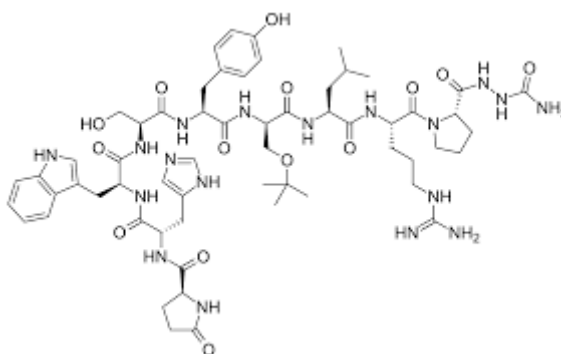
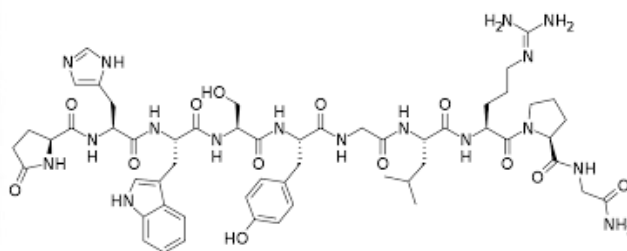
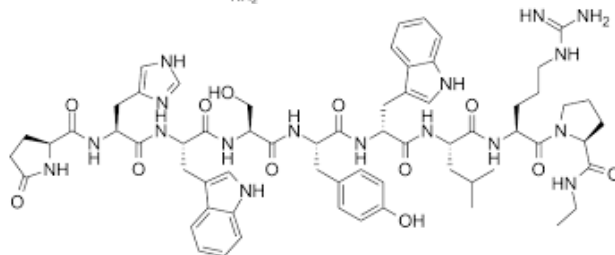
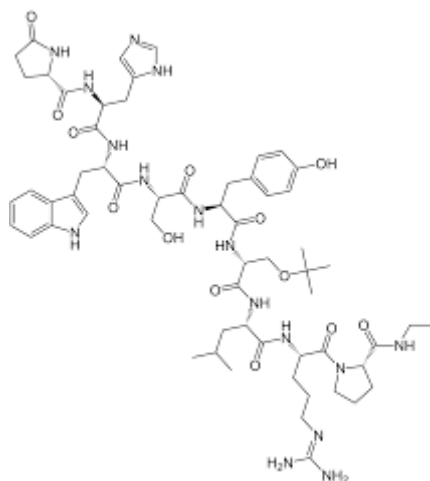




Figure 9: Peptide hormones and releasing factors.

Corticotrophins and their releasing factors, e.g. corticorelin.

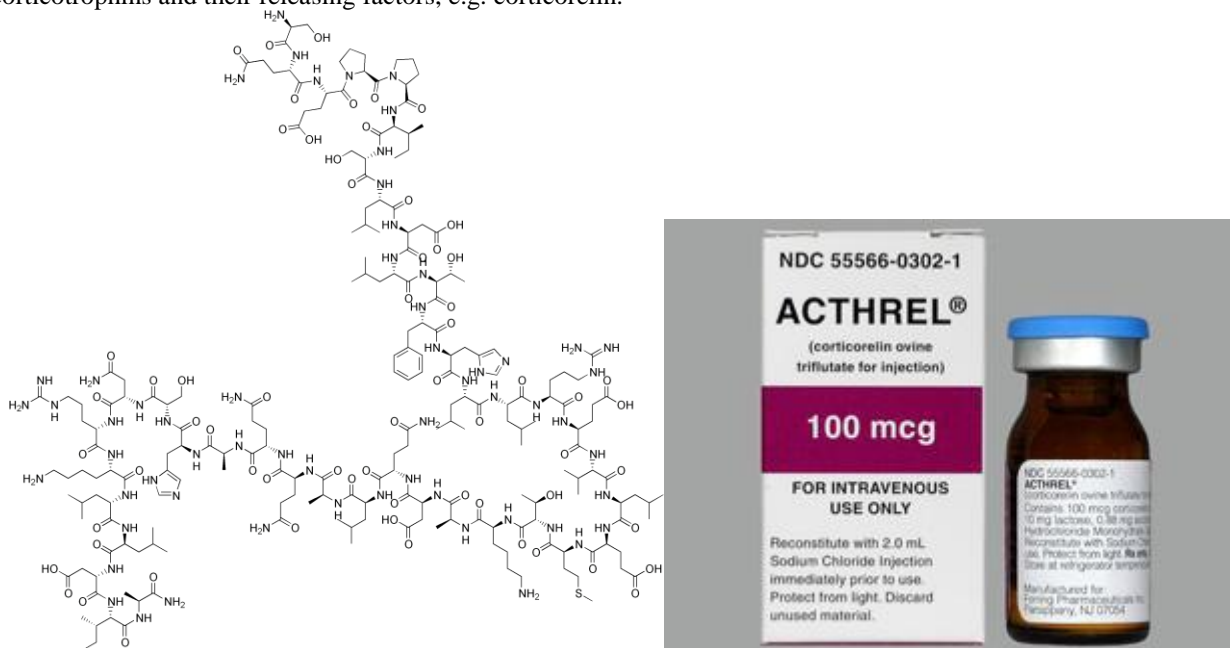
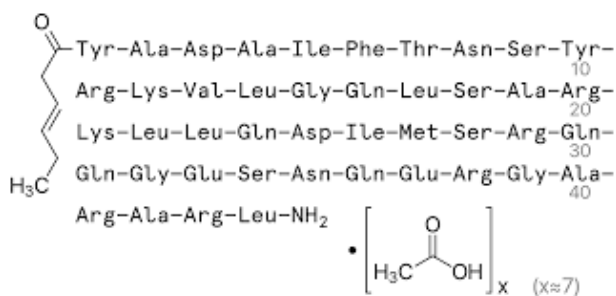
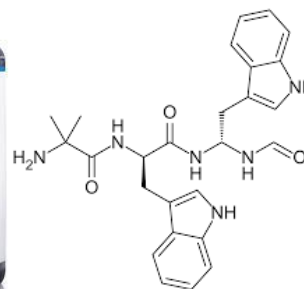
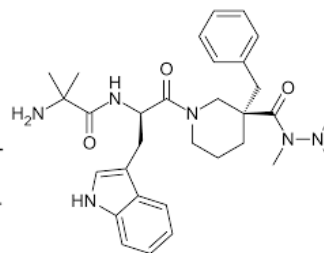


Figure 10: Corticotrophins and releasing factor.

Growth hormone (GH), its fragments and releasing factors, including, but not limited to: growth hormone fragments, e.g. AOD-9604 and hGH 176-191; growth hormone-releasing hormone (GHRH) and its analogues, e.g. CJC-1293, CJC-1295, sermorelin and tesamorelin; growth hormone secretagogues (GHS), e.g. lenomorelin

(ghrelin) and its mimetics, e.g. anamorelin, ipamorelin, macimorelin and tabimorelin; GH-releasing peptides (GHRPs), e.g. alexamorelin, GHRP-1, GHRP-2 (pralmorelin), GHRP-3, GHRP-4, GHRP-5, GHRP-6, and examorelin (hexarelin).^[11-14]





Rx only NDC 71090-002-02

Macrilen[™]
(macimorelin) for oral solution

60 mg

Must administer dose within 30 minutes after reconstitution. Discard unused portion.

For oral use only.



193277/1

Each pouch contains:

60 mg macimorelin (equivalent to 68 mg macimorelin acetate), which corresponds to 60 mg/120 mL (0.5 mg/mL), when reconstituted.

Directions for use: See enclosed package insert.

Must be refrigerated, store at 2-8 °C (36-46 °F).

Keep out of reach of children.

Manufactured by: Alplharma Pharbil Arzneimittel GmbH, Goettingen, Germany

Distributed by: Strongbridge U.S. Inc., Trevose, PA 19053

193278/1

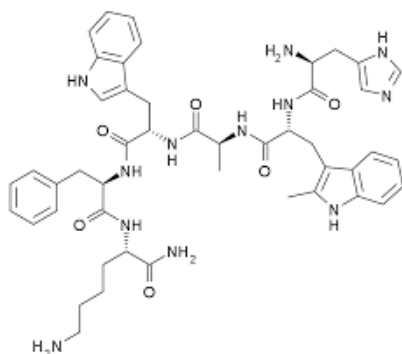
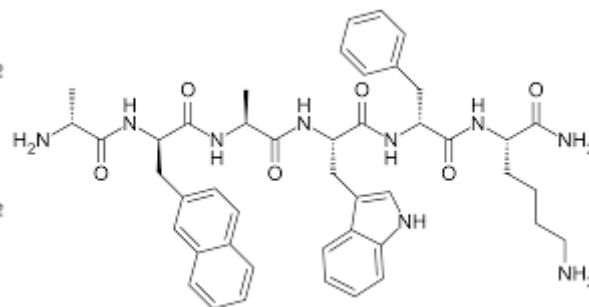
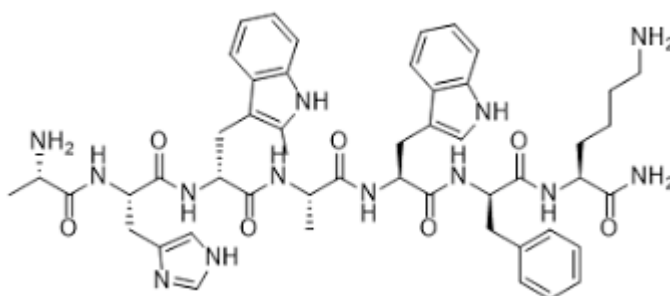
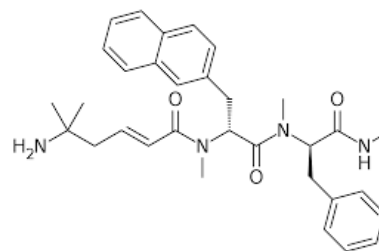


Figure 11: Growth hormone and releasing factor.

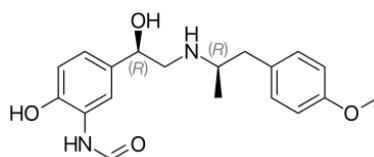
Growth factors and modulators**Including, but not limited to**

Fibroblast growth factors (FGFs), Hepatocyte growth factor (HGF), Insulin-like growth factor 1 (IGF-1) and its analogues, Mechano growth factors (MGFs), Platelet-derived growth factor (PDGF), Thymosin- β 4 and its derivatives e.g. TB-500, Vascular endothelial growth factor (VEGF) and other growth factors or growth factor

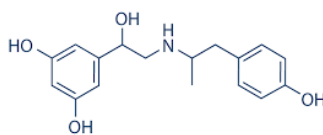
modulators affecting muscle, tendon or ligament protein synthesis/degradation, vascularisation, energy utilization, regenerative capacity or fibre type switching. All selective and non-selective beta-2 agonists, including all optical isomers, are prohibited.^[15]

Including, but not limited to

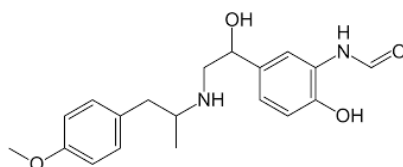
Arformoterol (N-[2-hydroxy-5-[(1R)-1-hydroxy-2-[[2-(4-methoxyphenyl)propan-2-yl]amino]ethyl] phenyl] formamide)



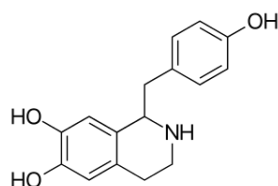
Fenoterol ((RR,SS)-5-(1-hydroxy-2-[[2-(4-hydroxyphenyl)-1-methylethyl]amino]ethyl)benzene-1,3-diol)



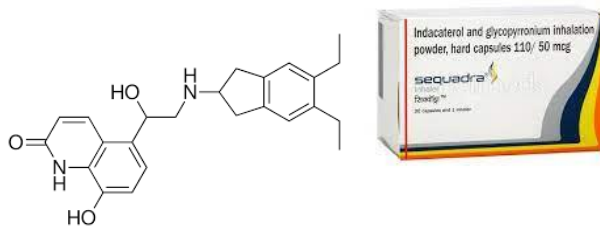
Formoterol ((RR,SS)-N-[2-hydroxy-5-[1-hydroxy-2-[1-(4-methoxyphenyl) propan-2-ylamino]ethyl] phenyl]formamide)



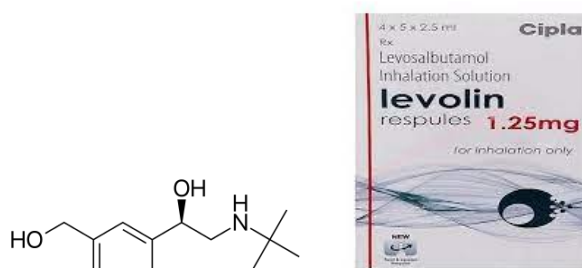
Higenamine (1-[(4-Hydroxyphenyl)methyl]-1,2,3,4-tetrahydroisoquinoline-6,7-diol)



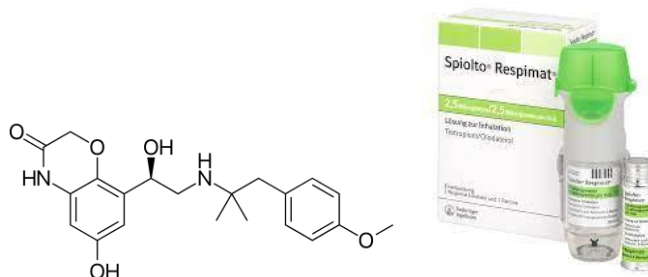
Indacaterol (5-[2-[(5,6-Diethyl-2,3-dihydro-1H-inden-2-yl)amino]-1-hydroxyethyl]-8-hydroxyquinolin-2(1H)-one)



Levosalsbutamol (4-[(1R)-2-(tert-butylamino)-1-hydroxyethyl]-2-(hydroxymethyl)phenol)



Olodaterol (6-hydroxy-8-[(1R)-1-hydroxy-2-[[1-(4-methoxyphenyl)-2-methylpropan-2-yl]amino]ethyl]-4H-1,4-benzoxazin-3-one)



Procaterol ((±)-(1R,2S)-rel-8-Hydroxy-5-[1-hydroxy-2-(isopropylamino)butyl]-quinolin-2(1H)-one)

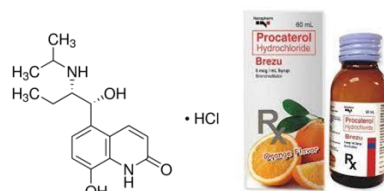
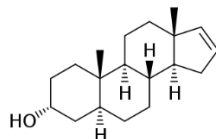


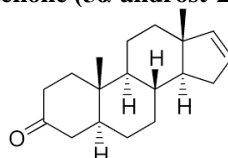
Figure 12: Growth factors modulators.

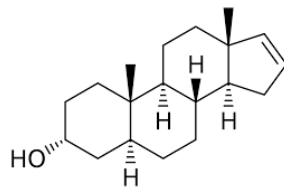
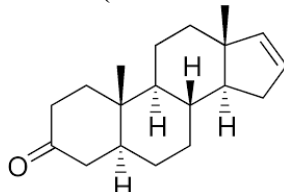
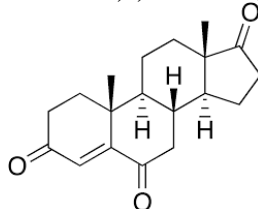
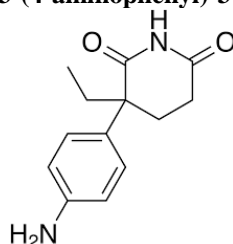
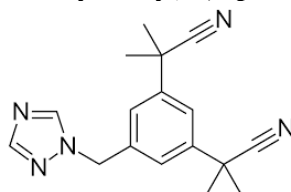
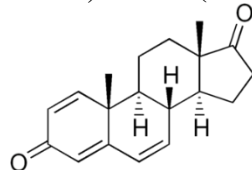
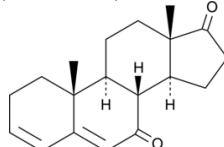
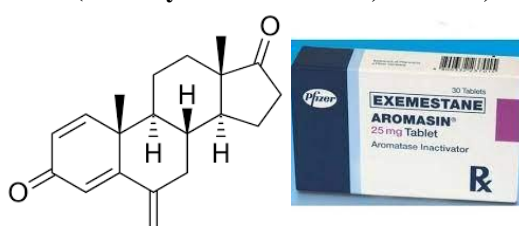
Aromatase inhibitors

2-Androst-enol (5 α -androst-2-en-17-ol)

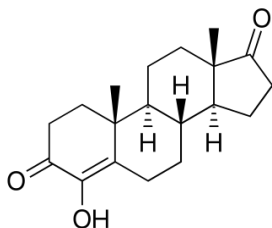


2-Androst-enone (5 α -androst-2-en-17-one)

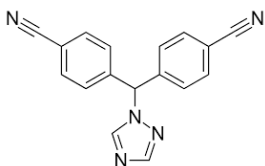


3-Androstenol (5 α -androst-3-en-17-ol)**3-Androstenone (5 α -androst-3-en-17-one)****4-Androstene-3,6,17 trione (6-oxo)****Aminogluthethimide ((RS)-3-(4-aminophenyl)-3-ethyl-piperidine-2,6-dione)****Anastrozole (2,2'-[5-(1H-1,2,4-triazol-1-ylmethyl)-1,3-phenylene]bis(2-methylpropanenitrile)****Androsta-1,4,6-triene-3,17-dione (androstatrienedione)****Androsta-3,5-diene-7,17-dione (arimistane)****Exemestane (6-Methylideneandrosta-1,4-diene-3,17-dione)**

Formestane (4-Hydroxyandrost-4-ene-3,17-dione)



Letrozole (4,4'-((1H-1,2,4-triazol-1-yl)methylene)dibenzonitrile)



Testolactone (13-Hydroxy-3-oxo-13,17-secoandrosta-1,4-dien-17-oic acid δ-lactone)

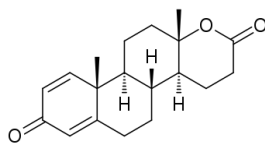
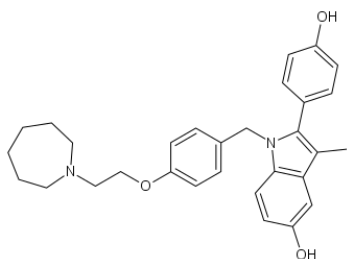


Figure 13: Aromatase inhibitors.

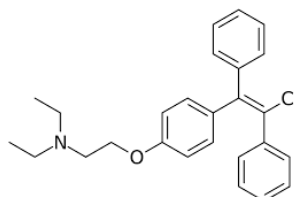
Anti-estrogenic substances [anti-estrogenic and selective estrogenic receptor modulators SERMS

Including, but not limited to

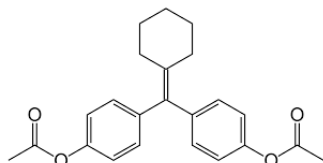
Bazedoxifene (1-[4-[2-(azepan-1-yl)ethoxy]benzyl]-2-(4-hydroxyphenyl)-3-methyl-1H-indol-5-yl)



Clomifene ((E,Z)-2-(4-(2-chloro-1,2-diphenylethenyl)phenoxy)-N,N-diethylethanamine)



Cyclofenil ([4-[(4-Acetoxyphenyl)-cyclohexylidene-methyl]phenyl] acetate)



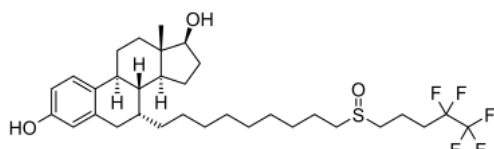
Cyclofenil

Must Read

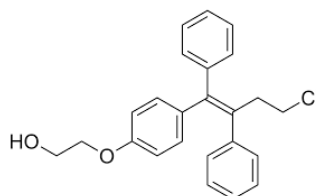


Whatsteroids.com

Fulvestrant (Pentafluoropentyl-sulfinyl]nonyl]estra-1,3,5(10)-triene-3,17β-diol)



Ospemifene (2-(p-((Z)-4-chloro-1,2-diphenyl-1-butenyl)phenoxy)ethanol)

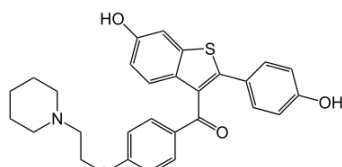


Ospemifene

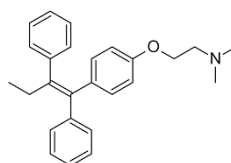
Ospemifene



Raloxifene ([6-hydroxy-2-(4-hydroxyphenyl)-benzothiophen-3-yl]-[4-[2-(1-piperidyl)ethoxy]phenyl]-methanone)



Tamoxifen ((Z)-2-[4-(1,2-Diphenylbut-1-enyl)phenoxy]-N,N-dimethylethanamine)



Toremifene (2-[4-[(1Z)-4-Chloro-1,2-diphenyl-but-1-en-1-yl]phenoxy]-N,N-dimethylethanamine)

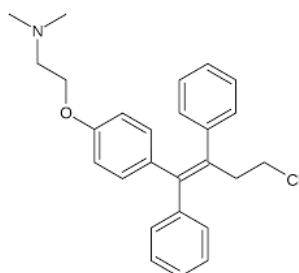
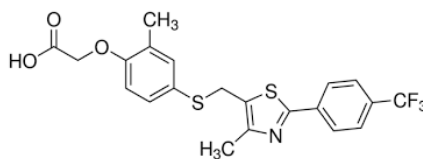


Figure 14: Antiestrogenic substances.

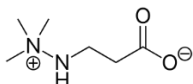
Metabolic modulators: Activators of the AMP-activated protein kinase (AMPK), e.g. AICAR, SR9009; and peroxisome proliferator-activated receptor delta (PPAR δ) agonists, e.g.

2-(2-methyl-4-((4-methyl-2-(4-(trifluoromethyl)phenyl)thiazol-5-yl)methylthio)phenoxy) acetic acid (GW1516, GW501516)



Insulins and insulin-mimetic

Meldonium (2-(2-Carboxylato-ethyl)-1,1,1-trimethylhydrazinium)



Trimetazidine (1-(2,3,4-trimethoxybenzyl)piperazine)

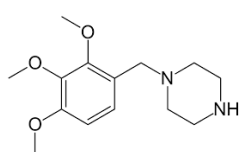
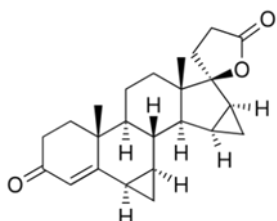


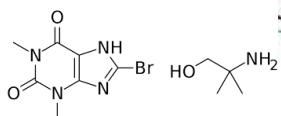
Figure 15: Metabolic modulators.

Exceptions

Drospirenone (17 β -Hydroxy-6 β ,7 β :15 β ,16 β -dimethylene-3-oxo-17 α -pregn-4-ene-21-carboxylic acid)



Pamabrom (1:1 mixture of 2-amino-2-methyl-1-propanol and 8-bromotheophyllinate)



and topical ophthalmic administration of carbonic anhydrase inhibitors (e.g. dorzolamide, brinzolamide); Local administration of felypressin in dental anaesthesia.

The detection in an Athlete's Sample at all times or In-Competition, as applicable, of any quantity of the following substances subject to threshold limits: formoterol, salbutamol, cathine, ephedrine, methylephedrine and pseudoephedrine, in conjunction

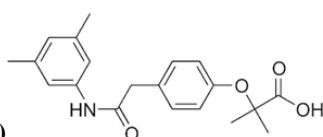
with a diuretic or masking agent, will be considered as an Adverse Analytical Finding (AAF) unless the Athlete has an approved Therapeutic Use Exemption (TUE) for that substance in addition to the one granted for the diuretic or masking agent.

Prohibited methods

Manipulation of blood & blood components: The Administration or reintroduction of any quantity of autologous, allogenic (homologous) or heterologous blood, or red blood cell products of any origin into the circulatory system.

Artificially enhancing the uptake, transport or delivery of oxygen. Including, but not limited to:

Perfluorochemicals; **efaproxiral (RSR13)** and modified haemoglobin products, e.g. haemoglobin-based blood substitutes and microencapsulated haemoglobin products, excluding supplemental oxygen by inhalation. Any form of intravascular manipulation of the blood or blood components by physical or chemical means.^[16-18]



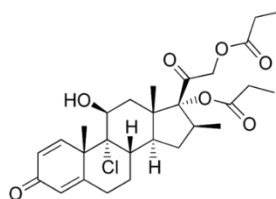
Efaproxiral (RSR13)

Stimulants

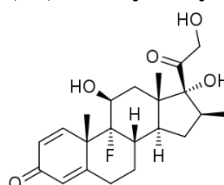
All glucocorticoids are prohibited when administered by oral, intravenous, intramuscular or rectal route.

Including, but not limited to

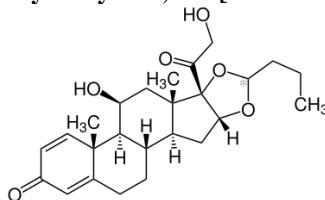
Beclomethasone



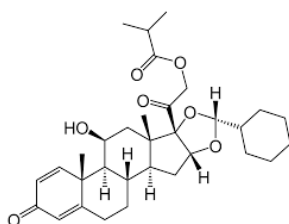
Betamethasone ((11β,16β)-9-Fluoro-11,17,21-trihydroxy-16-methylpregna-1,4-diene-3,20-dione)



Budesonide (1β,21-Dihydroxy-16α,17α-[butane-1,1-diylbis(oxy)]pregna-1,4-diene-3,20-dione)



Ciclesonide ([2-[(1S,2S,4R,6R,8S,9S,11S,12S,13R)-6-cyclohexyl-11-hydroxy-9,13-dimethyl-16-oxo-5,7-dioxapentacyclo[10.8.0.02,9.04,8.013,18]icosa-14,17-dien-8-yl]-2-oxoethyl] 2-methylpropanoate)



2-[4-[2-[(3,5-dimethylphenyl)amino]-2-oxoethyl]phenoxy]-2-methylpropanoic acid

Chemical & physical manipulation: Tampering, or Attempting to Tamper, to alter the integrity and validity of Samples collected during Doping Control.

Including, but not limited to:

Sample substitution and/or adulteration, e.g. addition of proteases to Sample.

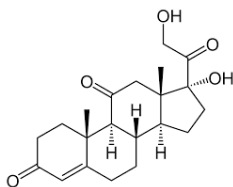
Intravenous infusions and/or injections of more than a total of 100 mL per 12-hour period except for those legitimately received in the course of hospital treatments, surgical procedures or clinical diagnostic investigations.

Gene & cell doping: The following, with the potential to enhance sport performance, are prohibited:

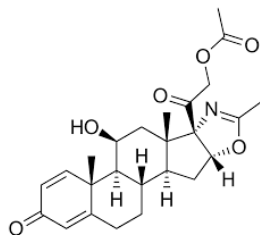
The use of nucleic acids or nucleic acid analogues that may alter genome sequences and/ or alter gene expression by any mechanism. This includes but is not limited to gene editing, gene silencing and gene transfer technologies.

The use of normal or genetically modified cells.

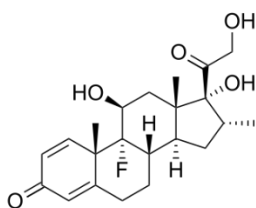
Cortisone (17 α ,21-Dihydroxypregn-4-ene-3,11,20-trione)



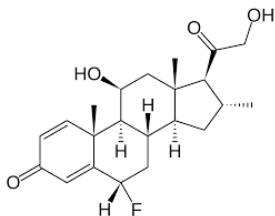
Deflazacort ([2-[(1S,2S,4R,8S,9S,11S,12S,13R)-11-hydroxy-6,9,13-trimethyl-16-oxo-5-oxa-7-azapentacyclo[10.8.0.02,9.04,8.013,18]icosa-6,14,17-trien-8-yl]-2-oxoethyl] acetate)



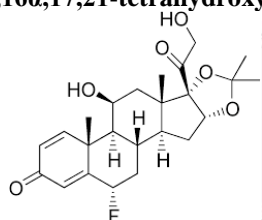
Dexamethasone (8S,9R,10S,11S,13S,14S,16R,17R)-9-Fluoro-11,17-dihydroxy-17-(2-hydroxyacetyl)-10,13,16-trimethyl-6,7,8,9,10,11,12,13,14,15,16,17-dodecahydro-3H-cyclopenta[a]phenanthren-3-one



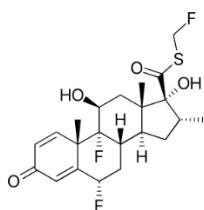
Flucortolone (6S,8S,9R,10S,11S,13S,14S,16R,17S)-6-fluoro-11-hydroxy-17-(2-hydroxyacetyl)-10,13,16-trimethyl-6,7,8,9,11,12,14,15,16,17-decahydrocyclopenta[a]phenanthren-3-one



Flunisolide (6 α -Fluoro-11 β ,16 α ,17,21-tetrahydroxypregna-1,4-diene-3,20-dione acetone cyclic 16,17-acetal)



Fluticasone (6 α ,9 α -Difluoro-11 β ,17 α -dihydroxy-16 α -methyl-21-thia-21-fluoromethylpregna-1,4-dien-3,20-dione



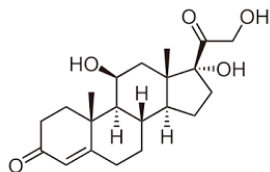
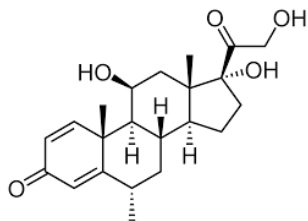
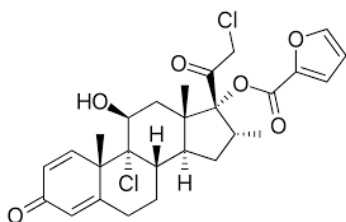
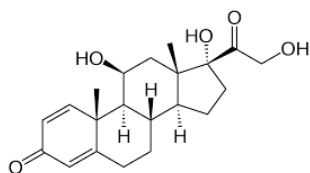
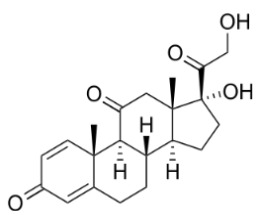
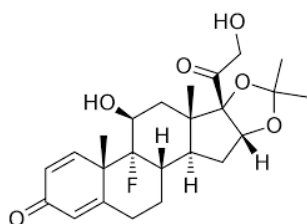
Hydrocortisone (11 β ,17 α ,21-Trihydroxypregna-4-ene-3,20-dione)**Methylprednisolone (11 β ,17,21-Trihydroxy-6 α -methylpregna-1,4-diene-3,20-dione)****Mometasone (9 α ,21-Dichloro-11 β ,17 α -dihydroxy-16 α -methylpregna-1,4-diene-3,20-dione 17 α -(2-furoate))****Prednisolone ((11 β)-11,17,21-Trihydroxypregna-1,4-diene-3,20-dione)****Prednisone (17,21-dihydroxypregna-1,4-diene-3,11,20-trione)****Triamcinolone acetonide (4 α S,4 β R,5S,6 α S,6 β S,9 α R,10 α S,10 β S)-4 β -fluoro-6 β -glycoloyl-5-hydroxy-4 α ,6 α ,8,8-tetramethyl-4 α ,4 β ,5,6,6 α ,6 β ,9 α ,10,10 α ,10 β ,11,12-dodecahydro-2H-naphtho[2',1':4,5]indeno[1,2-d][1,3]dioxol-2-one)**

Figure 16: Glucocorticoids.

Beta-blockers are prohibited In-Competition only, in the following sports, and also prohibited Out-of-Competition where indicated (*).

Archery (WA)*, Automobile (FIA), Billiards (all disciplines) (WCBS), Darts (WDF), Golf (IGF), Shooting (ISSF, IPC)*. *Also prohibited Out-of-Competition: Skiing/Snowboarding (FIS) in ski jumping, freestyle aerials/halfpipe and snowboard halfpipe/big air.^[19,20]

Underwater sports (CMAS) in constant-weight apnoea with or without fins, dynamic apnoea with and without fins, free immersion apnoea, Jump Blue apnoea, spearfishing, static apnoea, target shooting, and variable weight apnoea

Including, but not limited to: Acebutolol, Alprenolol, Atenolol, Betaxolol, Bisoprolol, Bunolol, Carteolol, Carvedilol, Celiprolol, Esmolol, Labetalol, Metipranolol, Metoprolol, Nadolol, Nebivolol, Oxprenolol, Pindolol, Propranolol, Sotalol, Timolol



Figure 17: Dope & Race.

Dope testing: The National Dope Testing Laboratory is equipped with state-of-the-art technologies and the most modern equipment. The use of Gas Chromatography coupled with Mass Spectrometry (GC-MS) is the most common and the oldest technology being used worldwide for dope testing. Nowadays, the use of liquid chromatography coupled to tandem mass spectrometry (LC-MS/MS) has become quite widespread. This technique has helped detect the difficult drugs falling into various categories of banned substances and is becoming increasingly more important in the fight against doping. Apart from GC-MS and LC-MS/MS, the use of Gas Chromatography coupled with tandem Mass Spectrometry (GC-MS/MS) and Isotope-ratio mass spectrometry (IRMS) is also very prevalent in sports dope testing. Both GC-MS/MS and LC-MS/MS are used primarily to analyse urine samples. The analysis of the blood matrix requires a completely different type of equipment which is commonly used in hospital laboratories.^[21]

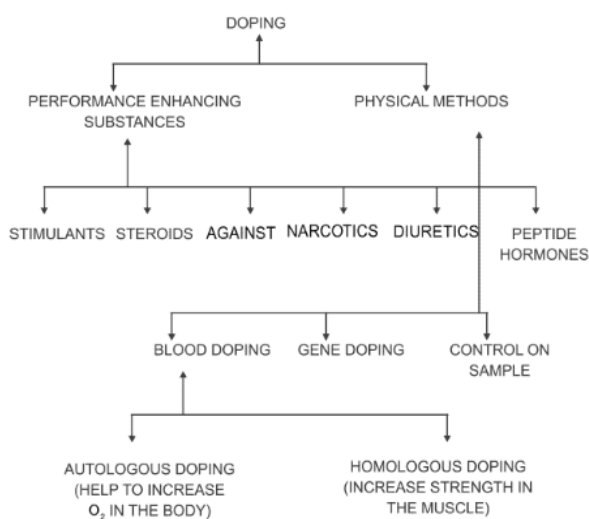


Figure 18: Dope testing.

Analogy of dope drugs with medication therapy: Mostly all dope drugs are cyclopentanoperhydrophenanthrene structure in nature. All are highly lipid soluble in nature and having high potency anabolic in nature. The four quadrants of this steroidal nature of all doping drugs are capable to be

soluble in lipid layer of muscle due to high logP values. The cumulative nature of all these agents shows anabolic reflections towards pharmacokinetic profile because their absorption, distribution, metabolism and excretion parameters are also of high extent due to solubilisation in adipose tissues can show high activation in energy

profile in body homeostasis that can produce extra energy to body which helps to athletes to cross the milestones of success. Steroidal nature of doping agents can be detected by chromatography with molar mass content in urine sample [LC-MS] because after ADME the excretions of metabolites of these agents are also of steroidal units because after metabolism the structural units are also remain same but their substitutions at different parts get into polar units to be excreted easily from body. Cholestane, Coprosten, Pregnane, Androsten, Estrane, etc steroidal units which are present in doping drugs are lipid soluble which prefer adipose tissues to accumulate but after metabolic steps after phase-I & II these are converted into nonpolar entity to polar entity which can be easily excreted from body that can be detected by chromatography.

CONCLUSION

Depending on the sport practiced and the physical attributes it requires, the athletes will look for one or more of the following benefits of doping: recovering from an injury, increasing body recovery capacity after training, increasing muscle mass and strength, decreasing fat tissue, increasing endurance. After critically looking at all aspects of performance-enhancing drug use, we have concluded, as a team, that doping is negatively affecting sports. Performance-enhancing drugs are a bad thing for several reasons. They have terrible side effects on athletes and destroy their bodies in the long run. They give athletes an unfair advantage in sports which is a form of cheating in our opinion. They also promote a do-anything-to-win attitude and an attitude that "ideal" bodies cannot be achieved through hard work and effort. Most importantly, this topic is becoming a social issue because of the prevalence and the effect that it is having on our youth and teenagers. Athletes using drugs are encouraging young people, who view them as role models, to use these drugs to improve their performance and the looks of their bodies. Science has created many drugs and made the population aware of the harm they can do. They have promoted the use in the medical field to speed the healing of injuries. They have in no way recommended the use of these drugs as performance-enhancing drugs in sports. Science has played a huge role in the use of performance-enhancing drugs but they have not made them for this reason. Doping should be banned in all sports leagues, and a no tolerance policy should be enforced.

REFERENCES

- Rashika S Shah, Nirav S Patel, Noopur V Rami, Paresh V Parmar, Chirag M Rathod, Mit H Vaidya and Dhruvo Jyoti Sen; Dope as a dose of physiological enhancers; *International Journal of Comprehensive Pharmacy*, 2016; 07(03): 1-10.
- Ulrich, R.; et al. "Doping in Two Elite Athletics Competitions Assessed by Randomized-Response Surveys" (PDF). *Sports Medicine*, 2017; 48(1): 1-9.
- Connor, James; Woolf, Jules; Mazanov, Jason. "Would they dope? Revisiting the Goldman dilemma" (PDF). *British Journal of Sports Medicine*, 2013; 47(11): 697-700.
- Connor, J. M; Mazanov, J. "Would you dope? A general population test of the Goldman dilemma". *British Journal of Sports Medicine*, 2009; 43(11): 871-872.
- Landy Justin F., Walco Daniel K., Bartels Daniel M. "What's Wrong with using Steroids? Exploring Whether and Why People Oppose the use of Performance Enhancing Drugs". *Journal of Personality and Social Psychology*, 2017; 113(3): 377-392.
- Piacentino Daria, Casale Antonio, Aromatario Maria, Pomara Cristoforo, Girardi Paolo, Sani Gabriele. "Anabolic-androgenic Steroid use and Psychopathology in Athletes. A Systematic Review". *Current Neuropharmacology*, 2015; 13(1): 101-21.
- Pope Harrison G., Wood Ruth I., Rogol Alan, Nyberg Fred, Bowers Larry, Bhasin Shalender. "Adverse Health Consequences of Performance-Enhancing Drugs: An Endocrine Society Scientific Statement". *Endocrine Reviews*, 2014; 35(3): 341-75.
- Woodland, Les. *Dope, the use of drugs in Sport*. UK: David and Charles, 1980. ISBN 978-0-7153-7894-6.
- Deventer, K; Roels, K; Delbeke, FT; Van Eenoo, P. "Prevalence of legal and illegal stimulating agents in sports". *Analytical and Bioanalytical Chemistry*, 2011; 401(2): 421-32. Kanayama G, Hudson JJ, Pope HG. "Long-term psychiatric and medical consequences of anabolic-androgenic steroid abuse: A looming public health concern?". *Drug and Alcohol Dependence*, November, 2008; 98(1-2): 1-12.
- Yesalis CE, Anderson WA, Buckley WE, Wright JE. "Incidence of the nonmedical use of anabolic-androgenic steroids" (PDF). *NIDA Research Monograph*, 1990; 102: 97-112. Walker Jennifer. "Cutaneous Manifestations of Anabolic-Androgenic Steroid Use in Athletes". *International Journal of Dermatology*, 2009; 48(10): 1044-1048.
- Nicole Thualagant. "The conceptualization of fitness doping and its limitations". *Sport in Society*, 2012; 15(3): 409-419.
- Kicman, A T. "Pharmacology of anabolic steroids". *British Journal of Pharmacology*, 2008; 154(3): 502-521.
- Barrett-Connor EL. "Testosterone and risk factors for cardiovascular disease in men". *Diabetes Metab*, 1995; 21(3): 156-61.
- Yamamoto Y, Moore R, Hess HA, Guo GL, Gonzalez FJ, Korach KS, Maronpot RR, Negishi M. "Estrogen receptor alpha mediates 17alpha-ethynylestradiol causing hepatotoxicity". *J Biol Chem.*, 2006; 281(24): 16625-31.

15. De Piccoli B, Giada F, Benettin A, Sartori F, Piccolo E. "Anabolic steroid use in body builders: an echocardiographic study of left ventricle morphology and function". *Int J Sports Med.*, 1991; 12(4): 408–12.
16. Green GA. "Performance-enhancing drug use". *Orthopedics*, 2009; 32(9): 647–649.
17. Turillazzi E, Perilli G, Di Paolo M, Neri M, Riezzo I, Fineschi V. "Side effects of AAS abuse: an overview". *Mini Rev Med Chem.*, 2011; 11(5): 374–89.
18. Hartgens F, Kuipers H. "Effects of androgenic-anabolic steroids in athletes". *Sports Med.*, 2004; 34(8): 513–54.
19. Kicman AT, Gower DB. "Anabolic steroids in sport: biochemical, clinical and analytical perspectives". *Ann. Clin. Biochem*, 2003; 40(Pt 4): 321–56.
20. Basaria S, Wahlstrom JT, Dobs AS. "Clinical review 138: Anabolic-androgenic steroid therapy in the treatment of chronic diseases". *J. Clin. Endocrinol. Metab.*, 2001; 86(11): 5108–17.
21. Ranke MB, Bierich JR. "Treatment of growth hormone deficiency". *Clinics in Endocrinology and Metabolism*, 1986; 15(3): 495–510.