ABRIDGED PRESCRIBING INFORMATION FOR MEDICAL PROFESSIONALS

Tadalex
(Tadalafil U.S.P.)

ٹیڈالیکس
(ٹیڈالافل یو۔ایس۔پی۔)

COMPOSITION
Tadalex 2.5 mg tablet: Each film coated tablet contains Tadalafil ..... 2.5 mg.
Tadalex 5 mg tablet: Each film coated tablet contains Tadalafil ..... 5 mg.
Tadalex 10 mg tablet: Each film coated tablet contains Tadalafil ..... 10 mg.
Tadalex 20 mg tablet: Each film coated tablet contains Tadalafil ..... 20 mg.
Product Specs.: U.S.P.
"This product contains Lactose"

MECHANISM OF ACTION
Penile erection during sexual stimulation is caused by increased penile blood flow resulting from the relaxation of penile arteries and corpus cavernosal smooth muscle. This response is mediated by the release of nitric oxide (NO) from nerve terminals and endothelial cells, which stimulates the synthesis of cGMP in smooth muscle cells. Cyclic GMP causes smooth muscle relaxation and increased blood flow into the corpus cavernosum. The inhibition of phosphodiesterase type 5 (PDE5) enhances erectile function by increasing the amount of cGMP. Tadalafil inhibits PDE5. Because sexual stimulation is required to initiate the local release of nitric oxide, the inhibition of PDE5 by tadalafil has no effect in the absence of sexual stimulation. The effect of PDE5 inhibition on cGMP concentration in the corpus cavernosum and pulmonary arteries is also observed in the smooth muscle of the prostate, the bladder and their vascular supply. The mechanism for reducing BPH symptoms has not been established.

PHARMACOKINETICS
Absorption: After single oral-dose administration, the maximum observed plasma concentration (Cmax) of tadalafil is achieved between 30 minutes and 6 hours (median time of 2 hours). Absolute bioavailability of tadalafil following oral dosing has not been determined. The rate and extent of absorption of tadalafil are not influenced by food; thus tadalafil may be taken with or without food.
Distribution: At therapeutic concentrations, 94% of tadalafil in plasma is bound to proteins.
Metabolism: Tadalafil is predominantly metabolized by CYP3A4 to a catechol metabolite. In vitro data suggests that metabolites are not expected to be pharmacologically active at observed metabolite concentrations.
Excretion: The mean terminal half-life is 17.5 hours in healthy subjects. Tadalafil is excreted predominantly as metabolites, mainly in the feces (approximately 61% of the dose) and to a lesser extent in the urine (approximately 36% of the dose).
Patients with Diabetes Mellitus: No dose adjustment is warranted.
Patients with BPH: No dose adjustment is warranted.

INDICATIONS AND USAGE
Erectile Dysfunction: Tadalex is indicated for the treatment of erectile dysfunction (ED).
Benign Prostatic Hyperplasia: Tadalex is indicated for the treatment of the signs and symptoms of benign prostatic hyperplasia (BPH).
Erectile Dysfunction and Benign Prostatic Hyperplasia: Tadalex is indicated for the treatment of ED and the signs and symptoms of BPH (ED/BPH).

DOSAGE AND ADMINISTRATION
Do not split **Tadalex** tablets; entire dose should be taken.
Tadalex for Use as Needed for Erectile Dysfunction
- The recommended starting dose of tadalafil for use as needed in most patients is 10 mg, taken prior to anticipated sexual activity.
- The dose may be increased to 20 mg or decreased to 5 mg, based on individual efficacy and tolerability. The maximum recommended dosing frequency is once per day in most patients.
- Tadalafil for use as needed was shown to improve erectile function compared to placebo up to 36 hours following dosing. Therefore, when advising patients on optimal use of tadalafil, this should be taken into consideration.

Tadalex for Once Daily Use for Erectile Dysfunction
- The recommended starting dose of tadalafil for once daily use is 2.5 mg, taken at approximately the same time every day, without regard to timing of sexual activity.
- The tadalafil dose for once daily use may be increased to 5 mg, based on individual efficacy and tolerability.

Tadalex for Once Daily Use for Benign Prostatic Hyperplasia
- The recommended dose of tadalafil for once daily use is 5 mg, taken at approximately the same time every day.
- When therapy for BPH is initiated with tadalafil and finasteride, the recommended dose of **Tadalex** for once daily use is 5 mg, taken at approximately the same time every day for up to 26 weeks.

Tadalex for Once Daily Use for Erectile Dysfunction and Benign Prostatic Hyperplasia
The recommended dose of tadalafil for once daily use is 5 mg, taken at approximately the same time every day,without regard to timing of sexual activity.
Renal Impairment
Tadalex for Use as Needed:
- Creatinine clearance 30 to 50 mL/min: A starting dose of 5 mg not more than once per day is recommended, and the maximum dose is 10 mg not more than once in every 48 hours.
- Creatinine clearance less than 30 mL/min or on hemodialysis: The maximum dose is 5 mg not more than once in every 72 hours.

Tadalex for Once Daily Use:
Erectile Dysfunction:
- Creatinine clearance less than 30 mL/min or on hemodialysis: Tadalafil for once daily use is not recommended.

Benign Prostatic Hyperplasia and Erectile Dysfunction/Benign Prostatic Hyperplasia
- Creatinine clearance 30 to 50 mL/min: A starting dose of 2.5 mg is recommended. An increase to 5 mg may be considered based on individual response.
- Creatinine clearance less than 30 mL/min or on hemodialysis: **Tadalex** for once daily use is not recommended.

Hepatic Impairment
Tadalex for Use as Needed:
- Mild or moderate: The dose should not exceed 10 mg once per day. The use of tadalafil once per day has not been extensively evaluated in patients with hepatic impairment and therefore, caution is advised.
- Severe: The use of tadalafil is not recommended.

Tadalex for Once Daily Use:
- Mild or moderate (Child Pugh Class A or B): Tadalafil for once daily use has not been extensively evaluated in patients with hepatic impairment. Therefore, caution is advised if tadalafil for once daily use is prescribed to these patients.
- Severe: The use of tadalafil is not recommended.

ADVERSE REACTIONS
Most common adverse reactions (≥2%) include headache, dyspepsia, back pain, myalgia, nasal congestion, flushing, and pain in limb.

WARNINGS AND PRECAUTIONS
- Patients should not use tadalafil if sex is inadvisable due to cardiovascular status.
- Use of tadalafil with alpha-blockers, antihypertensives or substantial amounts of alcohol (≥5 units) may lead to hypotension.
- Tadalafil is not recommended in combination with alpha-blockers for the treatment of BPH because efficacy of the combination has not been adequately studied and because of the risk of blood pressure lowering. Caution is advised when tadalafil is used as a treatment for ED in men taking alpha-blockers.
- Patients should seek emergency treatment if an erection lasts >4 hours. Use tadalafil with caution in patients predisposed to priapism.
- Patients should stop tadalafil and seek medical care if a sudden loss of vision occurs in one or both eyes, which could be a sign of non-arteritic anterior ischemic optic neuropathy (NAION). Tadalafil should be used with caution, and only when the anticipated benefits outweigh the risks, in patients with a history of NAION. Patients with a "crowded" optic disc may also be at an increased risk of NAION.
- Patients should stop tadalafil and seek prompt medical attention in the event of sudden decrease or loss of hearing.
- Prior to initiating treatment with tadalafil for BPH, consideration should be given to other urological conditions that may cause similar symptoms.

USE IN SPECIFIC POPULATIONS
Pregnancy: Tadalafil is not indicated for use in females. There are no data with the use of tadalafil in pregnant women to inform any drug-associated risks for adverse developmental outcomes.
Lactation: Tadalafil is not indicated for use in females. There is no information on the presence of tadalafil and/or metabolites in human milk, the effects on the breastfed child, or the effects on milk production.
Pediatric Use: Tadalafil is not indicated for use in pediatric patients. Safety and efficacy in patients below the age of 18 years have not been established.
Geriatric Use: No dose adjustment is warranted based on age alone. However, a greater sensitivity to medications in some older individuals should be considered.

DRUG INTERACTIONS
Nitrates: Concomitant use of nitrates in any form is contraindicated.
Alpha-Blockers: ED - When tadalafil is coadministered with an alpha-blocker in patients being treated for ED, patients should be stable on alpha-blocker therapy prior to initiating treatment, and tadalafil should be initiated at the lowest recommended.
BPH - tadalafil is not recommended for use in combination with alpha-blockers for the treatment of BPH.
CYP3A4 Inhibitors: Tadalafil for Use as Needed - For patients taking concomitant potent inhibitors of CYP3A4, such as ketoconazole or ritonavir, the maximum recommended dose is 10 mg, not to exceed once every 72 hours.
Tadalafil for Once Daily Use - For patients taking concomitant potent inhibitors of CYP3A4, such as ketoconazole or ritonavir, the maximum recommended dose is 2.5 mg.

CONTRAINDICATIONS
- Administration of tadalafil to patients who are using any form of organic nitrate, either regularly and/or intermittently, is contraindicated. In clinical pharmacology studies, tadalafil was shown to potentiate the hypotensive effect of nitrates.
- Tadalafil is contraindicated in patients with a known serious hypersensitivity to tadalafil.
- Do not use tadalafil in patients who are using a Guanylate Cyclase stimulator, such as riociguat. PDE5 inhibitors, including tadalafil, may potentiate the hypotensive effects of GC stimulators.

OVERDOSAGE
Single doses up to 500 mg have been given to healthy subjects, and multiple daily doses up to 100 mg have been given to patients. Adverse events were similar to those seen at lower doses. In cases of overdose, standard supportive measures should be adopted as required. Hemodialysis contributes negligibly to tadalafil elimination.

PRESENTATION
Tadalex Tablet 2.5 mg : 10's pack
Tadalex Tablet 5 mg : 10's pack
Tadalex Tablet 10 mg : 10's pack
Tadalex Tablet 20 mg : 10's pack

DOSAGE
As directed by the physician.

INSTRUCTIONS
Store below 30°C.
Protect from sunlight & moisture.
Keep out of reach of children.
To be sold on the prescription of registered medical practitioner only.

خوراک:
ڈاکٹر کی ہدایت کے مطابق استعمال کریں۔

ہدایات:
دوا کو ۳۰ ڈگری سینٹی گریڈ سے کم درجہ حرارت پر رکھیں۔
دھوپ اور نمی سے بچائیں۔
بچوں کی پہنچ سے دور رکھیں۔
صرف مستند ڈاکٹر کے نسخے پر فروخت کی جائے۔

Detailed prescribing information is available on request.

Manufactured for:
AsianContinental (Pvt.) Ltd.
D-32, S.I.T.E. II, Super Highway,
Karachi-Pakistan.
www.asiancontinental.com

ISO 9001/ 14001 BUREAU VERITAS Certification
An ISO Certified Company

Manufactured by:
Wimits Pharmaceuticals (Pvt.) Ltd.
129, Sunder Industrial Estate,
Lahore.