LUVOX * (fluvoxamine maleate) Tablets

Brief Summary (For full Prescribing Information refer to package insert.)

INDICATIONS AND USAGE

INDRATIONS AND USAGE UWX folders on advoced for the teamment of classissions and compulsions in portients with Obsessive Computive Disorder (DOL), as defined in the DSMIHR. The obser-sions or computions cause model distress, are mine-consuming, or significantly interfere with spoil or occupational functioning. The efficient (UVX) folders was esti-blicat in two I)-over this with observative computies augmentary with the disposis of Dissessive Computies Disorder additional DSMIHR. Observative Computies Disorder is is distributed by recurrent and parsistent idees, thoughts, impacts or images (observing) that are age-dystenic and/or repetitive, purposed), and internitional behaviors (computions) that are recognized by the person as excessive or unreasonable. The efficiences of UVXX folders for admeter periods should repetid the no. 10 weeks, has not been systematically evoluted in particularity and the distributed and because laters to use UVXX folders for admeter periods and periods repositioned admeters. m usefulness of the drug for the individual potient

CONTRAINDICATIONS

Condiministration of terfenortime astemizate or risonade with LLVDX Tablets is contraindicated (see WARNINGS and PRECAUTIONS) LLVDX Tablets are contraindicated in ents with a history of hypersensitivity to fluvoxamine male panents with a WARNINGS

Potential for later n with M. ine Oxidase inhibitors. In patients receiving another serot in reuntake inhibitor drug in con rthermia, rigidity agitamon pro-on started on nied that HIVOX Tabl ets, at leas

Potential for Interaction with Manacamber Oxidese inhibitors. In partients receiving enother serotomin respirate inhibitor drug in combination menomiane axidese inhibitors (MAOI), there have been reports of serices, sometimes fard, reactions including hyperthermica, it synodause, activations instability with possible region flacturitors of virial sprus, and menuti strutus druges that include exteme egistritie grassing to delifiam and come. These mentions have data been reported in partients who have discontinued that drug parethermica, its a MAOI. Some asses presented with features resembling neurologitic malignant syndrome. Therefore, it is recommanded that UVOX is are be seed in combination with a MAOI. Attribute MAOI and MAOI attribute and MAOI attribute stapping UVOX Tables, at 2 weeks should be allowed before storting a MAOI. Potential farbanding, Astenzizado, and Cosprible Interactions. Terfenadine, astenzizade, and its substantial attribute and attribute and attribute that be and a constrained with a theorem and the transformation with a matching theorem and the set proteing in horased plasme concentrative of parent drugs, increased plasme constraints of terfenadine, astenzizado by the UXOX is proteogetion and torsades do pointers trugt and trady attribute tradycardis, astenzizade, a potent inhibitor of IIIAA, blacks the metabolized by the cytool Potenzial farbandes states and a potent tradycardis, astenziade, a potent inhibitor of IIIAA, blacks the interbolized by the UXOX is be see to be and devisioned by the MAOI environments fard. As anote below, a statemidal by the MAOI environment tordycardis, astenziade, and below, astenziade by the MAOI Exercention is anotempotenze and the trade and any distribute of the flow astential historics to the statemation in terretorial to vorxumine with adprazolean. Consequently, it is recommended that throw to be metabolized by the MAOI extension or chardia (see CONTRAINOKATIONE and PRECAUTIONE). izole, and cisapride cause QT pharmacokinetic international nah i mial interaction of flu

Voltament with detections. Labelogency is a technological text introducting by the pool in commendation with entry entry in the continuation of the pool in commendation with entry entry intervence, are dispetite loss (CORTRANOCATONS) and RECAUTIONS). Other Potentially Important Drug Interactions: (Moss ee PKEUMINGS- log interactions) Bearcadargeniess: Bearcadargenies metabolized by linearation metabel (100 mg d) and alprazolam, indication, etc.) should be used with contion because the decorate of these drugs is likely to be reduced by filearation. The decorate of bearcadargenies metabolized by glacuroindition (e.g., learategon, autogon, temacegon) is calledy to be differed by havaname. Alprazolam - When havaname metabel (100 mg d) and alprazolam (ing gid) were do administered to stored state, poron commonitors and them paramoticines (commetes (AUL Cymps, Tr.)) of deprazolam were approximately twice faces observed where adprazolam was administered advec, and dearate was reduced by door SDN. The eleveted plasma diparazolam concentrations resulted in discussed systemmetal. No dosogn dyname were approximately twice faces observed where adprazolam designs advalde on the text banded of timuton to the lowest effective does is recommended. No dosogn dyname were proximitely twice faces data advective of UNOX Tabless and diagrapm advalde on advalde in an observed where a transmission of UNOX Tabless and diagrapm to especial from a study in which healty volumes tables to continuitation to advalde and text state of apprazolam (1000X Tables) and diagrapm advective of diagrapm was reduced by 50% and that of Mexamine were doministered in statementations to advant the text state state advalde and text state state state. The experimentation is the down different diagrapm and the data statement advantame metabolize, the distribution of the species different data significanty universitations that days of the data statement advantame metabolize and diagrapm advantame text advantame text advantad to the data statement and th period concentration of integrations about the normalised in a backgroup operation is reported to carrier and the second of the

PRECAUTIONS

General

General Activation of Mania/Myponamile: During premoketing studies involving primarily degressed patients, hypornania or mariae occurred in approximately 1% of patients heated with fluoraamine. Activation of mania/hypornania has also been reported in a small proportion of patients with major diffective disarder who were treated with other motated antidegressoms. As while all antidegressoms, UVIXI tables shauld be used cantidacylin patients with antiport diffective disarder who were treated with other ess servers were reported in 0.25 of Monomimentered patients. UVIXI tables should be used cantidacylin patients with degressor in associations of patients with a patients. Servers, Braine Schwarz, Braines, UVIXI tables should be used cantidacylin patients with degression and instruct of servers. Servers were reported in 0.25 of Monomimentered patients. UVIXI tables should be used cantidacylin patients with degression and association with another parave static servers. Braines and begression and association with another parave static servers. Braines and Brai

apprecisie screen in potents with a revent hotory of myocradia infraction or unstable heard desses. Privatels with these dognoses were systematicable excluded from many direct studies during the product's permokeling testing. Evolution of the electrocardograms for potents with depression or OCD who potificipated in premokeling studies revended no differences between thorounding and the emergence of clinicably important ISC damps. In potents with lives dystruction, but and bufferences between thorounding and produce in the emergence of clinicably important ISC damps. In potents with lives dystruction, but and bufferences between thorounding and the emergence of clinicably important ISC damps. In potents with lives dystruction, but and bufferences between thorounding and the locable stable with potents for whom the prescribe UNOX liables. Interference with Cognetive ar Meter Performance: Since any psychocytrie drug may impair judgement, thinking, or moto skills, patients should be contioned about operating lazandous mochine-ery, including automobiles, until they are certain that UNOX liables therapy does not obviersally differs their diality to engage in such achieving. Preparatory: Patients Should be achieved to only their physicians if they are bears freeding on infant. Cee PEKGAUINOS: Nusing Morthers). Concentration Mechanics: Privates should be achieved to only their physicians if they are bears freeding on infant. Cee PEKGAUINOS: Nusing Morthers). Concentration Mechanics: Privates should be achieved to only their physicians if they are bears freeding on infant. See PEKGAUINOS: Nusing Worthers). Concentration for discould yimportant interactions: whitUNOX liables. Acknobel: with other psychotogic meckanics, patients should be achieved to only their physicians if they are bears freeding on infant. See PEKGAUINOS: Nusing Worthers). Concentration Mechanics: Patients should be achieved to only their physicians if they are bears freeding on infant. See PEKGAUINOS: Nusing Worthers). Concentration Mechanics: Privat Laboratory Tests

specific laboratory tests recommended There are a

Parients isolad be obiest for notify freig physicians if they develop a rash, heres, or a related allergic phenomenon during theopy with LUNOX fables. **Checretery Pist**These are no specific laboratory tests recommended. **Presential Instructions with Drags that halbet or are Matcholized by Cytochrome P450 (szygmes:** Multiple hepotic cytochrome P450 (VPF450)
mayness are model in the oxidante bottonoformation of a large nurber of structurily different dugs and endogenous compounds. It is evolvelia in the isothery of the dustice of the oxidante bottonoformation of a large nurber of structurily different dugs and endogenous compounds. It is evolvelia in the indexed of the dustice of the oxidante bottonoformation of a large nurber of structurily different dugs and endogenous compounds. It is evolvelia in the indexed of the dustice of th

the maximum human daily dose as a mg/m¹ bass. *Meragenesis*: we evidence of mutogenic patential was observed in a mass microardeus test, an in who chro-macane aluaration test, an the Ames microbid mutogen test with an without metabolic activation. *Impairment of Fartility*: In tentility states and all and end tende test, as plo 83 mg/s/g/s/gy and/of fluxowane models. (approxamative) J times the maximum human daily does an a mg/m¹ basis, had ne flet on maniger deformance, up to 80 uration of aestation, or preanancy rate

Pregr

Travitations). Travitagenic Effects - Prognancy Category C. In teratology studies in nots and nabitris, daily and doses of fluoraamine malente of up to 80 and 40 mg/kg, respec Tively (opproximately 2 times the maximum human daily dose on a mg/m² basis) caused no fead malformations. Howeve, in other reproduction studies in which preg Herargement criters' or regionery currency or the introducy stoles in tractical toxics, an inclusion with the intervention in the intervention of the interventing of the intervention of ikely occurred seconomy. ies in program women. Flowcarnine mnosuw ... **Labor and Delivery** ¹¹ Affort of fluwcarnine on labor and delivery in humans is unknown.

As for more your drugs, fluxwonine is secreted in human breast milk. The decision of whether to discontinue runsing or to discontinue the drug should take into account the patential for serious adverse effects from exposure to fluxwarnine in the nursing infant as well as the patential benefits of UVOX^{IM} Tablets therapy to the mother. Pediatric Use

Safety and effectiveness of LUVOX Tablets in individuals below 18 years of age have not been established.

Geriatric Use

Approximately 230 patients participating in controlled premarketing studies with LUVOX Tablets were 65 years of age or over. No overall differences in safety were abserved hypocontrol is to blanch pompaning in the measurement and in process minit to only defined process or again the too the second second process or again the second process or again the second second second process or again the second second proces or again the second second process or again the se ADVERSE REACTIONS

Associated with **Discontinuation of Treatment** – Of the 1087 0CD and depressed potients treated with Euroacomiene molecte in contralled clinical trials conduct-ed in North America, 22% discontinued treatment due to an adverse event. The most common events (21%) associated with discontinuation and considered to be drug relat-

ed in Nath America, 22% Sconthaud hearment due to an obsess event. The most common events (21%) associated with dispont of learning principal sociated with dispont of learning branching. The most common events (21%) associated with dispont of learning principal sociated with the sociated sociated sociated with learning principal sociated sociated with the use of UNOX fables to the dispersive (PHI300). In greated, oriented of links of OMA sociated society and sociated ogy. The presence sends be write har these fluges control to easily to pread the indexice of size affects in the could or usion mending principation write principation treatists and other factors may differ than those that prevaid the directions. The cited figures, however, do provide the prescribing physician with some basis for estimating the relative contribution of drug and non-drug factors to the side-affect indexicer are in the populations studied. Adverse Events in OOP Precise Controlled Studies: Which are Markadly Different (defined as at least a two-fold difference) in Rate from the Pooled Event Rates in OOD and Depression Physicab Controlled Studies: The events in OOC studies with a two-fold difference) in Rate from the Pooled Event Rates in OOD and Depression Physicab Controlled Studies: The events in OOC studies with a two-fold difference) in Rate from the Pooled Event Rates in OOD and Depression Physicab Controlled Studies: The events in OOC studies with a two-fold difference) in Rate from time to Physicab controlled resolution amblypig (mstry blured vision). Additionally, there was on opproximate 25% decrease in noces. The events in OO studies with a two-fold more large the factor difference in the factor the two-fold more the provide the event mate in OO studies with a two-fold more factor the the factor difference in the compared to the factor the factor the factor the two-fold more the event in the compared to the event the top of the event to the compared to the event the top of the event to the compared to the event to the top of the event to the top of the event to the top of the event top of the even to even track in the constant water and the set of the

Vital Sign Changes Comparisons of fluvoxam Comparisons of Ruoxiamine maleate and placebo groups in separate pools of short+tern DCD and depression trials on (1) median change from baseline on various vital signs variables and on (2) incidence of parients meeting criteria for potentially important changes from baseline on various vital signs variables revealed no important differences between fluvoxamine malente and placeba

Cervent invocamine molece on paceto. Laborator of Maccanine molecte and paceto grazas in separate pools of short-term OCD and depression thicks on (1) median change from baseline on various serum chemistry, hematology, and utinalysis variables and on (2) indicence of patients meeting criteria for patentially important changes from baseline on various serum chemistry, hematolysy, and utinalysis variables and on (2) indicence of patients meeting criteria for patentially important changes from baseline on various serum chemistry.

ECG Che

ECG Changes Comportions of invoxomine maleote and plexable groups in separate pools of shortNerm OCD and depression hids on (1) mean change from baseline on various ECG vari-debs and on (2) incidence of priferins meeting afferers for potentially important changes from baseline on various ECG variables revealed no important differences between nte and ala

Tedie 2: TREATMENT-EMERGENT ADVERSE EVENT INCIDENCE RATES BY BODY SYSTEM IN OCD AND DEPRESSION POPULATIONS COM-BINED¹ (throadmine sc, diotedo by primer percentage): BODY AS WHOLE: Hoodade (25 sc. 20), Ashenia (14 sc. 6), Fill-Syndmine (35 sc. 2), Follis (27 sc. 1), CARDOVASCULAR: Flohtmins (35 sc. 2), INGENTY SYSTEM: Associated (40 sc. 14), Burnel (11 sc. 2), Carasiton (10 sc. 3), Expension (10 sc. 3), Expension (10 sc. 3), Expension (21 sc. 3), Following (3 sc. 2), INERVOUS SYSTEM: Somolence (22 sc. 8), Hauman (21 sc. 3), Expension (21 sc. 3), Following (3 sc. 3), Distance (24 sc. 3), Expension (24 sc. 1), NERVOUS SYSTEM: Somolence (22 sc. 8), Hauman (22 sc. 3), Following (3 sc. 2), Distance (24 sc. 3), Expension (2 sc. 1), INERVOUS SYSTEM: Somolence (22 sc. 8), Hauman (24 sc. 1), Expension (2 sc. 1), INERVOUS SYSTEM: Symposine (22 sc. 1), Expension (2 sc. 1), Interviewed (23 sc. 2), Depression (2 sc. 1), INERVOUS SYSTEM: Symposine (2 sc. 1), Symposine (2 sc. 1), Team (2 sc. 2), Somoling (3 sc. 2), Database (2 sc. 3), Following (3 sc. 2), Database (2 sc. 3), Interviewed (2 sc. 1)), Depression (2 sc. 1)), INERVOUS SYSTEM: Upper Respiratory Intertion (9 sc. 5), Diganee (2 sc. 1)), Team (2 sc. 3), SPECIAL SENSES, Toste Provesion (3 sc. 1), Hauter (3 sc. 2), LINDGENTIAL: Alternard Bocluthrial (4 sc. 1), Dimory Frequency (3 sc. 2), Interviewer (2 sc. 1)), Anotypoint (2 sc. 1)), LINDGENTIAL: Alternard Bocluthrial (4 sc. 1), Binary Frequency (3 sc. 2), Interviewer (2 sc. 1)), Anotypoint (2 sc. 1)), LINDGENTIAL: Alternard teams, appetite increase, bock pair, confiction, dysmenorthen, (ever, inflation, leg comps, migraine, myldia, pair, appetite, increase, bock pair, and, confistion, dysmenorthen, alter, and index (3 sc. 1), Houter (3 sc. 3), SPECIAL SENSES, Toste Provession (3 sc. 3), Lindows, the indevent (3 sc. 3), SPECIAL SENSES, Toste Provession (3 sc. 3), Toble 2: TREATMENT-EMERGENT ADVERSE EVENT INCIDENCE RATES BY BODY SYSTEM IN OCD AND DEPRESSION POPULATIONS COM

vision ", Washy" deviged exolution: ", Incodence tasked on number of male potents. Other Events Observed During the Premarketing Evaluation of LUVOX Tablets During premarketing clinical indix concluded in North America and Europe, multiple does of fluxouonine maleate were administered for a combined total of 2737 patient exposures in patients suffering OLD or Major Depressive Boorder. Untowed events associated with this exposure were recorded by clinical investigations using description ter-minology of their own choosing. Consequently, it is not possible to provide a meaningful estimate of the proportion of individual experiencing objects events without first grouping similar types of untowerd events into a limited (i.e., leakced) number of similar event chargives. In the tabletations which follow, a standard COSIARRhosed Dictionary terminology his been used to classify reported obvece events. The COSIAR term for an event was so general as to be uninformative, it was replaced with a music information. The Thomponer executed the patient observation of the 272 officient event events and the event of the 273 officient event events and the event events the source to the patient observation event events and there of the provide event was source to the major terminology have events. The COSIAR term is the constraint event events and there of the provide events and the event events the source to the patient observation. Transport of the two construction. Consequently, its hold possible to provide a meaning or simule on the program of informations which follow, a strandord CSDRF location and provide simulations which follow, a strandord CSDRF location and the strandord event decrepse. In the tobulations which follow, a strandord CSDRF location and the strandord event decrepse. In the tobulations which follow, a strandord CSDRF location are event with the location and the strandord event decrepse. In the tobulations which follow, a strandord CSDRF location are event with the location and the strandord cSDRF location are event of the type ched on or least one excasion while receiving fluxournine nudexis. All reported events are included in the last below, with the following excaptions. In those events for which a duo graces was considered entero (Le., neglistica) and events on the locations are evented with a graces events of which a duo graces was considered entero (Le., neglistica) and events on the locations are evented in the last periods of the strandord controlling control and depression direct thinks, are excluded. If here periods and any constraint and presents of the location and unintered pregnancy) are antitled, and sign extension in the strandord material service are included. It is important to manipulate the dustified within body system cotapolises and enumerate in arder of decreasing frequency using the following derivation and unintered pregnancy are antiticated and exact and and any constraints, and are advected present present in the strandord in the strandord material service are antiticated and and antitication and event and and decrease events are those accurating and error accusions in a fast 1/100 prients. Body are a MMode: Encendent cocale failership in molecus, thereaut and events are for a cortanic and present an

1 Based on the number of females.; 2 Based on the number of males

Function in remains of manaces, is cause of the native of mines. MonUS ensurance international properties to king UUVX Tables that have been received since marker introduction and ore of unknown causal relationship to UUVX Tables use include: taxis epidermol nerophysis, Stevens-Johnson syndrome, Henach/Scheenlein puppuro, ballaus exuption, priopism, agranulocytosis, neuropathy, splastic one-mic, acquisity lattic receition, hypometrimor, outer end Tables, and severe claimski with fever when fluorounnine was condministered with antipychoits methodurta. CAUTION: Federal low prohibits dispensing without prescription 4E1252 Rev 9/95

Reference: 1. Data on file. Solvay Pharmaceuticals. Inc.

Pharmacia & Upjohn



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ESTABLISHED THERAPY FOR OCD

EFFECTIVE CONTROL OF OBSESSIONS AND COMPULSIONS^{1*}

LOW INCIDENCE OF AGITATION

 $(2\% vs 1\% \text{ for placebo})^1$

LOW INCIDENCE OF SEXUAL DYSFUNCTION

LUVOX® Tablets vs placebo[†]: decreased libido 2% vs 1%; delayed ejaculation 8% vs 1%; anorgasmia 2% vs 0%; impotence 2% vs 1%

FAVORABLE SAFETY PROFILE

- Relatively low incidence of anticholinergic side effects in controlled trials of OCD and depression, LUVOX® Tablets vs placebo¹: dizziness 11% vs 6%; constipation 10% vs 8%; dry mouth 14% vs 10%
- The most commonly observed adverse events compared to placebo were somnolence 22% vs 8%, insomnia 21% vs 10%, nervousness 12% vs 5%, nausea 40% vs 14%, abnormal ejaculation 8% vs 1%, asthenia 14% vs 6%¹
- Concomitant use of LUVOX® Tablets and monoamine oxidase inhibitors is not recommended¹

FLEXIBLE DOSING

Initial Dose: 50 mg once a day HS Dose Range: 100 to 300 mg/day

COMPREHENSIVE SAFETY DATABASE

(Worldwide Exposure for Reporting Overdose[‡])¹

- Data from 40 countries
- Over 12 million patients treated
- More than 37,000 patients studied in clinical trials

LUVOX fluvoxamine maleate 50 mg & 100 mg SORED TABLETS **A SELECTIVE SEROTONIN REUPTAKE INHIBITOR**

> *Effectiveness not established beyond 10 weeks in controlled trials. [↑]Parameters occurring ≥ 1% with fluvoxamine maleate. [‡]Prescribers should write the smallest tablet quantity consistent with good patient management to reduce overdose risk. *Please see brief summary of prescribing information on adjacent page.*

DOUBTS