THE BRITISH JOURNAL OF PSYCHIATRY

MAY 1998 VOL. 172

EDITORIALS

REVIEW ARTICLES

Self-injurious behaviour in people with a learning disability

PAPERS

PRELIMINARY REPORT

COLUMNS

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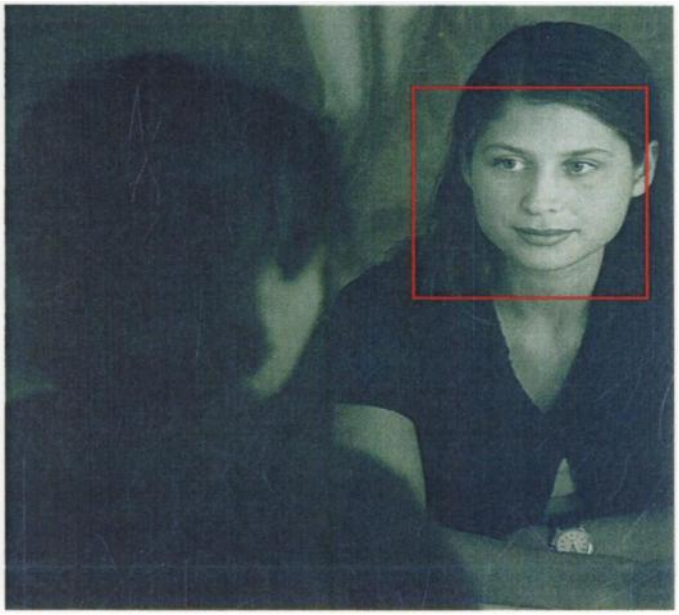
Edward H. Hare	1973–77	Hugh L. Freeman	1984–93

Founded by J. C. Bucknill in 1853 as the *Asylum Journal* and known as the *Journal of Mental Science* from 1858 to 1963.

Debbie doesn't know that Cipramil is now indicated for panic disorder



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As a patient with Panic Disorder, Debbie is beginning to appreciate the value of the Cipramil treatment that her doctor has newly prescribed.

Of course, Debbie would no more talk of the recently extended indication for Cipramil than its high selectivity[1,2], good tolerability[3], and low risk of drug interactions[4,5,6]. She just recognises the difference that Cipramil makes to the stability and quality of her life.

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citalopram
now indicated for panic disorder

Presentation: 'Cipramil' tablets 10 mg: PL 0458/0057, each containing 10 mg of citalopram as the hydrobromide. 28 (OP) 10 mg tablets £12.77. 'Cipramil' tablets 20 mg: PL 0458/0058, each containing 20 mg of citalopram as the hydrobromide. 28 (OP) 20 mg tablets £21.28. **Indications:** Treatment of depressive illness in the initial phase and as maintenance against relapse/recurrence. Treatment of panic disorder, with or without agoraphobia. **Dosage:** ***Treating depression: Adults:*** 20 mg a day. Depending upon individual patient response, this may be increased in 20 mg increments to a maximum of 60 mg. Tablets should not be chewed, and should be taken as a single oral daily dose, in the morning or evening without regard for food. Treatment for at least 6 months is usually necessary to provide adequate maintenance against the potential for relapse. ***Treating panic disorder:*** 10 mg daily for the first week, increasing to 20 mg daily. Depending upon individual patient response, dosage may be further increased to a maximum of 60 mg daily. Depending upon individual patient response, it may be necessary to continue treatment for several months. ***Elderly:*** 20 mg a day increasing to a maximum of 40 mg dependent upon individual patient response. ***Children:*** Not recommended. ***Reduced hepatic/renal function:*** Restrict dosage to lower end of range in hepatic impairment. Dosage adjustment not necessary in cases of mild/moderate renal impairment. No information available in severe renal impairment (creatinine clearance <20ml/min). **Contra-indications:** Combined use of 5-HT agonists. Hypersensitivity to citalopram. ***Pregnancy and Lactation:*** Safety during human pregnancy and lactation has not been established. Use only if potential benefit outweighs possible risk. **Precautions:** Driving and operating machinery. History of mania. Caution in patients at risk of cardiac arrhythmias. Do not use with or within 14 days of MAO inhibitors: leave a seven day gap before starting MAO inhibitor treatment. Use a low starting dose for panic disorder, to reduce the likelihood of an initial anxiogenic effect (experienced by some patients) when starting pharmacotherapy. **Drug Interactions:** MAO inhibitors (see Precautions). Use lithium and tryptophan with caution. Routine monitoring of lithium levels need not be adjusted. **Adverse Events:** Most commonly nausea, sweating, tremor, somnolence and dry mouth. With citalopram, adverse effects are in general mild and transient. When they occur, they are most prominent during the first two weeks of treatment and usually attenuate as the depressive state improves. **Overdosage:** Symptoms have included somnolence, coma, sinus tachycardia, occasional nodal rhythm, episode of grand mal convulsion, nausea, vomiting, sweating and hyperventilation. No specific antidote. Treatment is symptomatic and supportive. Early gastric lavage suggested. **Legal Category:** POM 24.1.95. Further information available upon request. Product licence holder: Lundbeck Ltd., Sunningdale House, Caldecotte Lake Business Park, Caldecotte, Milton Keynes, MK7 8LF. ® 'Cipramil' is a Registered Trade Mark. © 1997 Lundbeck Ltd. Date of preparation: April 1997. 0897/CIP/501/044

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A NEW SELECTIVE NARI. LIFTS DEPRESSION.
HELPS RESTORE SOCIAL INTERACTION.

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ABBREVIATED PRESCRIBING INFORMATION
Presentation: Tablets containing 4mg reboxetine. **Indications:** Use in the acute treatment of depressive illness, and maintenance of clinical benefit in patients responsive to treatment. **Posology and method of administration:** *Adults* 4 mg b.i.d. (8 mg/day) administered orally. After 3-4 weeks, can increase to 10 mg/day. *Elderly and children* Elderly patients have been studied in comparative clinical trials at doses of 2 mg b.i.d., although not in placebo controlled conditions. There is no experience in children and therefore reboxetine cannot be recommended in either of these groups. *Renal/Hepatic* **Special warnings and precautions for use:** Close supervision is required for subjects with a history of convulsive disorders and must be discontinued if the patient develops seizures. Avoid concomitant use with MAO inhibitors. Close supervision of bipolar patients is recommended. Close supervision should be applied in patients with current evidence of urinary retention, glaucoma, prostatic hypertrophy and cardiac disease. At doses higher than the maximum recommended, orthostatic hypotension has been observed with greater frequency. Particular attention should be paid when administering reboxetine with other drugs known to lower blood pressure. **Interactions with other medicaments** that have a narrow therapeutic margin and are metabolised by CYP3A4 or CYP2D6 e.g. anti-arrhythmics (flecainide), anti-psychotic drugs and tricyclic anti-depressants. No pharmacokinetic interaction with lorazepam. Reboxetine does not appear to potentiate the effect of alcohol. **Pregnancy and lactation:** Reboxetine is contraindicated in pregnancy and lactation. **Effects on ability to drive and use machines:** Reboxetine is not sedative per se. However, as with all psychoactive drugs, caution patients about operating machinery and driving. **Undesirable effects:** Adverse events occurring more frequently than placebo are: dry mouth, constipation, insomnia, paraesthesia, increased sweating, required. **Package and NHS Price:** Pack of 60 tablets in blisters £19.80. **Legal Category:** POM **Marketing Authorisation Holder:** Pharmacia & Upjohn Limited, Davy Avenue, Milton Keynes, MK5 8PH, UK. **Marketing Authorisation Number:** PL 0032/0216, **Date of Preparation:** October 1997. **References: 1.** Montgomery SA. *Journal of Psychopharmacology* 1997 (in press). **2.** Dubini A. et al. *European Neuropsychopharmacol.* 1997; **7** (Suppl 1): S57-S70. **3.** Bosc M. et al. *European Neuropsychopharmacol.* 1997; **7** (Suppl 1): S57-S70. Further information is available from Pharmacia & Upjohn Limited, Davy Avenue, Knowlhill, Milton

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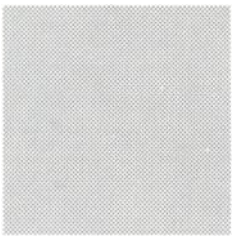

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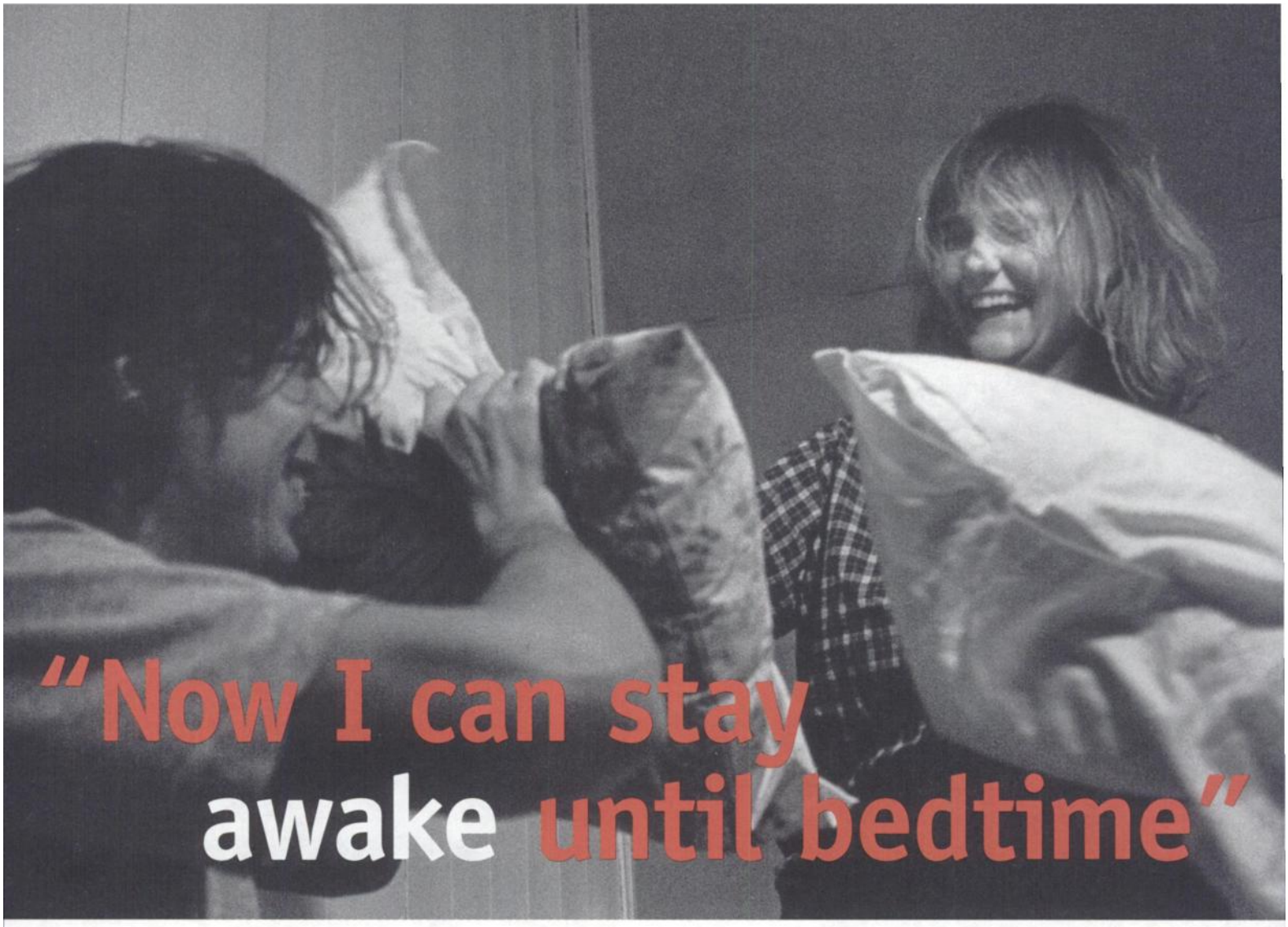

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REFERENCES

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Serdolect is not sedative, however, patients should be advised not to drive or operate machinery until their individual susceptibility is known. History of diabetes, seizures, Parkinson's disease. Symptoms of orthostatic hypotension may occur and blood pressure should be monitored during initial dose titration and in early maintenance phase. In common with other antipsychotic drugs, Serdolect lengthens the QT interval in some patients (<1.7% of patients). Electrolyte imbalance or combined use of other drugs that inhibit Serdolect metabolism can increase the risk of occurrence of prolonged QT interval. An ECG should be performed prior to use with periodic ECG monitoring during treatment. Serdolect should not be initiated or should be discontinued if the QTC_2 interval exceeds 520 msec. Hypokalaemia and hypomagnesaemia should be corrected and maintained within normal limits during treatment. If signs and symptoms of tardive dyskinesia appear, consider dose reduction or discontinuation. **Drug interactions:** [illegible] (see Precautions). Combined use of agents known to inhibit hepatic isoenzymes may necessitate lower maintenance doses. Combined use of agents

prolonged QT interval. Incidence of EPS adverse events similar to placebo. Overdosage: Symptoms have included somnolence, slurred speech, tachycardia, hypotension and transient prolongation of QT interval. There is no specific antidote. Treatment is supportive and symptomatic. Epinephrine and dopamine should not be used (may exacerbate hypotension). Cardiovascular monitoring recommended. Administration of activated charcoal and laxative should be considered. **Package quantities and basic NHS price:** 4mg tablets, £36.63 for 30 tablet pack, 12mg tablets, £102.55 for 28 tablet calendar pack. 16mg tablets, £102.55 for 28 tablet calendar pack. 20mg tablets, £102.55 for 28 tablet calendar pack. Legal category: POM. **Product Licence numbers:** 4mg: 13761/0001. 12mg: 13761/0003, 16mg: 13761/0004. 20mg: 13761/0005. **Date of last review:** April 1997. Further information is available on request from Lundbeck Limited, Sunningdale House, Caldecott Lake Business

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CLOZARIL®
clozapine

CLOZARIL ABBREVIATED PRESCRIBING INFORMATION. The use of CLOZARIL is restricted to patients registered with the CLOZARIL Patient Monitoring Service. Indication Treatment-resistant schizophrenia (patients non-responsive to, or intolerant of, conventional neuroleptics). **Presentations** 25mg and 100 mg clozapine tablets. **Dosage and Administration** Initiation must be in hospital in-patients and is restricted to patients with normal white blood cell and differential counts. Initially, 12.5 mg once or twice on the first day, followed by one or two 25 mg tablets on the second day. Increase dose slowly, by increments to reach a therapeutic dose within the range of 200 - 450mg daily (see data sheet). The total daily dose should be divided and a larger portion of the dose may be given at night. Once control is achieved a maintenance dose of 150 to 300 mg daily may suffice. At daily doses not exceeding 200mg, a single administration in the evening may be appropriate. Exceptionally, doses up to 900 mg daily may be used. Patients with a history of epilepsy should be closely monitored during CLOZARIL therapy since dose-related convulsions have been reported. Patients with a history of seizures, as well as those suffering from cardiovascular, renal or hepatic disorders, together with the elderly need lower doses (12.5 mg given once on the first day) and more gradual titration. **Contra-Indications** Allergy to any constituents of the formulation. History of drug-induced neutropenia/agranulocytosis, myeloproliferative disorders, uncontrolled epilepsy, alcoholic and toxic psychoses, drug intoxication, comatose conditions, circulatory collapse and/or CNS depression of any cause, severe renal or cardiac failure, active liver disease, progressive liver disease or hepatic failure. **Warning** CLOZARIL can cause agranulocytosis. A fatality rate of up to 1 in 300 has been estimated when CLOZARIL was used prior to recognition of this risk. Since that time strict haematological monitoring of patients has been demonstrated to be effective in markedly reducing the risk of fatality. Therefore, because of this risk its use is limited to treatment-resistant schizophrenic patients:- 1. who have normal leucocyte findings and 2. in whom regular leucocyte counts can be performed weekly during the first 18 weeks and at least every two weeks thereafter for the first year of therapy. After one year's treatment, monitoring may be changed to four weekly intervals in patients with stable neutrophil counts. Monitoring must continue throughout treatment and for four weeks after complete discontinuation of CLOZARIL. Patients must be under specialist supervision and CLOZARIL supply is restricted to pharmacies registered with the CLOZARIL Patient Monitoring Service. Prescribing physicians must register themselves, their patients and a nominated pharmacist with the CLOZARIL Patient Monitoring Service. This service provides for the required leucocyte counts as well as a drug supply audit so that CLOZARIL treatment is promptly withdrawn from any patient who develops abnormal leucocyte findings. Each time CLOZARIL is prescribed, patients should be reminded to contact the treating physician immediately if any kind of infection begins to develop, especially any flu-like symptoms. **Precautions** CLOZARIL can cause agranulocytosis. Perform pre-treatment white blood cell count and differential count to ensure only patients with normal findings receive CLOZARIL. Monitor white blood cell count weekly for the first 18 weeks and at least two-weekly for the first year of therapy. After one year's treatment, monitoring may change to four weekly intervals in patients with stable neutrophil counts. Monitoring must continue throughout treatment and for four weeks after complete discontinuation. If signs or symptoms of infection develop an immediate differential count is necessary. If the white blood count falls below 3.0 x 10^9/L and/or the absolute neutrophil count drops below 1.5 x 10^9/L, withdraw CLOZARIL immediately and monitor the patient closely, paying particular attention to symptoms suggestive of infection. Re-evaluate any patient developing an infection, or when a routine white blood count is between 3.0 and 3.5 x 10^9/L and/or a neutrophil count between 1.5 and 2.0 x 10^9/L, with a view to discontinuing CLOZARIL. Any further fall in white blood/neutrophil count below 1.0 x 10^9/L and/or 0.5 x 10^9/L respectively, after drug withdrawal requires immediate specialised care, where protective isolation and administration of GM-CSF or G-CSF and broad spectrum antibiotics may be indicated. Colony stimulating factor therapy should be discontinued when the neutrophil count returns above 1.0 x 10^9/L. CLOZARIL lowers the seizure threshold. Orthostatic hypotension can occur therefore close medical supervision is required during initial dose titration. Patients affected by the sedative action of CLOZARIL should not drive or operate machinery, administer with caution to patients who participate in activities requiring complete mental alertness. Monitor hepatic function regularly in liver disease. Investigate any signs of liver disease immediately with a view to drug discontinuation. Resume only if LFTs return to normal, then closely monitor patient. Use with care in prostatic enlargement, narrow-angle glaucoma and paralytic ileus. Patients with fever should be carefully evaluated to rule out the possibility of an underlying infection or the development of agranulocytosis. Avoid immobilisation of patients due to increased risk of thromboembolism. Do not give CLOZARIL with other drugs with a substantial potential to depress bone marrow function. CLOZARIL may enhance the effects of alcohol, MAO inhibitors, CNS depressants and drugs with anticholinergic, hypotensive or respiratory depressant effects. Caution is advised when CLOZARIL therapy is initiated in patients who are receiving (or have recently received) a benzodiazepine or any other psychotropic drug as these patients may have an increased risk of circulatory collapse, which, on rare occasions, can be profound and may lead to cardiac and/or respiratory arrest. Caution is advised with concomitant administration of therapeutic agents which are highly bound to plasma proteins. Clozapine binds to and is partially metabolised by the isoenzymes cytochrome P450 1A2 and P450 2D6. Caution is advised with drugs which posses affinity for these isoenzymes. Concomitant cimetidine and high dose CLOZARIL was associated with increased plasma clozapine levels and the occurrence of adverse effects. Concomitant fluoxetine and fluvoxamine have been associated with elevated clozapine levels. Discontinuation of concomitant carbamazepine resulted in increased clozapine levels. Phenytoin decreases clozapine levels resulting in reduced effectiveness of CLOZARIL. No clinically relevant interactions have been noted with antidepressants, phenothiazines and type Ic antiarrhythmics, to date. Concomitant use of lithium or other CNS-active agents may increase the risk of neuroleptic malignant syndrome. The hypertensive effect of adrenaline and its derivatives may be reversed by CLOZARIL. Do not use in pregnant or nursing women. Use adequate contraceptive measures in women of child bearing potential. **Side-Effects** Neutropenia leading to agranulocytosis (See Warning and Precautions). Rare reports of leucocytosis including eosinophilia. Isolated cases of leukaemia and thrombocytopenia have been reported but there is no evidence to suggest a causal relationship with the drug. Most commonly fatigue, drowsiness, sedation. Dizziness or headache may also occur. CLOZARIL lowers the seizure threshold and may cause EEG changes and delirium. Myoclonic jerks or convulsions may be precipitated in individuals who have epileptogenic potential but no previous history of epilepsy. Rarely it may cause confusion, restlessness, agitation and delirium. Extrapyramidal symptoms are limited mainly to tremor, akathisia and rigidity. Tardive dyskinesia reported very rarely. Neuroleptic malignant syndrome has been reported. Transient autonomic effects eg dry mouth, disturbances of accommodation and disturbances in sweating and temperature regulation. Hypersalivation. Tachycardia and postural hypotension, with or without syncope, and less commonly hypertension may occur. In rare cases profound circulatory collapse has occurred. ECG changes, arrhythmias, pericarditis and myocarditis (with or without eosinophilia) have been reported, some of which have been fatal. Rare reports of thromboembolism. Isolated cases of respiratory depression or arrest, with or without circulatory collapse. Rarely aspiration may occur in patients presenting with dysphagia or as a consequence of acute overdosage. Nausea, vomiting and usually mild constipation have been reported. Occasionally obstipation and paralytic ileus have occurred. Asymptomatic elevations in liver enzymes occur commonly and usually resolve. Rarely hepatitis and cholestatic jaundice may occur. Very rarely fulminant hepatic necrosis reported. Discontinue CLOZARIL if jaundice develops. Rare cases of acute pancreatitis have been reported. Both urinary incontinence and retention and priapism have been reported. Isolated cases of interstitial nephritis have occurred. Benign hyperthermia may occur and isolated reports of skin reactions have been received. Rarely hyperglycaemia has been reported. Rarely increases in CPK values have occurred. With prolonged treatment considerable weight gain has been observed. Sudden unexplained deaths have been reported in patients receiving CLOZARIL. **Package Quantities and Price** Community pharmacies only 28 x 25mg tablets: £12.52 (Basic NHS) 28 x 100mg tablets: £50.05 (Basic NHS) Hospital pharmacies only 84 x 25 mg tablets: £37.54 (Basic NHS) 84 x 100 mg tablets: £150.15 (Basic NHS) Supply of CLOZARIL is restricted to pharmacies registered with the CLOZARIL Patient Monitoring Service. **Product Licence Numbers** 25 mg tablets: PL 0101/0228 100 mg tablets: PL 0101/0229 **Legal Category**: POM. CLOZARIL is a registered Trade Mark. Date of preparation, August 1997. Full prescribing information, including Product Data Sheet is available from Novartis Pharmaceuticals UK Ltd. Trading as: SANDOZ PHARMACEUTICALS, Frimley Business Park, Frimley, Camberley, Surrey, GU16 5SG.

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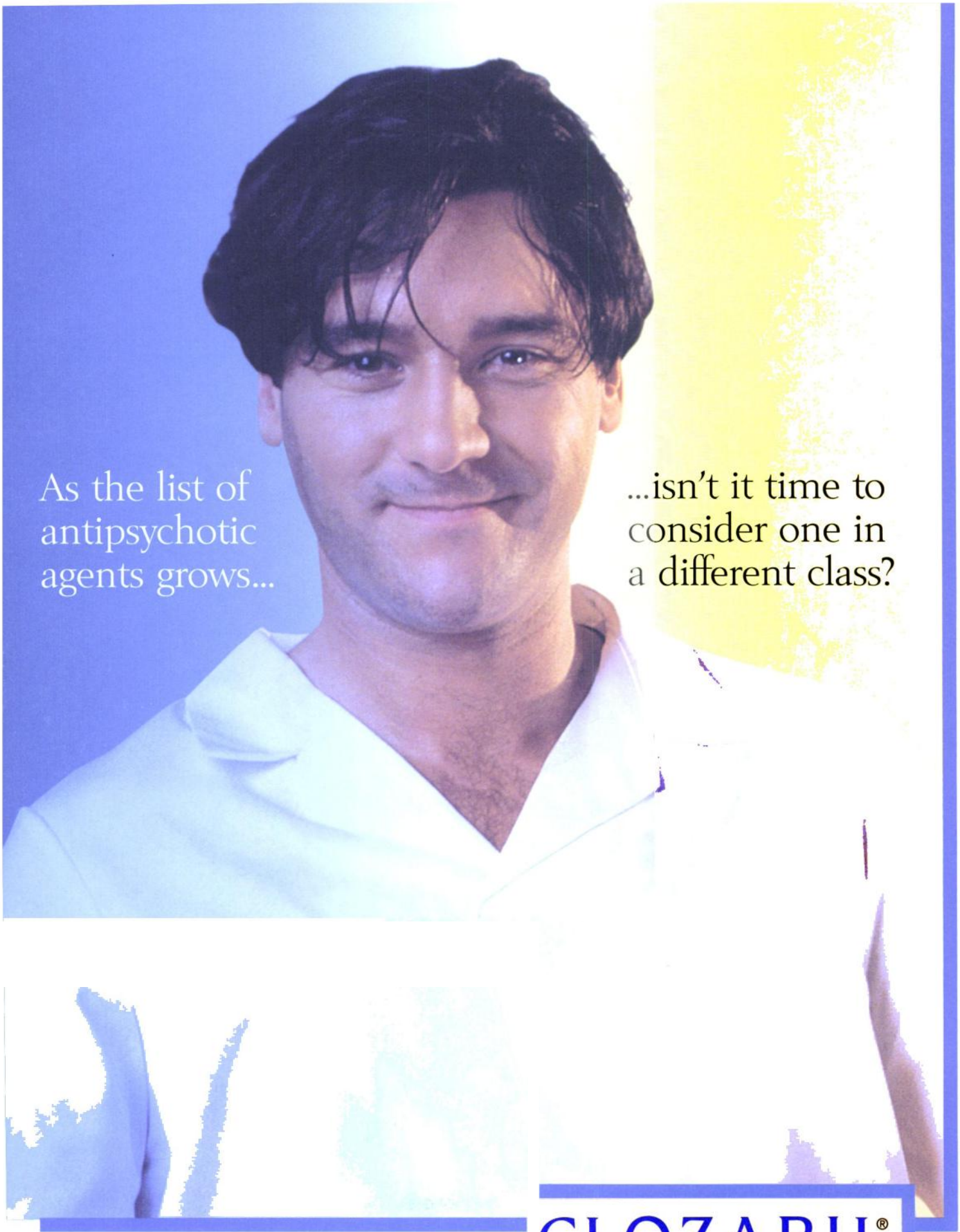

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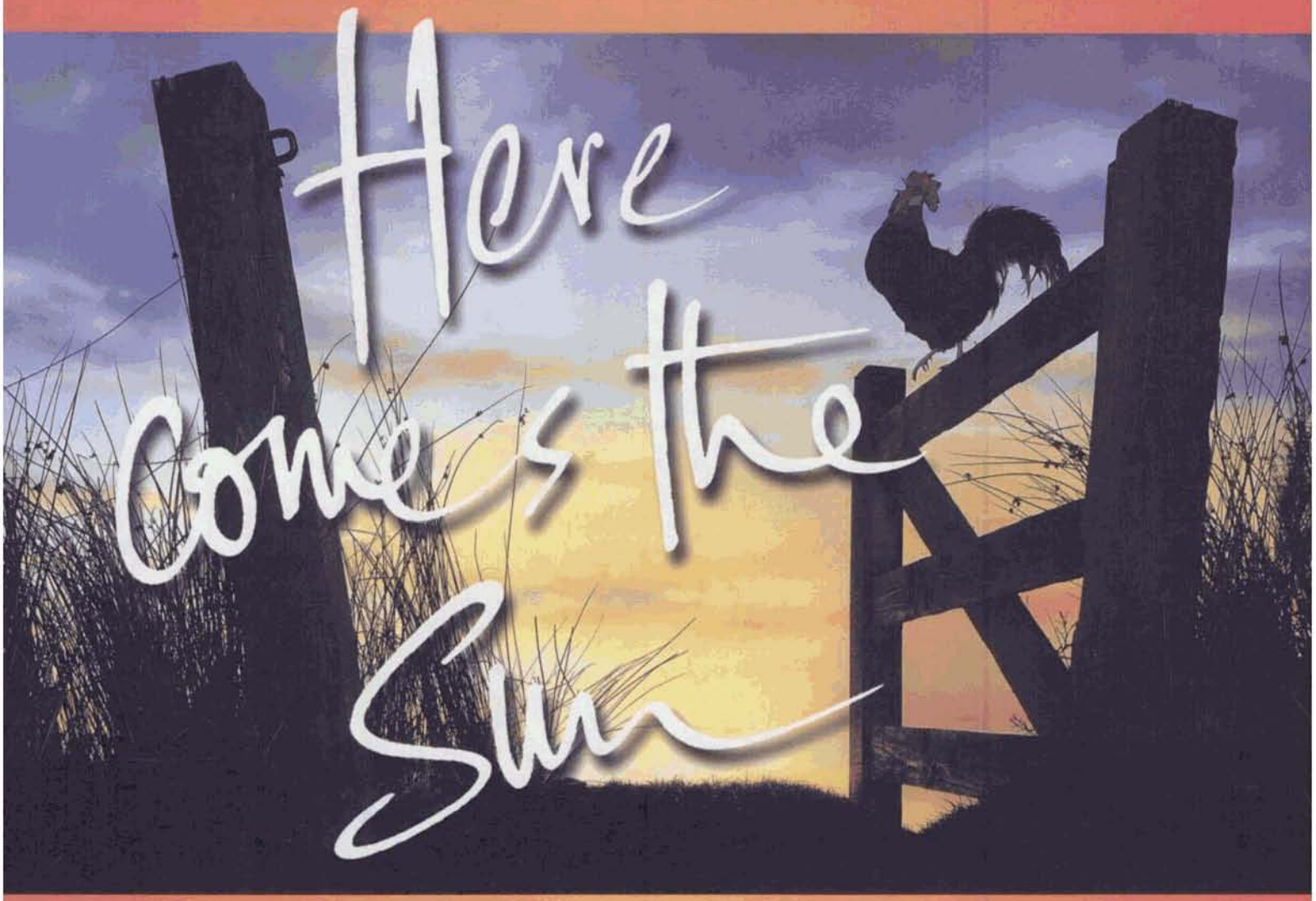
Efexor® XL venlafaxine - Prescribing information Presentation: Capsules containing 75mg or 150mg venlafaxine (as hydrochloride) in an extended release formulation. **Use:** Treatment of depressive illness. **Dosage:** *Adults (including the elderly):* Usually 75mg, given once daily with food, increasing to 150mg once daily if necessary. The dose can be increased further to 225mg once a day. Dose increments should be made at intervals of approximately 2 weeks or more, but not less than 4 days. Discontinue gradually to avoid possibility of discontinuation effects. *Children:* Contra-indicated below 18 years of age. *Moderate renal or moderate hepatic impairment:* Doses should be reduced by 50%. Not recommended in severe renal or severe hepatic impairment. **Contra-indications:** Pregnancy, lactation, concomitant use with MAOIs, hypersensitivity to venlafaxine or other components, patients aged below 18 years. **Precautions:** Use with caution in patients with myocardial infarction, unstable heart disease, renal or hepatic impairment, or a history of epilepsy (discontinue in event of seizure). Patients should not drive or operate machinery if their ability to do so is impaired. Possibility of postural hypotension (especially in the elderly). Women of child-bearing potential should use contraception. Prescribe smallest quantity of tablets according to good patient management. Monitor blood pressure with doses >200mg/day. Advise patients to notify their doctor should an allergy develop or if they become or intend to become pregnant. Patients with a history of drug abuse should be monitored carefully. **Interactions:** MAOIs: do not use Efexor XL in combination with MAOIs or within 14 days of stopping MAOI treatment. Allow 7 days after stopping Efexor XL before starting an MAOI. Use with caution in elderly or hepatically-impaired patients taking cimetidine, in patients taking other CNS-active drugs, and in patients taking drugs which inhibit *both* CYP2D6 and CYP3A4 hepatic enzymes. **Side-effects:** Nausea, insomnia, dry mouth, somnolence, dizziness, constipation, sweating, nervousness, asthenia, abnormal ejaculation/orgasm, anorexia, abnormal vision/accommodation, impotence, vomiting, tremor, abnormal dreams, vasodilatation, hypertension, rash, agitation, hypertonia, paraesthesia, postural hypotension, reversible increases in liver enzymes, slight increase in serum cholesterol, weight gain or loss, hyponatraemia. **Basic NHS price:** 75mg capsule (PL 00011/0223) - blister pack of 28 capsules: £23.97. 150 mg capsule (PL 00011/0224) - blister pack of 28 capsules: £39.97. **Legal category:** POM. Further information is available upon request from the Product Licence holder: Wyeth Laboratories, Taplow, Maidenhead, Berkshire, SL6 0PH. Date of preparation: August 1997. * trade mark Code no Z777440/0897. WEFX3-UK-JA. References: 1. Muth EA *et al.* Biochem Pharmacol 1986; 35(24): 4493-4497. 2. Muth EA *et al.* Drug Development Research 1991; 23: 191-199. 3. Rudolph R *et al.* Poster presented at the New Clinical Drug Evaluation Unit (National Institute of Mental Health). Boca Raton, Florida 1997. 4. McPartlin GM *et al.* Poster at the 10th European College of Neuropsychopharmacology meeting, Vienna, September 13th-17th, 1997. 5. Salinas E. Biol Psychiatry 1997; 42(Suppl. 1): 244S.



- EFEXOR XL ACTS DIRECTLY ON BOTH SEROTONIN AND NORADRENALINE[1,2]
- PROVEN EFFICACY VS LEADING SSRIs[3,4]
- TOLERABILITY[3,4,5] AND CONVENIENCE YOU EXPECT FROM A FIRST-LINE THERAPY

NEW ONCE DAILY

EFEXOR XL*

VENLAFAXINE 75 mg o.d.

Simply effective

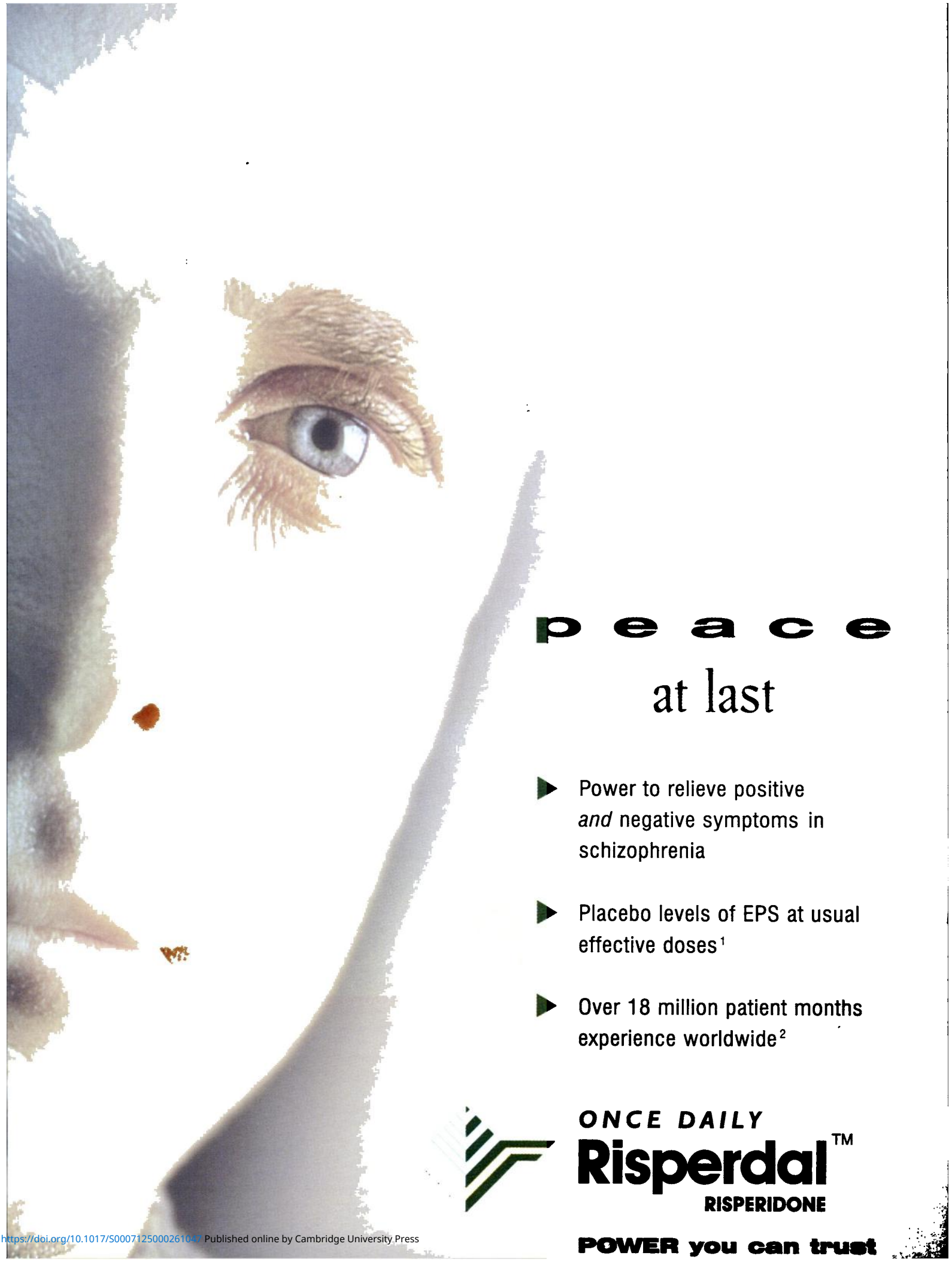
Please refer to Summary of Product Characteristics before prescribing Risperdal (risperidone). **USES** The treatment of acute and chronic schizophrenia, and other psychotic conditions, in which positive and/or negative symptoms are prominent. Risperdal also alleviates affective symptoms associated with schizophrenia. **DOSAGE** Where medically appropriate, gradual discontinuation of previous antipsychotic treatment while Risperdal therapy is initiated is recommended. Where medically appropriate, when switching patients from depot antipsychotics, consider initiating Risperdal therapy in place of the next scheduled injection. The need for continuing existing antiparkinson medication should be re-evaluated periodically. **Adults:** Risperdal may be given once or twice daily. All patients, whether acute or chronic, should start with 2 mg/day. This should be increased to 4 mg/day on the second day and 6 mg/day on the third day. However, some patients such as first-episode psychotic patients may benefit from a slower rate of titration. From then on the dosage can be maintained unchanged, or further individualised if needed. The usual effective dosage is 4 to 8 mg/day although in some patients an optimal response may be obtained at lower doses. Doses above 10 mg/day may increase the risk of extrapyramidal symptoms and should only be used if the benefit is considered to outweigh the risk. Doses above 16 mg/day should not be used. **Elderly, renal and liver disease:** A starting dose of 0.5 mg bd is recommended. This can be individually adjusted with 0.5 mg bd increments to 1 to 2 mg bd. Risperdal is well tolerated by the elderly. Use with caution in patients with renal and liver disease. Not recommended in children aged less than 15 years. **CONTRA-INDICATIONS, WARNINGS, ETC. Contra-indications:** Known hypersensitivity to Risperdal. **Precautions:** Orthostatic hypotension can occur (alpha-blocking effect). Use with caution in patients with known cardiovascular disease. Consider dose reduction if hypotension occurs. For further sedation, give an additional drug (such as a benzodiazepine) rather than increasing the dose of Risperdal. Drugs with dopamine antagonistic properties have been associated with tardive dyskinesia. If signs and symptoms of tardive dyskinesia appear, the discontinuation of all antipsychotic drugs should be considered. Caution should be exercised when treating patients with Parkinson's disease or epilepsy. Patients should be advised of the potential for weight gain. Risperdal may interfere with activities requiring mental alertness. Patients should be advised not to drive or operate machinery until their individual susceptibility is known. **Pregnancy and lactation:** Use during pregnancy only if the benefits outweigh the risks. Women receiving Risperdal should not breast feed. **Interactions:** Use with caution in combination with other centrally acting drugs. Risperdal may antagonise the effect of levodopa and other dopamine agonists. On initiation of carbamazepine or other hepatic enzyme-inducing drugs, the dosage of Risperdal should be re-evaluated and increased if necessary. On discontinuation of such drugs, the dosage of Risperdal should be re-evaluated and decreased if necessary. **Side effects:** Risperdal is generally well tolerated and in many instances it has been difficult to differentiate adverse events from symptoms of the underlying disease. Common adverse events include: insomnia, agitation, anxiety, headache. Less common adverse events include: somnolence, fatigue, dizziness, impaired concentration, constipation, dyspepsia, nausea/vomiting, abdominal pain, blurred vision, priapism, erectile dysfunction, ejaculatory dysfunction, orgasmic dysfunction, urinary incontinence, rhinitis, rash and other allergic reactions. The incidence and severity of extrapyramidal symptoms are significantly less than with haloperidol. However, the following may occur: tremor, rigidity, hypersalivation, bradykinesia, akathisia, acute dystonia. If acute, these symptoms are usually mild and reversible upon dose reduction and/or administration of antiparkinson medication. Rare cases of Neuroleptic Malignant Syndrome have been reported. In such an event, all antipsychotic drugs should be discontinued. Occasionally, orthostatic dizziness, hypotension (including orthostatic), tachycardia (including reflex) and hypertension have been observed. An increase in plasma prolactin concentration can occur which may be associated with galactorrhoea, gynaecomastia and disturbances of the menstrual cycle. Oedema and increased hepatic enzyme levels have been observed. A mild fall in neutrophil and/or thrombocyte count has been reported. Rare cases of water intoxication with hyponatraemia, tardive dyskinesia, body temperature dysregulation and seizures have been reported. **Overdosage:** Reported signs and symptoms include drowsiness and sedation, tachycardia and hypotension, and extrapyramidal symptoms. A prolonged QT interval was reported in a patient with concomitant hypokalaemia who had ingested 360 mg. Establish and maintain a clear airway, and ensure adequate oxygenation and ventilation. Gastric lavage and activated charcoal plus a laxative should be considered. Commence cardiovascular monitoring immediately, including continuous electrocardiographic monitoring to detect possible arrhythmias. There is no specific antidote, so institute appropriate supportive measures. Treat hypotension and circulatory collapse with appropriate measures. In case of severe extrapyramidal symptoms, give anticholinergic medication. Continue close medical supervision and monitoring until the patient recovers. **PHARMACEUTICAL PRECAUTIONS** Tablets: Store below 30°C. Liquid: Store below 30°C, protect from freezing. **LEGAL CATEGORY** POM. **PRESENTATIONS, PACK SIZES, PRODUCT LICENCE NUMBERS & BASIC NHS COSTS** White, oblong tablets containing 1 mg risperidone in packs of 20. PL 0242/0186 £13.45. Pale orange, oblong tablets containing 2 mg risperidone in packs of 60. PL 0242/0187 £79.56. Yellow, oblong tablets containing 3 mg risperidone in packs of 60. PL 0242/0188 £117.00. Green, oblong tablets containing 4 mg risperidone in packs of 60. PL 0242/0189 £154.44. Yellow, circular tablets containing 6 mg risperidone in packs of 28. PL 0242/0317 £109.20. Starter packs containing 6 Risperdal 1 mg tablets are also available £4.15. Clear, colourless solution containing 1 mg risperidone per ml in bottles containing 100 ml. PL 0242/0199 £65.00. **FURTHER INFORMATION IS AVAILABLE FROM THE PRODUCT LICENCE HOLDER:** Janssen-Cilag Ltd, Saunderton, High Wycombe, Buckinghamshire HP14 4HJ. APIVER 140797. **References:** 1. Brecher M, Lemmens P, Van Baelen B. Presented at the Annual Meeting of the American College of Neuropsychiatry, December 9-13, 1996, San Juan, Puerto Rico. 2. Data on file, Janssen-Cilag Ltd. MJE 12/97.

Date of preparation: December 1997.

Campral EC acamprosate

Presentation: Off-white round enteric-coated tablets, containing 333mg acamprosate calcium. Printed on one side with 333. **Properties:** Acamprosate may act by stimulating GABAergic inhibitory neurotransmission and antagonising excitatory amino acids, particularly glutamic acid. **Indication:** Maintenance of abstinence in alcohol dependent patients. It should be combined with counselling. **Dosage and Administration:** *Adults ≥ 60kg:* 6 tablets per day (2 tablets taken three times daily with meals) *Adults < 60kg:* 4 tablets per day (2 tablets in the morning, 1 at noon and 1 at night with meals). Recommended treatment period one year, starting as patient relapses. *Elderly:* Not recommended. *Children:* Not recommended. **Contraindications:** Known hypersensitivity to the drug, renal insufficiency (serum creatinine > 120 micromol/L), severe hepatic failure (Childs-Pugh classification C), pregnancy, lactation. **Precautions and Warnings:** Campral EC does not constitute treatment during the withdrawal period. **Interactions:** None observed in studies with diazepam, disulfiram or imipramine. The concomitant intake of alcohol and acamprosate does not affect the pharmacokinetics of either alcohol or acamprosate. **Side Effects:** Diarrhoea, and less frequently nausea, vomiting and abdominal pain; pruritus. These are usually mild and transient. An occasional maculopapular rash and rare reported. Campral EC should not impair the patient's ability to drive or operate machinery. **Overdose:** Gastric lavage; should hypercalcaemia occur, treat patient for acute hypercalcaemia. **Legal Category:** POM. **Pharmaceutical Precautions:** None. **Package Quantities and Basic NHS Price:** 84 blister packed tablets £24.95. **Marketing Authorisation Number/Holder:** 13466/0001, Lipha SA, Lyon, France. **Date of Preparation:** August 1997. Further information is available on request from Merck Pharmaceuticals, Harrier House, High Street, West Drayton, Middlesex, UB7 7QG. **Date of Preparation:** March 1998.
March 1998.ZZ10104

https://doi.org/10.1017/S0007125000261047 Published online by Cambridge University Press



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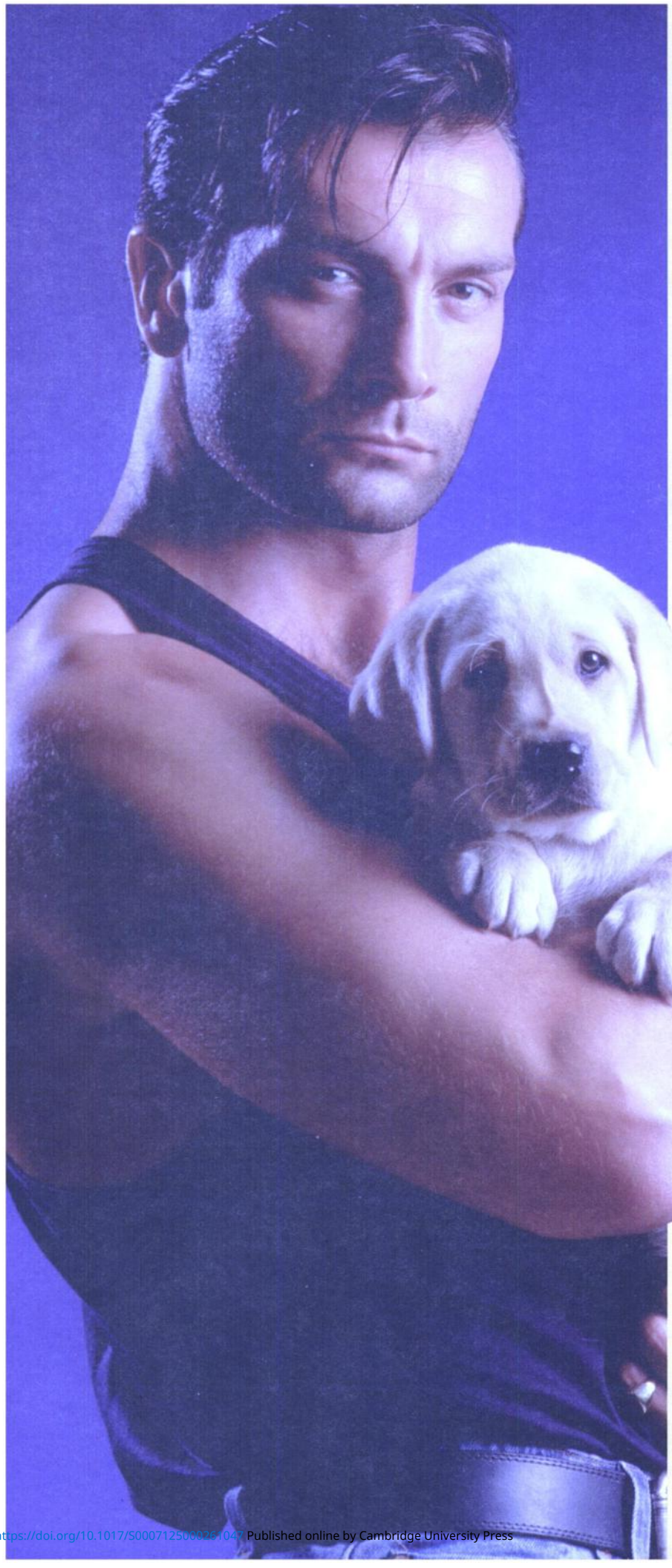

MIRTAZAPINE

ZISPIN 30 mg

The NaSSA

Strong yet gentle in depression

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Add life to living with schizophrenia

Solian is a new benzamide antipsychotic, with the ability to treat both the positive[1] and negative[2] symptoms of schizophrenia.

Solian offers a lower incidence of EPS than standard neuroleptics such as haloperidol,[3] as well as avoiding some of the drawbacks of certain atypicals: it does not require routine cardiovascular[4,5] or haematological[4,6] monitoring and patients gain significantly less weight than those treated with risperidone.[2]

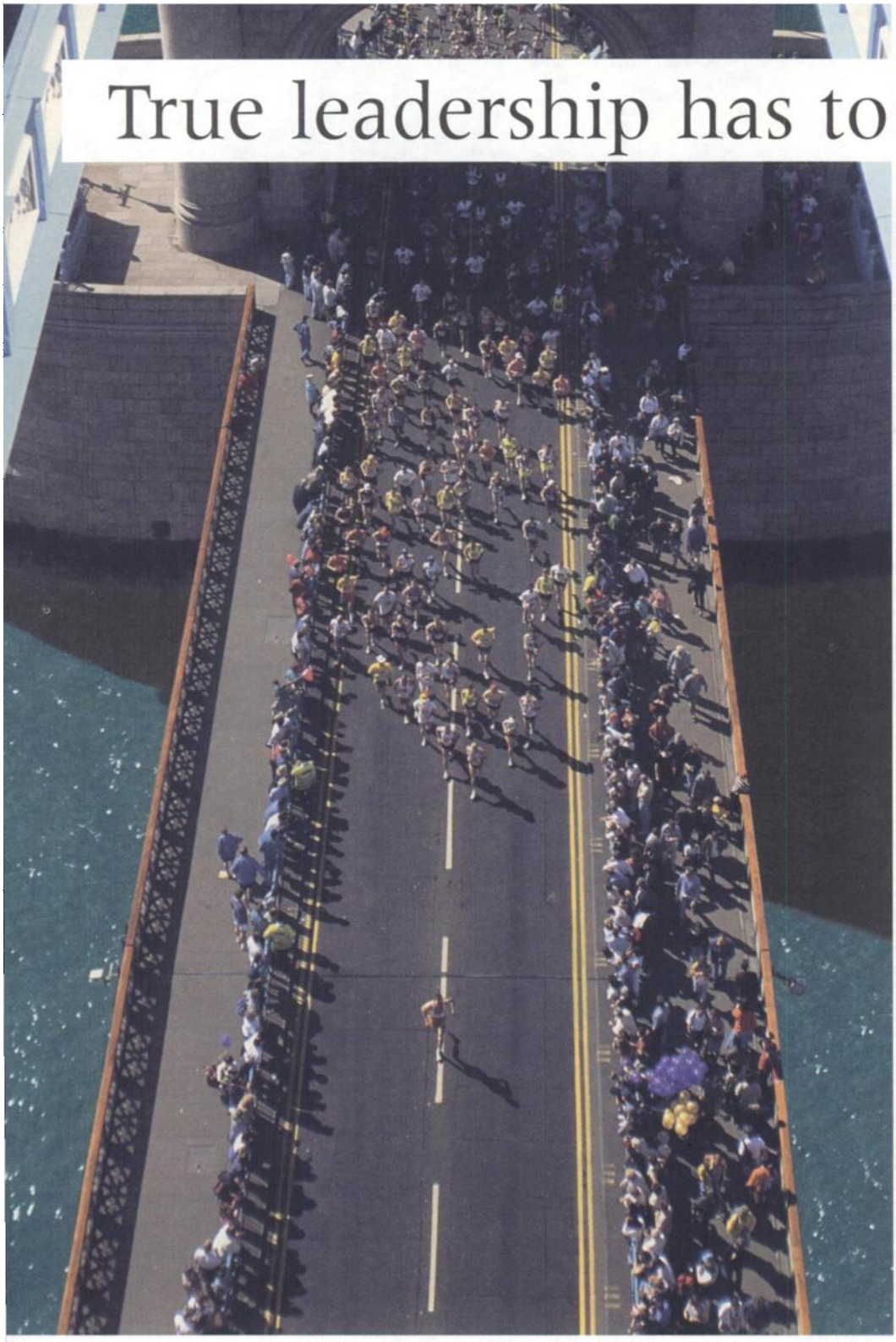
So when patients need the ability to cope with their condition, Solian has the power to treat their positive[1] *and* their negative[2] symptoms whilst still allowing them to do the everyday things that the rest of us take for granted.

Solian®
AMISULPRIDE

Efficacy that patients can live with

Prescribing Information - Solian 200 and Solian 50 ▼ Presentation: Solian 200mg tablets contain 200mg amisulpride and Solian 50mg tablets contain 50mg amisulpride. **Indication:** Acute and chronic schizophrenia in which positive and/or negative symptoms are prominent. **Dosage:** Acute psychotic episodes: 400-800mg/day, increasing up to 1200mg/day according to individual response (dose titration not required), in divided doses. Predominantly negative symptoms: 50-300mg once daily adjusted according to individual response. Elderly: administer with caution due to the risk of hypotension or sedation. Renal insufficiency: reduce dose and consider intermittent therapy. Hepatic insufficiency: no dosage adjustment necessary. Children: contraindicated in children under 15 years (safety not established). **Contraindications:** Hypersensitivity; concomitant prolactin-dependent tumours e.g. pituitary gland prolactinaemias and breast cancer; phaeochromocytoma; children under 15 years; pregnancy; lactation; women of child-bearing potential unless using adequate contraception. **Warning and Precautions:** As with all neuroleptics, neuroleptic malignant syndrome may occur (discontinue Solian). Caution in patients with a history of epilepsy and Parkinson's disease. **Interactions:** Caution in hypotensive medications, and dopamine agonists. **Side Effects:** Insomnia, anxiety, agitation. Less commonly somnolence and GI disorders. In common with other neuroleptics: Solian causes a reversible increase in plasma prolactin levels; Solian may also cause weight gain, acute dystonia, extrapyramidal symptoms, tardive dyskinesia, hypotension and bradycardia; rarely, allergic reactions, seizures and neuroleptic malignant syndrome have been reported. **Basic NHS Cost:** Blister packs of: 200mg x 60 tablets - £60.00; 200mg x 90 tablets - £90.00; 50mg x 60 tablets - £16.45; 50mg x 90 tablets - £24.69. **Legal Category:** POM. **Product Licence Numbers:** Solian 200 - PL 15819/0002, Solian 50 - PL 15819/0001. **Product Licence Holder:** Lorex Synthélabo UK and Ireland Ltd, Foundation Park, Roxborough Way, Maidenhead, Berks, SL6 3UD. **References:** 1. Freeman HL. Int Clin Psychopharmacol 1997;12(Suppl 2):S11-S17. 2. Möller HJ. 6th World Congress of Biological Psychiatry, Nice, France, June 22-27 1997. 3. Coukell AJ, Spencer CM, Benfield P. CNS Drugs (Adis) 1996 Sep 6 (3):237-256. 4. Solian SPC. Lorex Synthélabo. 5. Sertindole SPC. Lundbeck Ltd. 6. Clozapine SPC.

SYNTHELABO
CNS DIVISION



True leadership has to be earned.

ASSOCIATED ANXIETY

Prozac has a proven record of efficacy in depression,[1,2,3] with a confirmed indication in depression with or without associated anxiety symptoms.[4]

A possible reason why Prozac has earned its status around the world.

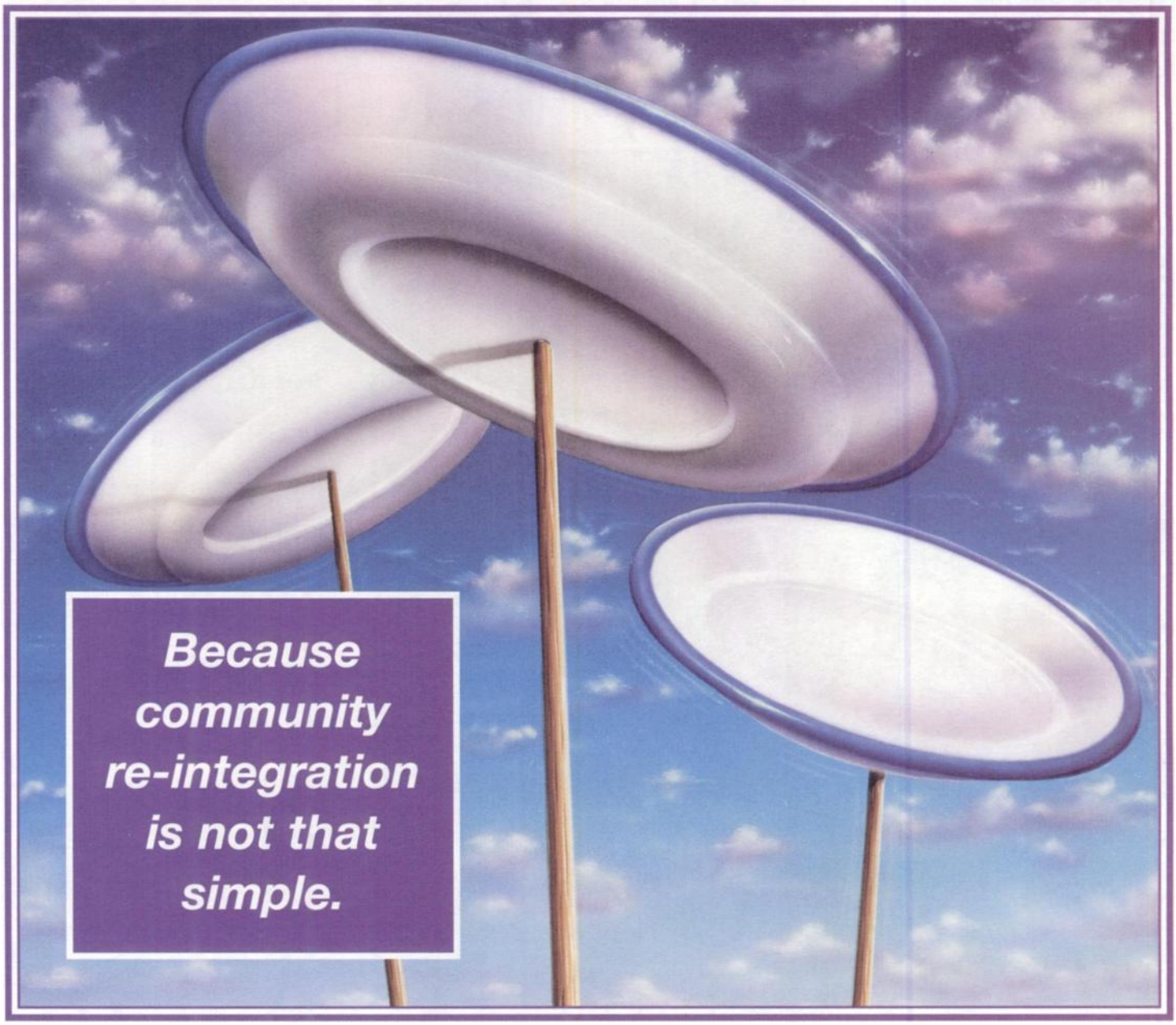
PROZAC fluoxetine

The World's No.1 prescribed antidepressant brand.[1]

'PROZAC' ABBREVIATED PRESCRIBING INFORMATION (FLUOXETINE HYDROCHLORIDE)
Presentation Capsules containing 20mg or 60mg fluoxetine, as the hydrochloride. Liquid containing 20mg fluoxetine, as the hydrochloride, per 5ml syrup. USES Depression: TREATMENT OF THE SYMPTOMS OF DEPRESSIVE ILLNESS, WITH OR WITHOUT ASSOCIATED ANXIETY SYMPTOMS. *Obsessive-compulsive disorder. Bulimia nervosa:* For the reduction of binge-eating and purging activity. **Dosage and Administration** (For full information, see data sheet.) For oral administration to adults only. *Depression, with or without associated anxiety symptoms - adults and the elderly:* A dose of 20mg/day is recommended. *Obsessive-compulsive disorder:* 20mg/day to 60mg/day. A dose of 20mg/day is recommended as the initial dose. *Bulimia - adults and the elderly:* A dose of 60mg/day is recommended. Because of the long elimination half-lives of the parent drug (1-3 days after acute administration; may be prolonged to 4-6 days after chronic administration) and its major metabolite (average 9.3 days), active drug substance will persist in the body for several weeks after dosing is stopped. The capsule and liquid dosage forms are bioequivalent. *Children:* Not recommended. *Patients with renal and/or hepatic dysfunction:* See 'Contra-indications' and 'Precautions' sections. **Contra-indications** Hypersensitivity to fluoxetine. Prozac should not be administered to patients with severe renal failure (GFR <10ml/min). *Usage in nursing mothers:* Prozac should not be prescribed to nursing mothers. *Monoamine oxidase inhibitors:* At initiation of therapy with an MAOI. Serious, sometimes fatal reactions (including hyperthermia, rigidity, myoclonus, autonomic instability and mental status changes that include extreme agitation, progressing to delirium and coma) have been reported with concomitant use or when fluoxetine had been recently discontinued and an MAOI started. Some cases presented with features resembling neuroleptic malignant syndrome. **Warnings** *Rash and allergic reactions:* Angioneurotic oedema, urticaria and other allergic reactions have been reported. Upon appearance of rash, or of other allergic phenomena for which an alternative aetiology cannot be identified, Prozac should be discontinued. *Pregnancy:* Use of Prozac should be avoided unless there is no safer alternative. **Precautions** Prozac should be discontinued in any patient who develops seizures. Prozac should be avoided in patients with unstable epilepsy; patients with controlled epilepsy should be carefully monitored. There have been rare reports of prolonged seizures in patients on fluoxetine receiving ECT treatment. A lower dose of Prozac, eg, alternate day dosing, is recommended in patients with significant hepatic dysfunction or mild to moderate renal failure (GFR 10-50ml/min). Caution is advisable when Prozac is used in patients with acute cardiac disease. Prozac may cause weight loss which may be undesirable in underweight depressed patients. In diabetics, fluoxetine may alter glycaemic control. There have been reports of abnormal bleeding in several patients, but causal relationship to fluoxetine and clinical importance are unclear. *Drug interactions:* cytochrome P450IID6 isoenzyme system, concomitant therapy with other drugs also metabolised by this system, and which have a narrow therapeutic index (eg, carbamazepine, tricyclic antidepressants), should be initiated at or adjusted to the low end of their dose range. Greater than 2-fold increases of previously stable plasma levels of cyclic antidepressants have been observed when Prozac has been administered in combination. Agitation, restlessness and gastro-intestinal symptoms have been reported in a small number of patients receiving fluoxetine in combination with tryptophan. Patients on stable phenytoin doses have developed elevated plasma concentrations and clinical phenytoin toxicity after starting fluoxetine. *For further information, see data sheet.* **Adverse Effects** Asthenia, fever, nausea, diarrhoea, dry mouth, appetite loss, dyspepsia, vomiting, rarely abnormal LFTs, headache, nervousness, insomnia, drowsiness, anxiety, tremor, dizziness, fatigue, decreased libido, seizures, hypomania or mania, dyskinesia, movement disorders, neuroleptic malignant syndrome-like events, pharyngitis, dyspnoea, pulmonary events (including inflammatory processes and/or fibrosis), rash, urticaria, vasculitis, excessive sweating, arthralgia, myalgia, serum sickness, anaphylactoid reactions, hair loss, sexual dysfunction. The following have been reported in association with fluoxetine but no causal relationship has been established: aplastic anaemia, cerebral vascular accident, confusion, ecchymoses, eosinophilic pneumonia, gastro-intestinal haemorrhage, hyperprolactinaemia, immune-related Hyponatraemia (including serum sodium below 110mmol/l) has been rarely reported. This appears to be reversible upon discontinuation. **Overdosage** On the evidence available, fluoxetine has a wide margin of safety in overdose. Since introduction, reports of death, attributed to overdosage of fluoxetine alone, have been extremely rare. One patient who reportedly took 3000mg of fluoxetine experienced 2 grand mal seizures that remitted spontaneously. **Legal Category** POM **Product Licence Numbers** 0006/0195 0006/0198 0006/0272 **Basic NHS Cost** £20.77per pack of 30 capsules (20mg). £67.85 per pack of 98 capsules (20mg). £62.31 per pack of 30 capsules (60mg). £19.39 per 70ml bottle. **Date of Preparation or Last Review** October 1996. **Full Prescribing Information is Available From** Dista Products Limited, Dextra Court, Chapel Hill, Basingstoke, Hampshire, RG21 5SY. Telephone: Basingstoke (01256) 52011
'PROZAC' is a Dista trademark

References:1. Data on file, Dista Products Ltd. 2. Tignol J. *J Clin Psychopharm* 1993; **13** (6, suppl. 2): 18S-22S. 3. Bennie EH, Mullin JM, Martindale JJ. *J Clin Psychiatry* 1995; **56**: 229-237. 4. Prozac Data Sheet 24M.

Date of preparation: May 1997 PZ 906



Antipsychotic Efficacy for First-line Use

Zyprexa Olanzapine

Making Community Re-integration the Goal

ABBREVIATED PRESCRIBING INFORMATION: **Presentation:** Coated tablets containing 5mg, 7.5mg or 10mg of olanzapine. The tablets also contain lactose. **Uses:** Schizophrenia, both as initial therapy and for maintenance of response. **Further Information:** In studies of patients with schizophrenia and associated depressive symptoms, mood score improved significantly more with olanzapine than with haloperidol. **Pharmacodynamics:** Olanzapine was associated with significantly greater improvements in both negative and positive schizophrenic symptoms than placebo or comparator in most studies. **Dosage and Administration:** 10mg/day orally, as a single dose without regard to meals. Dosage may subsequently be adjusted within the range of 5-20mg daily. An increase to a dose greater than the routine therapeutic dose of 10mg/day is recommended only after clinical assessment. *Children:* Not recommended under 18 years of age. *The elderly:* A lower starting dose (5mg/day) is not routinely indicated but should be considered when clinical factors warrant. *Hepatic and/or renal impairment:* A lower starting dose (5mg) may be considered. When more than one factor is present which might result in slower metabolism (female gender, elderly age, non-smoking status), consideration should be given to decreasing the starting dose. Dose escalation should be conservative in such patients. **Contra-indications:** Known hypersensitivity to any ingredient of the product. Known risk for narrow-angle glaucoma. **Warnings and Special Precautions:** Caution in patients with prostatic hypertrophy, or paralytic ileus and related conditions. Caution in patients with elevated ALT and/or AST, signs and symptoms of hepatic impairment, pre-existing conditions associated with limited hepatic functional reserve, and in patients who are being treated with potentially hepatotoxic drugs. As with other neuroleptic drugs, caution in patients with low leucocyte and/or neutrophil counts for any reason, a history of drug-induced bone marrow depression/toxicity, bone marrow depression caused by concomitant illness, radiation therapy or chemotherapy and in patients with hypereosinophilic conditions or with myeloproliferative disease. Thirty-two patients with clozapine-related neutropenia or agranulocytosis histories received olanzapine without decreases in baseline neutrophil counts. Although, in clinical trials, there were no reported cases of NMS in patients receiving olanzapine, if such an event occurs, or if there is unexplained high fever, all antipsychotic drugs, including olanzapine, must be discontinued. Caution in patients who have a history of seizures or have conditions associated with seizures. If signs or symptoms of tardive dyskinesia appear a dose reduction or drug discontinuation should be considered. Caution when taken in combination with other centrally acting drugs and alcohol. Olanzapine may antagonise the effects of direct and elderly. However, blood pressure should be measured periodically in patients over 65 years, as with other antipsychotics. As with other antipsychotics, caution when prescribed with drugs known to increase QTc interval, especially in the elderly. In clinical trials, olanzapine was not associated with a persistent increase in absolute QT intervals. **Interactions:** Metabolism may be induced by concomitant smoking or carbamazepine therapy. **Pregnancy and Lactation:** Olanzapine had no teratogenic effects in animals. Because human experience is limited, olanzapine should be used in pregnancy only if the potential benefit justifies the potential risk to the foetus. Olanzapine was excreted in the milk of treated rats but it is not known if it is excreted in human milk. Patients should be advised not to breast feed an infant if they are taking olanzapine. **Driving, etc:** Because olanzapine may cause somnolence, patients should be cautioned about operating hazardous machinery, including motor vehicles. **Undesirable Effects:** The only frequent (>10%) undesirable effects associated with the use of olanzapine in clinical trials were somnolence and weight gain. Occasional undesirable effects included dizziness, increased appetite, peripheral oedema, orthostatic hypotension, and mild, transient anticholinergic effects, including constipation and dry mouth. Transient, asymptomatic elevations of hepatic transaminases, ALT, AST have been seen occasionally. Olanzapine-treated patients had a lower incidence of parkinsonism, akathisia and dystonia in trials compared with titrated doses of haloperidol. Photosensitivity reaction or high creatinine phosphokinase were reported rarely. Plasma prolactin levels were sometimes elevated, but associated clinical manifestations were rare. Asymptomatic haematological variations were occasionally seen in trials. *For further information see summary of product characteristics.* **Legal Category:** POM. **Marketing Authorisation Numbers:** EU/1/96/022/004 EU/1/96/022/006 EU/1/96/022/008 EU/1/96/022/009 EU/1/96/022/010. **Basic NHS Cost:** £52.73 per pack of 28 x 5mg tablets. £105.47 per pack of 28 x 10mg tablets. £158.20 perpack of 56 x 7.5mg tablets. £210.93 per pack of 56 x 10mg tablets. **Date of Preparation or Last Review:** April 1997. **Full Prescribing Information is Available From:** Eli Lilly and Company Limited, Dextra Court, Chapel Hill, Basingstoke, Hampshire RG21 5SY. Telephone: Basingstoke (01256) 315000.

Agenda

18.00 PM **Cocktail Reception**

18.15 PM **Welcome & Introduction**
Alistair Burns, MD, Chairman
Manchester
United Kingdom

18.20 **Pediatric OCD:**
Characteristics and Treatment
John March, MD
Durham, North Carolina
USA

18.40 **The Prevalence and Treatment of Comorbid MDD and OCD**
Rudolf Hoehn-Saric, MD
Baltimore, Maryland
USA

19.00 **Epidemiologic Perspectives:**
Comorbidity of Panic Disorder and Depression
Borwin Bandelow, MD
Göttingen
Germany

19.20 **Effective and Comprehensive Management of Patients with Panic Disorder**
Christer Allgulander, MD
Huddinge
Sweden

19.40 **Late Life Depression:**
Improving Cognition, Anxiety, Energy, and Sleep
Bernard Groulx, MD
Ste-Anne de Bellevue, Quebec
Canada

20.00 **Question & Answer Session**
Faculty Panel

20.15 **Dinner Buffet**

20.45 **Adjournment**

July 13, 1998

Depression, Panic, and OCD:

Improving Patient Management *Through the Life Cycle*

Argyll Suite
Moat House Hotel

Glasgow
Scotland

XXIst Congress of the
Collegium Internationale
Neuro-Psychopharmacologicum

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Presentation: White to off-white tablets each containing modafinil 100 mg. **Indication:** Narcolepsy. **Dosage:** *Adults:* 200-400 mg daily either as two divided doses in the morning and at noon or as a single morning dose according to response. *Elderly:* Treatment should start at 100 mg daily which may be increased subsequently to the maximum adult daily dose in the absence of renal or hepatic impairment. *Severe renal or hepatic impairment:* Reduce dose by half (100-200 mg daily). *Children:* See contra-indications. **Contra-indications:** Pregnancy, lactation, use in children, moderate to severe hypertension, arrhythmia, hypersensitivity to modafinil or any excipients used in Provigil. **Warnings and precautions:** Patients with major anxiety should only receive Provigil treatment in a specialist unit. Sexually active women of child-bearing potential should be established on a contraceptive programme before starting treatment. Blood pressure and heart rate should be monitored in hypertensive patients. Provigil is not recommended in patients with a history of left ventricular hypertrophy or ischaemic ECG changes, chest pain, arrhythmia or other clinically significant manifestations of mitral valve prolapse in association with CNS stimulant use. Studies of modafinil have demonstrated a low potential for dependence although the possibility of this occurring with long-term use cannot be entirely excluded. **Drug interactions:** Induction of cytochrome P-450 isoenzymes has been observed *in vitro*. Effectiveness of oral no clinically relevant interaction was seen in a single dose interaction study of Provigil and clomipramine. However, patients receiving such medication should be carefully monitored. Care should be observed with co-administration of anti-convulsant drugs. **Side effects:** Nervousness, excitation, aggressive tendencies, insomnia, personality disorder, anorexia, headache, CNS stimulation, euphoria, abdominal pain, dry mouth, palpitation, tachycardia, hypertension and tremor have been reported. Nausea and gastric discomfort may occur and may improve when tablets are taken with meals. Pruritic skin rashes have been observed occasionally. Buccofacial dyskinesia has been reported very rarely. A dose related increase in alkaline phosphatase has been observed. **Basic NHS cost:** Packs of 30 blister packed 100 mg tablets: £60.00. **Marketing authorisation number:** 16260/0001. **Marketing authorisation holder:** Cephalon UK Ltd., 11/13 Frederick Sanger Road, Surrey Research Park, Guildford, GU2 5YD. **Legal category:** POM. **Date of preparation:** January 1998. Provigil and Cephalon are registered trademarks. **References: 1.** Mitler MM. Sleep 1994; **17**: S103-S106. **2.** Data on file, Cephalon [3]. **3.** Lin JS *et al. Proc Natl Acad Sci USA* 1996; **93** (24): 14128-14133. **4.** Simon P *et al.* Eur Neuropsychopharmacol 1995; **5**: 509-514.

WAKE UP LITTLE SUZIE, WAKE UP

Excessive sleepiness associated with narcolepsy frequently has a disastrous effect on patients' lives, by impairing their physical, social and emotional well being. Unfortunately, treatment with amphetamines is often associated with a high incidence of unpleasant side effects, which limit their overall benefit.[1]

Now Provigil (modafinil) – a novel wake promoting agent – offers new advantages in narcolepsy. The clinical efficacy of Provigil has been demonstrated in large controlled clinical studies. In one study,[2] one in five people with severe narcolepsy reached normal levels of daytime wakefulness while receiving Provigil.

Provigil selectively activates the hypothalamus[3] and differs greatly from amphetamines in its pharmacology.[4] Consequently the incidence of amphetamine like side effects is very low.

PROVIGIL®▼
MODAFINIL

DUTONIN™▼ Abbreviated Prescribing Information PRESENTATION: Tablets containing 50mg, 100mg and 200mg nefazodone hydrochloride. INDICATIONS: Symptomatic treatment of all types of depressive illness, including depressive syndromes accompanied by anxiety or sleep disturbances. DOSAGE: Usual therapeutic dose 200mg twice daily. Range – 100mg - 600mg daily, see Summary of Product Characteristics. Elderly: Usual therapeutic dose 50 - 200mg twice daily. Renal and Hepatic Impairment: Lower end of dose range. Children: Not recommended below the age of 18 years. CONTRA-INDICATIONS: Hypersensitivity to nefazodone hydrochloride, tablet excipients or phenylpiperazine antidepressants. WARNINGS/ PRECAUTIONS: Hepatic or renal impairment. Patients at high risk of self harm should be kept under close supervision during initial treatment phase. Modest decrease in some psychomotor function tests but no impairment of cognitive function. Not recommended in pregnancy and lactation. Use with caution in epilepsy, history of mania/hypomania, recent M.I., unstable heart disease. No clinical studies available on concurrent use of ECT and nefazodone. DRUG INTERACTIONS: Caution is advised when combining with other CNS medication, digoxin, products metabolised by Cytochrome $P_{450}III_{A4}$; see Summary of Product Characteristics. SIDE EFFECTS: Most frequently asthenia, dry mouth, nausea, constipation, somnolence, light-headedness and dizziness; see Summary of Product Characteristics. OVERDOSAGE: There is no specific antidote for nefazodone. Gastric lavage recommended for suspected overdose. Treatment should be symptomatic and supportive in the case of hypotension or excessive sedation. PRODUCT LICENCE NUMBERS: Dutonin Tablets 50mg PL 11184/0027; Dutonin Tablets 100mg PL 11184/0028; Dutonin Tablets 200mg PL 11184/0029, PRODUCT LICENCE HOLDER: Bristol-Myers Squibb Pharmaceuticals Ltd. BASIC NHS PRICE: Treatment Initiation Pack containing 50mg tablets 14, 100mg tablets 14, 200mg tablets 28 – £16.80; 100mg tablets 56 – £16.80; 200mg tablets 56 – £16.80. LEGAL CATEGORY: POM. Further information from: Medical Information, Bristol-Myers Squibb House, 141-149 Staines Road, Hounslow, Middlesex, TW3 3JA. Telephone: 0181-754-3740. Date of preparation: July 1997.

REFERENCES: 1. Armitage R. Journal of Psychopharmacology 1996; 10(suppl1): 22-25. 2. Sharpley AL *et al.* Psychopharmacology 1996; 126: 50-54. 3. Armitage R *et al.* J Clin Psychopharmacol 1997; 17(3): 161-168. 4. Armitage R *et al.* Presented at the European College of Neuropsychopharmacology (ECNP), 30 September - 4 October 1995, Venice, Italy. 5. Fontaine R *et al.* J Clin Psychiatry 1994; 55(6): 234-241. 6. Gillin JC *et al.* J Clin Psychiatry 1997; 58: 185-192.

Bristol-Myers Squibb Pharmaceuticals Limited

Waking up early should be her decision, not her problem.

It's not only depression that wakes patients up early. Sleep can also be disturbed by many SSRIs.[1-4]

Dutonin is an excellent choice. Not only does Dutonin effectively relieve depression,[5] it also normalises sleep patterns.[3,4,6]

Moreover, Dutonin lifts anxiety symptoms within the first week of treatment.[5]

Waking up early should always be your patient's choice, not their problem.

Makes the difference in depression

DUTONIN™▼

NEFAZODONE

https://doi.org/10.1017/S0007125000261047 Published online by Cambridge University Press

Mum has Alzheimer's

- The only once daily treatment in Alzheimer's disease
- Effective in mild to moderately severe stages [1-4]
- Improves cognitive symptoms and maintains global function [1-4]
- Well tolerated 5mg and 10mg once daily doses. [1-5]

but she knew I was calling today

Aricept®

donepezil hydrochloride

Once daily in Alzheimer's

BRIEF PRESCRIBING INFORMATION

ARICEPT® (donepezil hydrochloride). Please refer to the SmPC before prescribing ARICEPT 5mg or ARICEPT 10mg. **Indication:** Symptomatic treatment of mild to moderately severe Alzheimer's dementia. **Dose and administration:** ***Adults/elderly;*** 5mg daily which may be increased to 10mg once daily after at least one month. No dose adjustment necessary for patients with renal or mild-moderate hepatic impairment. ***Children;*** Not recommended. **Contra-Indications:** Pregnancy. Hypersensitivity to donepezil, piperidine derivatives or any excipients used in ARICEPT. **Lactation:** Excretion into breast milk unknown. Women on donepezil should not breast feed. **Warnings and Precautions:** Initiation and supervision by a physician with experience of Alzheimer's dementia. A caregiver should be available to monitor compliance. Regular monitoring to ensure continued therapeutic benefit, consider discontinuation when evidence of a therapeutic effect ceases. Exaggeration of succinylcholine-type muscle relaxation. Avoid concurrent use of may be particularly important with "sick sinus syndrome" and supraventricular conduction conditions. Careful monitoring of patients at risk of ulcer disease including those receiving NSAIDs. Cholinomimetics may cause bladder outflow obstruction. Seizures occur in Alzheimer's disease and cholinomimetics have the potential to cause seizures. Care in patients suffering asthma and obstructive pulmonary disease. As with all Alzheimer's patients, routine evaluation of ability to drive/operate machinery. **Drug Interactions:** Experience of use with concomitant medications is limited, consider possibility of as yet unknown interactions. Interaction possible with inhibitors or inducers of Cytochrome P450: use such combinations with care. Possible synergistic activity with succinylcholine-type muscle relaxants, beta-blockers, cholinergic or anticholinergic agents. **Side effects:** Most commonly diarrhoea, muscle cramps, fatigue, nausea, vomiting and insomnia. Other common effects in clinical trials (≥5% and ≥placebo) headache, pain, accident, common cold, abdominal **Presentation and basic NHS cost:** Blister packed in strips of 14. ARICEPT 5mg; white, film coated tablets marked 5 and ARICEPT, packs of 28 £68.32. ARICEPT 10mg; yellow, film coated tablets marked 10 and ARICEPT, packs of 28 £95.76. **Marketing authorisation numbers:** ARICEPT 5 mg; PL 10555/0006. ARICEPT 10mg; PL 10555/0007. **Marketing authorisation holder:** Eisai Ltd. **Further information from/Marketed by:** Eisai Ltd, Hammersmith International Centre, 3 Shortlands, London, W6 8EE and Pfizer Ltd, Sandwich, Kent, CT13 9NJ. **Legal category:** POM **Date of preparation:** January 1998. **References: 1.** Rogers SL et al. Neurology 1998; 50: 136-145. **2.** Study 301 (accepted for publication, Arch Int Med). **3.** Rogers SL, Friedhoff LT. Eur Neuropsychopharmacol 1998; 8 (1): 67-75. **4.** Rogers SL et al. Dementia 1996; 7: 293-303. **5.** Rogers SL & Friedhoff LT. Eur Neuropsychopharmacol 1997; 7 (suppl. 2): S251.

Eisai Pfizer

TOPAMAX®
topiramate

At the end of the day, it works.

Adjunctive treatment for partial seizures with or without secondary generalisation

TOPAMAX Abbreviated Prescribing Information
Please read the data sheet before prescribing

Presentation: Tablets each imprinted "TOP" on one side and strength on the other containing 25mg (white), 50mg (light yellow), 100mg (yellow), and 200mg (salmon) topiramate. **Uses:** Adjunctive therapy of partial seizures, with or without secondarily generalised seizures, in patients inadequately controlled on conventional first line antiepileptic drugs. **Dosage and Administration:** Adults and Elderly: Oral administration. Usual dose: 200mg - 400mg/day in two divided doses. Maximum recommended dose: 800mg/day. Initiate therapy at 50mg bd then titrate to an effective dose. See data sheet for titration. Do not break tablets. It is not necessary to monitor topiramate plasma concentrations. Patients with renal disease/haemodialysis may require a modified titration schedule. (See data sheet). Children: Not recommended **Contra-indications:** Hypersensitivity to any component of the product. **Precautions and Warnings:** Withdraw all antiepileptic drugs gradually. Maintain adequate hydration to reduce risk of nephrolithiasis (especially increased in those with a predisposition). Drowsiness likely. TOPAMAX may be more sedating than other antiepileptic drugs therefore caution in patients driving or operating machinery, particularly until patients' experience with the drug is established. Do not use in pregnancy unless potential benefit outweighs risk to foetus. Women of child bearing potential should use adequate contraception. Do not use if breastfeeding. **Interactions:** Other Antiepileptic Drugs: No clinically significant effect except in some patients on phenytoin where phenytoin plasma concentrations may increase or decrease. Monitor phenytoin plasma concentrations on sodium valproate addition or withdrawal. Digoxin: A decrease in serum digoxin occurs. Monitor serum digoxin on addition or withdrawal of TOPAMAX. Oral Contraceptives: Should contain not less than 50µg of oestrogen. Ask patients to report any change in bleeding patterns. Others: Avoid agents predisposing to nephrolithiasis. **Side Effects:** In 5% or more: ataxia, impaired concentration, confusion, dizziness, fatigue, paraesthesia, somnolence and abnormal thinking. May cause agitation and emotional lability (which may manifest as abnormal behaviour) and depression. Less commonly: amnesia, anorexia, aphasia, diplopia, nausea, nystagmus, speech disorder, taste perversion, abnormal vision and weight decrease. Increased risk of nephrolithiasis. Venous thromboembolic events reported - causal association not established. **Overdosage:** If ingestion recent, empty stomach. Activated charcoal not recommended. Supportive treatment as appropriate. Haemodialysis is effective in removing topiramate. **Pharmaceutical Precautions:** Store in a dry place at or below 25°C. **Legal Category:** POM **Package Quantities and Prices:** Bottles of 60 tablets. 25mg (PL0242/0301) = £22.02; 50mg (PL0242/0302) = £36.17; 100mg (PL0242/0303) = £64.80; 200mg (PL0242/0304) = £125.83.

Product Licence Holder: JANSSEN-CILAG LIMITED, SAUNDERTON, HIGH WYCOMBE, BUCKINGHAMSHIRE HP14 4HJ. API VER 210397.

Further information is available on request from the Marketing Authorisation Holder: Janssen-Cilag Limited, Saunderton, High Wycombe, Buckinghamshire HP14 4HJ.

ADVANCE DIARY DATES

Thinking about management issues in schizophrenia?

As part of a comprehensive programme of initiatives open to psychiatrists, CPNs and pharmacists, we are organising a series of one day multi-disciplinary workshops under the general heading **"Therapy Management"**.

Presentations and discussion groups will focus on the following:

- Factors influencing concordance
- Wider therapeutic options in the management of schizophrenia

Meeting Dates

20 May	Zeneca HQ, Cheshire	18 June	Birmingham
21 May	Southampton	18 June	Essex
22 May	Aylesbury	22 June	Bristol
27 May	London	23 June	Cardiff
8 June	Newcastle	24 June	Wembley
10 June	Cambridge	26 June	Totnes
17 June	Wigan	1 July	Belfast
17 June	Ashford	3 July	Leeds
17 June	Glasgow		

For more information on these multi-disciplinary workshops please call Sally Heap at Zeneca on 01625 712412.

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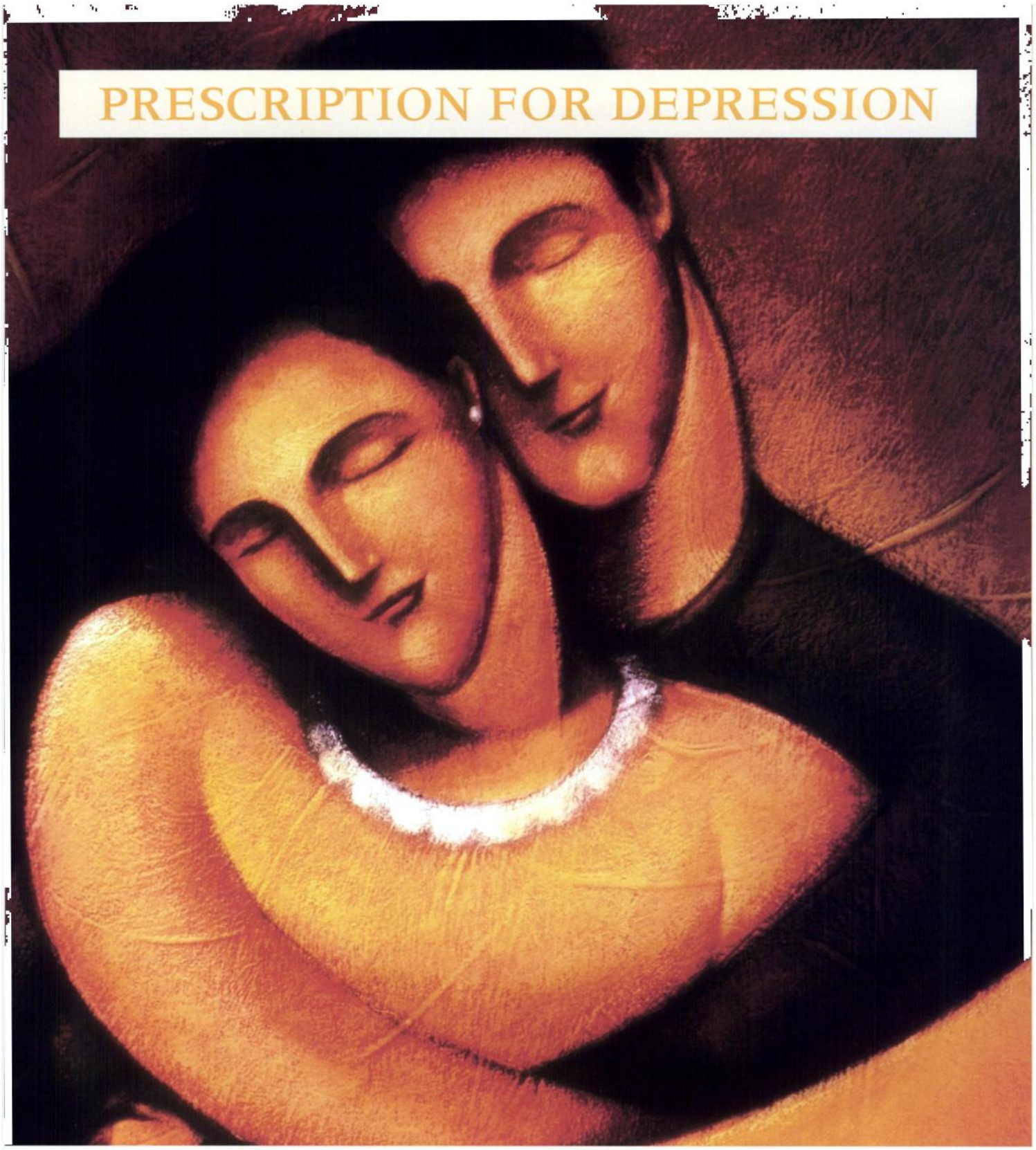
ZENECA

THINKING AHEAD IN PSYCHIATRY



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PRESCRIBING INFORMATION

Presentation: 'Seroxat' Tablets, PL 10592/0001-2, each containing either 20 or 30 mg paroxetine as the hydrochloride. 30 (OP) 20 mg tablets, £20.77; 30 (OP) 30 mg tablets, £31.16. 'Seroxat' Liquid, PL 10592/0092, containing 20 mg paroxetine as the hydrochloride per 10 ml. 150 ml (OP), £20.77.

Indications: Treatment of symptoms of depressive illness of all types including depression accompanied by anxiety. Following satisfactory response, continuation is effective in preventing relapse. Treatment of symptoms and prevention of relapse of obsessive compulsive disorder (OCD). Treatment of symptoms and prevention of relapse of panic disorder with or without agoraphobia.

Dosage: *Adults: Depression:* 20 mg a day. Review response within two to three weeks and if necessary increase dose in 10 mg increments to a maximum of 50 mg according to response.

Obsessive compulsive disorder: 40 mg a day. Patients should be given 20 mg a day initially and the dose increased weekly in 10 mg increments. Some patients may benefit from a maximum dose of 60 mg a day.

Panic disorder: 40 mg a day. Patients should be given 10 mg a day initially and the dose increased weekly in 10 mg increments. Some patients may benefit from a maximum dose of 50 mg a day.

Give orally once a day in the morning with food. The tablets should not be chewed. Continue treatment for a sufficient period, which may be several months for depression or longer for OCD and panic disorder. As with many psychoactive medications abrupt discontinuation should be avoided – see **Adverse reactions.**

Elderly: Dosing should commence at the adult starting dose and may be increased in weekly 10 mg increments up to a maximum of 40 mg a day according to response.

Children: Not recommended.

Severe renal impairment (creatinine clearance <30 ml/min) or severe hepatic impairment: 20 mg a day. Restrict incremental dosage if required to lower end of range.

Contra-indication: Hypersensitivity to paroxetine.

Precautions: History of mania. Cardiac conditions: caution. Caution in patients with epilepsy; stop treatment if seizures develop. Driving and operating machinery.

Drug interactions: Do not use with or within two weeks after MAO inhibitors; leave a two-week gap before starting MAO inhibitor treatment. Possibility of interaction with tryptophan. Great caution with warfarin and other oral anticoagulants. Use lower doses if given with drug metabolising enzyme inhibitors; adjust dosage if necessary with drug metabolising enzyme inducers. Alcohol is not advised. Use lithium with caution and monitor lithium levels. Increased adverse effects with phenytoin; similar possibility with other anticonvulsants.

Pregnancy and lactation: Use only if potential benefit outweighs possible risk.

Adverse reactions: In controlled trials most commonly nausea, somnolence, sweating, tremor, asthenia, dry mouth, insomnia, sexual dysfunction (including impotence and ejaculation disorders), dizziness, constipation and decreased appetite.

Also spontaneous reports of dizziness, vomiting, diarrhoea, restlessness, hallucinations, hypomania, rash including urticaria with pruritus or angioedema, and symptoms suggestive of postural hypotension. Extrapyramidal reactions reported infrequently; usually reversible abnormalities of liver function tests and hyponatraemia described rarely. Symptoms including dizziness, sensory disturbance, anxiety, sleep disturbances, agitation, tremor, nausea, sweating and confusion have been reported following abrupt discontinuation of 'Seroxat'. It is recommended that when antidepressant treatment is no longer required, gradual discontinuation by dose-tapering or alternate day dosing be considered.

Overdosage: Margin of safety from available data is wide. Symptoms include nausea, vomiting, tremor, dilated pupils, dry mouth, irritability, sweating and somnolence. No specific antidote. General treatment as for overdosage with any antidepressant. Early use of activated charcoal suggested.

Legal category: POM. 16.2.98

SmithKline Beecham Pharmaceuticals

Welwyn Garden City, Hertfordshire AL7 1EY.

SEROXAT

Rebuilding the lives of anxious depressed patients

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