THE PRO CHOICE IN ARTHRITIS

It is commonly agreed that prostaglandins confer upon the gastric mucosa a resistance to erosion and damage.[1] This so called mucoprotection is compromised by the use of antiarthritic drugs which are prostaglandin synthetase inhibitors. It has been suggested that this is of great relevance to the observed level of gastric side-effects observed with this class of drug.

Lederfen in the stomach has little or no effect on the mucoprotection conferred by the endogenous prostaglandins.[2] Simply, unmetabolised Lederfen is not a prostaglandin synthetase inhibitor.

The major metabolites of Lederfen are however most potent PSIs, shown by in vitro studies to be twice as potent as indomethacin.[2] Thus, once Lederfen has been absorbed – avoiding prostaglandin-associated gastric compromise – a powerful antiarthritic effect is observed. Lederfen has been found to be at least equal in clinical effectiveness to reference standards including naproxen,[3] indomethacin,[4] and aspirin.[5] But with significantly fewer side-effects.

Power without doubt. The pro-choice in arthritis.

Lederfen

fenbufen TABLETS

the pro-drug way to control pain, stiffness and side-effects

Indications: Rheumatoid arthritis, osteoarthritis, ankylosing spondylitis. **Dosage: Adults:** One tablet in the morning and two at night. LEDERFEN is not a gastric irritant and is converted, following absorption into long-acting active metabolites. Consequently LEDERFEN is extremely unlikely to cause gastro-intestinal bleeding or ulceration. Furthermore, LEDERFEN may be used without dosage modification in elderly patients in whom mild to moderate renal impairment commonly occurs. **Contra-indications:** Hypersensitivity to propionic acid anti-inflammatory drugs or aspirin. **Precautions:** In patients with a history or evidence of active peptic or intestinal ulceration, and only when considered essential in pregnant and nursing women. **Side-effects:** LEDERFEN is well tolerated by most patients. Gastro-intestinal symptoms, and occasionally skin rash are the most commonly reported. **Legal Category:** POM. **Presentation:** Light blue, film coated, capsule shaped 300 mg tablets engraved "LL300," or capsules printed "Lederle 300 mg," in bottles of 100. **Basic NHS Cost:** £16.24 per 100 tablets or capsules. PL/0095/0081. Further information is available on request to the company.

Lederle Laboratories, a division of Cyanamid of Great Britain Limited, Fareham Road, Gosport, Hants PO13 0AS. Tel: (0329) 236131.

References: 1. Guth PH, Ann Rev Med; 33 : 183-196 (1982) 2. Birnbaum J, et al, Pharmacology; 25 (Suppl. 1): 27-38 (1982) 3. Khan FM, to be published 4. Salzman RT, Reid RT, Eur J Rheum and Inflam; 5 (3) 318-325 (1982) 5. Roden DF, J Irish Med Assoc; 72 (No. 6.) 250-256 (1979).

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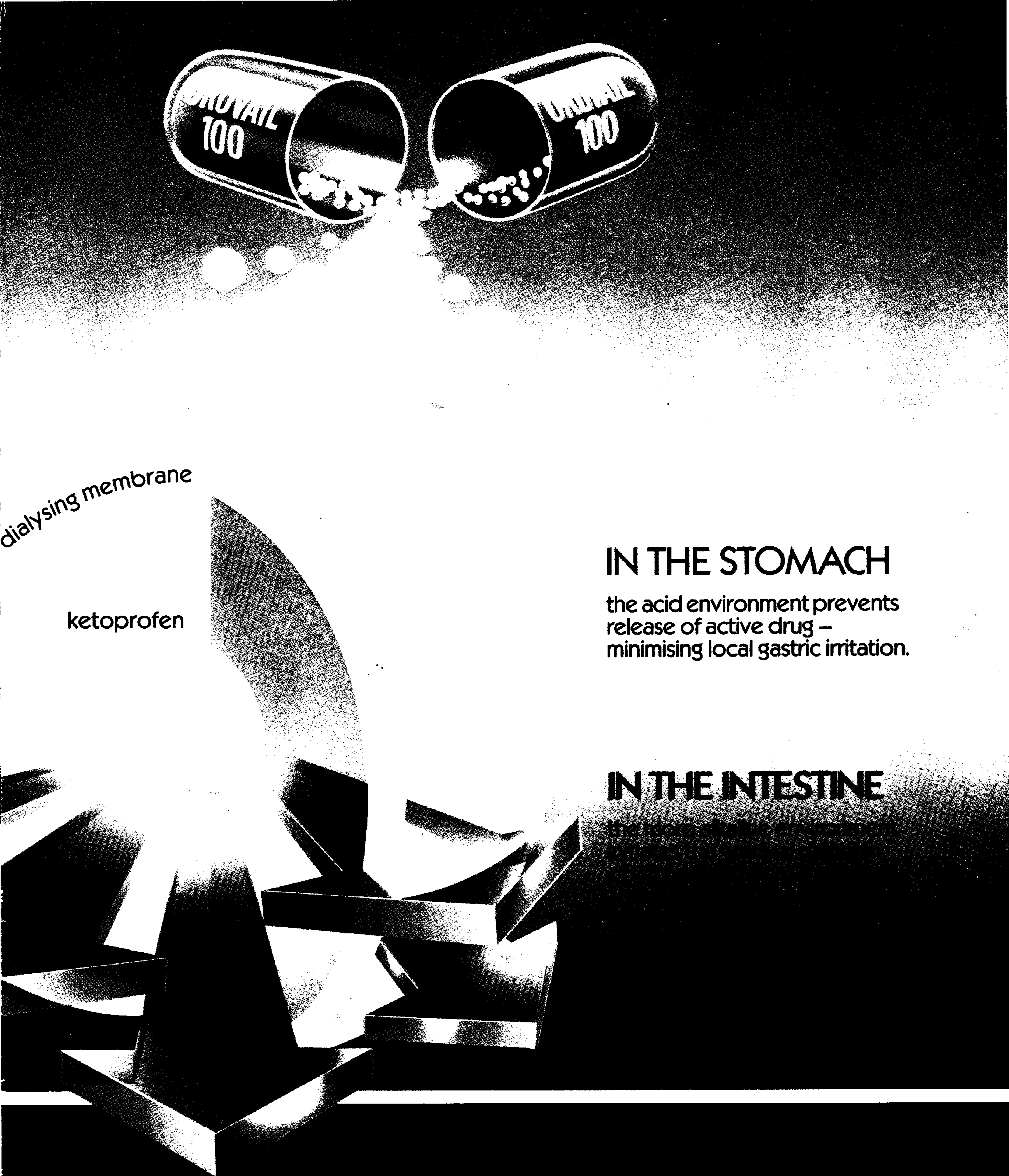

Prescribing information
Dosage: Orally with food, 100-200mg once daily.
Contra-indications: Recurring history of or peptic ulceration, chronic dyspepsia, use in children, in patients sensitive to aspirin or other non-steroidal anti-inflammatory drugs known to inhibit prostaglandin synthetase, or with bronchial asthma or allergic disease.
Precautions: Pregnancy, lactation. Dosage of concomitant protein-binding drugs may need modification.
Side-effects: Occasional gastro-intestinal intolerance, very rare gastro-intestinal haemorrhage, skin rashes.
Presentation: 100mg capsules PL [illegible]
Basic NHS Cost: [illegible] 100 x 100mg capsules [illegible]
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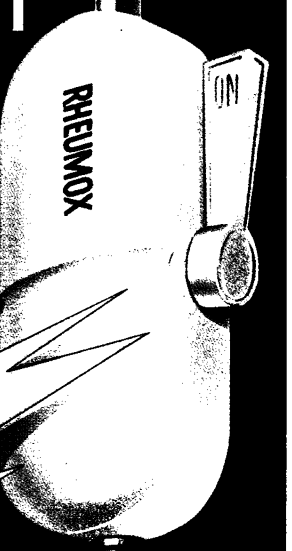

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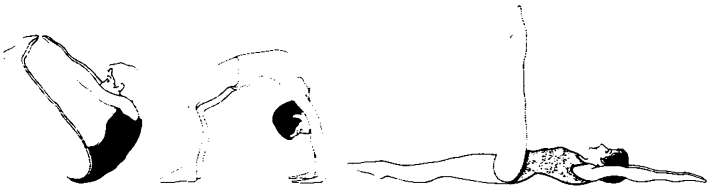

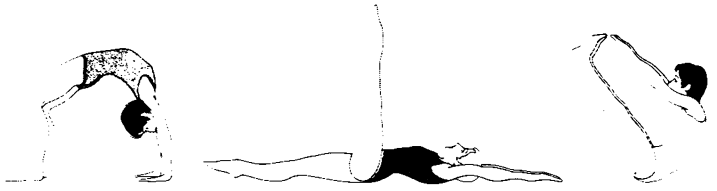

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GV 7

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towards mobility

Flosint offers major anti-inflammatory/analgesic activity WITHOUT an unacceptable level of adverse effects, possessing both the powerful anti-inflammatory properties of indomethacin and the potent analgesic effects of the propionic acid derivatives.

Flosint is particularly suitable for treatment of a wide range of arthritic disorders including:

- Osteoarthritis
- Rheumatoid arthritis
- Ankylosing spondylitis
- Non-articular rheumatism

Prescribing Information

Presentation
White, uncoated, biconvex, scored tablets containing 200mg indoprofen.

Indications
Rheumatoid arthritis, osteoarthritis, ankylosing spondylitis and other musculo-skeletal disorders.

Dosage
Osteoarthritis: the recommended dosage of indoprofen is two tablets in the morning and one tablet at night. The night-time dosage may be increased to two tablets for patients with more severe pain.
Rheumatoid Arthritis and Ankylosing Spondylitis: the recommended dosage of indoprofen is two tablets in the morning and two tablets at night.
A relatively small group of patients may prefer to take one tablet three times or even four times daily. Flosint allows for this flexibility.
As with similar drugs, the tablets should be taken with food to diminish the risk of gastric irritation.
Children: Paediatric usage has not been established.

Contra-Indications, Warnings, etc.
Sensitivity to this class of compounds, and active peptic ulceration.
Warnings: Bronchospasm may be precipitated in patients suffering from or with a previous history of bronchial asthma or allergic disease. Indoprofen is extensively protein-bound, and may displace oral anti-coagulants, sulphonylureas, etc. from binding sites. Dosage alterations may be needed in patients requiring concomitant treatment with these agents. Dosage reduction may be required in renal or hepatic impairment.
Pregnancy: The use of indoprofen during pregnancy should be avoided if possible, though animal experiments show no evidence of teratogenicity.
Adverse Effects: The most common adverse reactions caused by indoprofen are gastro-intestinal, including dyspepsia and nausea. Haemorrhage can occur rarely, as well as exacerbation of latent peptic ulceration. Other effects include mild central nervous system symptoms, such as dizziness and headache, and also skin rashes of various types. Decrease in platelet count has been observed rarely and blood dyscrasias may uncommonly occur. Limited data available suggest that abnormal liver function may be further impaired by treatment with indoprofen.
Overdosage: Gastric lavage and general supportive treatment should be carried out. There is no specific antidote.

Legal Category
POM.

Basic NHS Cost
Container of 100 tablets-£15.00.

Product Licence Number
PL3433/0034.

References
1. *Eur. J. Rheumatol. Inflamm.* 1981; **4**: 118. 2. *Ibid* 1981; **4**: 74. 3. *Ibid* 1981; **4**: 53. 4. *Brit. Med. J.* 1978; **1**: 274. 5. *Clin. Trials J.* 1982; **19**: 248. 6. *Eur. J. Rheumatol. Inflamm.* 1981; **4**: 126. 7. *Ibid* 1981; **4**: 135. 8. *Ibid* 1981; **4**: 144. 9. *Curr. Ther. Res.* 1978; **24**: 274. 10. *J. Oral Surg.* 1981; **39**: 21.

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without pain

Clinical evidence has confirmed that Flosint is at least as effective as other anti-inflammatory agents in relieving pain and inflammation[1-5] and is often better tolerated.[6-8]

FLOSINT
INDOPROFEN

- Highly effective anti-inflammatory properties[4,9]
- Powerful, rapid relief of pain – superior in pain relief to paracetamol (650mg) plus dextropropoxyphene (100mg)[10]
- Well tolerated by most patients – no reports of onycholysis or phototoxicity in long-term studies[6,7]
- Dosage tailored to meet the patients changing needs – simple b.d. dosage.

in arthritis

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fenclofenac

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Flenac is not only a first-class treatment for arthritic pain and inflammation; recent studies[1-4] point to an extra benefit not shared by other non-steroidal anti-inflammatories.

These studies show Flenac to have a beneficial effect on E.S.R. and on C-reactive protein and Rheumatoid Factor titres – the laboratory parameters considered to best reflect disease activity and correlate with radiological evidence of joint damage[5]

reducing eventual joint damage

relieving pain and inflammation

Prescribing information: Presentation Tablets of 300mg fenclofenac. **Indications** Chronic and sub-acute rheumatological conditions such as osteoarthrosis, rheumatoid arthritis, ankylosing spondylitis. **Dosage and Administration** Adults 600-1200mg (2-4 tablets) daily in divided doses morning and night with or after food. Flenac is not recommended for children. **Contra-indications** Active peptic ulceration or gastric bleeding. **Warnings** Flenac should not at present be prescribed for children or for pregnant or lactating women. Flenac should not be co-administered with anti-coagulants. Care should be taken when treating patients with known renal or hepatic dysfunction, eczema, asthma or sensitivity to other non-steroidal anti-inflammatory drugs. **Note** Flenac interferes with thyroid function tests. **Side-effects** Gastro-intestinal symptoms sufficient to require discontinuing treatment are rare. Rashes have occurred but have resolved shortly after withdrawal of the drug. **N.H.S. Price** £12.92 pack of 100. **P.L.** 44/0060. **References** 1. *Roy Soc Med* (CSS) (1980). No 28. p 95-100. 2. *Roy Soc Med* (CSS) (1980). No 28. p 17-22. 3. *Roy Soc Med* (CSS) (1980). No 28. p 57-60. 4. *Roy Soc Med* (CSS) (1980). No.28. p 109-112. 5. *Brit Med J* (1980). **280**. p 964. Additional information is available from: Reckitt & Colman Pharmaceutical Division, Hull HU8 7DS. Tel: 0482 26151. Flenac is a registered trade mark.

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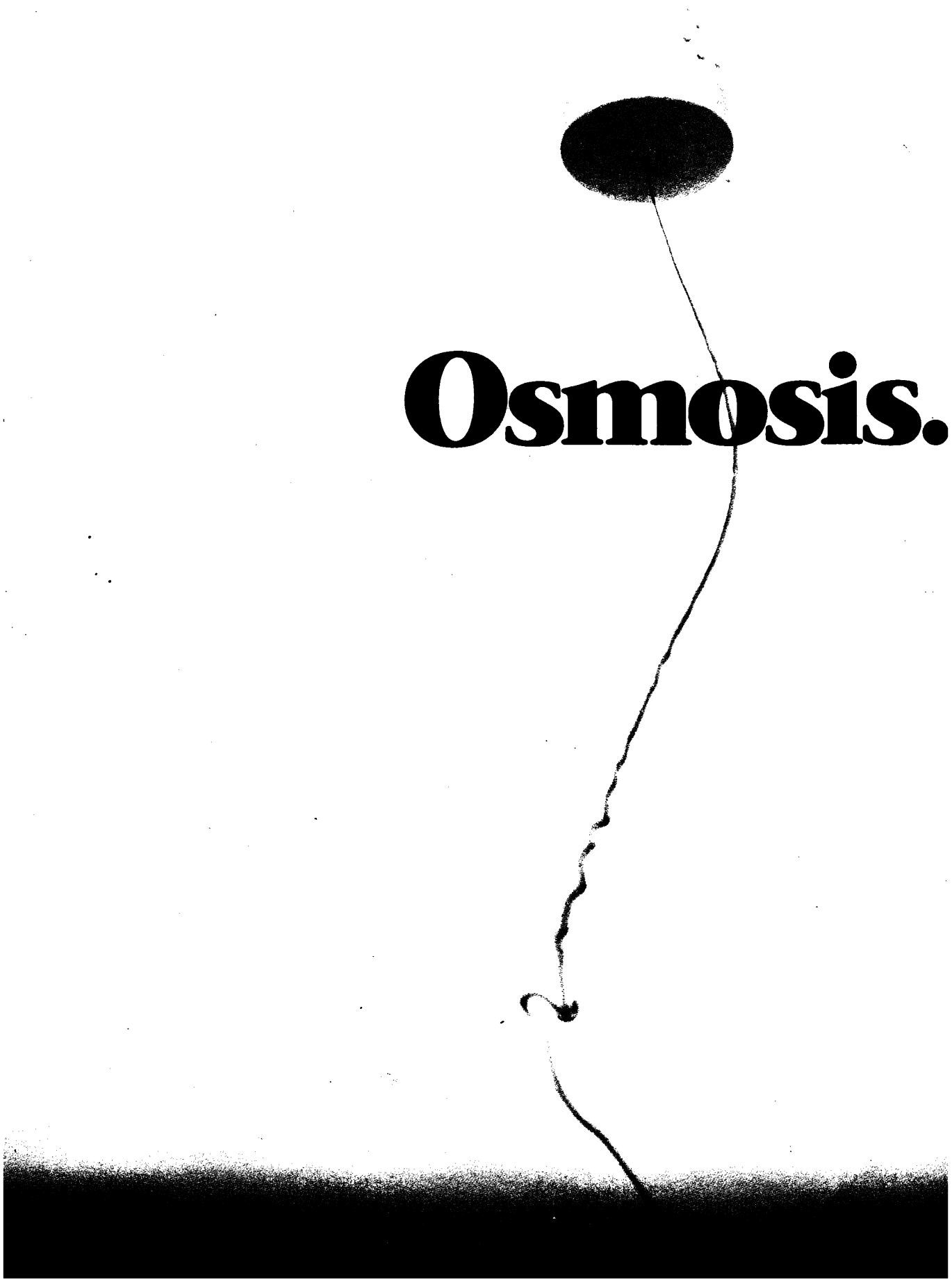
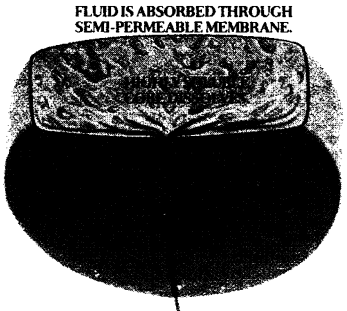


The tablet.



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With a precision and regularity that no other oral therapy can achieve.

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To help minimise GI and CNS side effects associated with conventional therapies.

One tablet daily will provide full 24-hour efficacy, for most patients.

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OSMOSIN: Highly effective relief from pain, inflammation and stiffness.

OSMOSIN: Minimises GI and CNS side effects.

℞ Osmosin 1 daily (30)

Abridged Product Information.

Full prescribing information is available on request and should be consulted before prescribing.

INDICATIONS Osteoarthritis; rheumatoid arthritis; ankylosing spondylitis; acute musculo-skeletal disorders and low back pain. Pain and associated symptoms of primary dysmenorrhoea.

DOSAGE Usually one daily. If required one twice daily, take whole - do not chew or crush - and take with food or immediately after a meal. The maximum dose is two per day.

CONTRA-INDICATIONS Active peptic ulcer, history of gastro-intestinal lesions, sensitivity to indomethacin or other non-steroidal anti-inflammatory agents, children, lactating women, and pregnancy.

PRECAUTIONS If GI symptoms occur, weigh benefits against risks of continuing. If GI bleeding occurs discontinue OSMOSIN. May mask the signs and symptoms of infection. Use cautiously in the elderly and in patients with a history of psychiatric disorders, epilepsy or parkinsonism. Monitor the prothrombin time when adding OSMOSIN to the treatment of patients on anticoagulants. Interactions: aspirin, probenecid, lithium, frusemide, thiazides, beta-blockers.

SIDE EFFECTS OSMOSIN is usually well tolerated. GI symptoms including nausea, dyspepsia, are most common. Isolated cases of peptic ulcer and bleeding have been reported with indomethacin as have hepatic CVS, and renal effects. CNS symptoms including headache, dizziness, rarely hypersensitivity - including skin rashes - and haematological reactions; ocular changes including blurred vision and corneal deposits have occurred.

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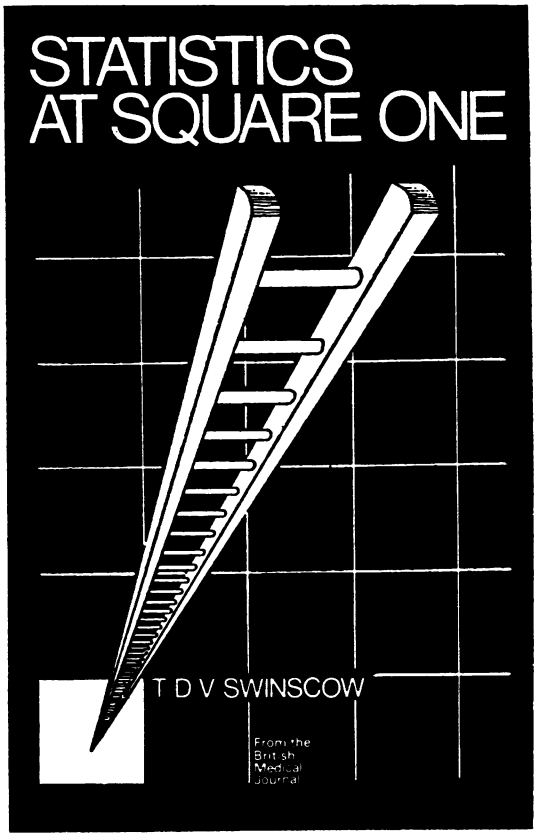
de Jong, Rudolph, H., *J.A.M.A.*, 1977, **237**, 1874.



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flurbiprofen

Active in Arthritis

Prescribing Information:
Presentation: Sugar-coated tablets, each containing either 50mg or 100mg of flurbiprofen. **Uses:** Froben is indicated in the treatment of rheumatoid disease, osteoarthrosis and ankylosing spondylitis. **Dosage:** 150mg to 200mg daily in 3 or 4 divided doses. In patients with severe symptoms or disease of recent origin, or during acute exacerbations, the total daily dose may be increased to 300mg in divided doses. **Contra-indications, Warnings etc:** Froben should not be given to patients with peptic ulceration. Care should be taken when administering the drug to patients with asthma or who have experienced bronchospasm with other anti-inflammatory or analgesic agents. The safety of Froben during pregnancy or lactation has not been established. In animal experiments, no teratogenic effects were demonstrated but parturition was delayed and prolonged. Side-effects: dyspepsia, heartburn and headache are the commonest encountered. Occasional skin rashes have been reported. Treatment of overdosage: gastric lavage and, if necessary, correction of serum electrolytes. There is no specific antidote. **Basic NHS Price:** 50mg tablets, 100 £8.24 100mg tablets, 100 £15.65. **Product Licence No:** 50mg tablets, PL0014/0167 100mg tablets, PL0014/0168. Further information is available on request.

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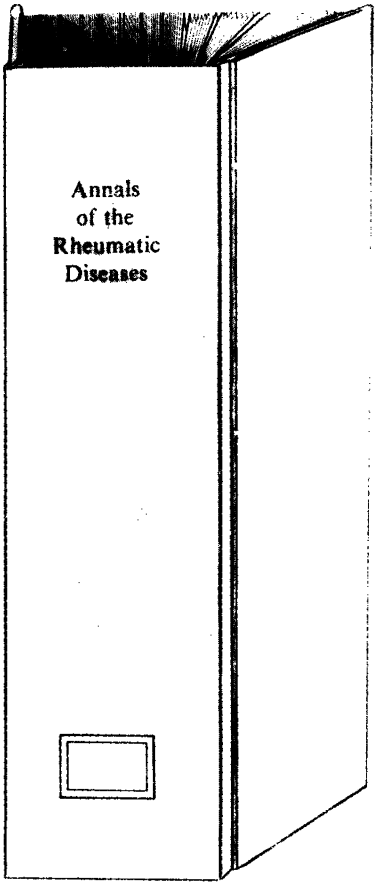

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