Spence’s polemic on duloxetine loses power because most of what it is based on is wrong. Making pronouncements in respected journals about good and bad medicine comes with the responsibility of knowing what you are talking about. Bad scholarship helps no one, especially those in pain; research in genetics, neurobiology, and psychology has contributed to huge advances in knowledge about the bio-psycho-social origins of pain.

Chronic pain is defined as pain of moderate or severe intensity that lasts for three months or more (think of a really bad headache lasting from Christmas to Easter). It affects one in five adults. Painful conditions are among the most prevalent conditions and are five of the top 11 in terms of years living with disability. Chronic pain destroys lives, has a huge negative impact on quality of life, is costly, reduces the ability to work or function inside the family, and may be associated with decreased life expectancy.

Here is a brief, non-comprehensive list of where Spence misses the point.

Women are not over-represented in trials. Chronic pain disproportionately affects women; their representation matches the epidemiology.

Pain is the most immediate of patient outcome measures, reported by patients themselves. Its subjectivity was recognised in the 1950s and dealt with. Different scales show excellent agreement, and patients in clinical trials record consistent pain levels over long periods, as in clinical practice.

Average benefits over placebo are in the order of 1 point on a 10 point rating score. But few patients are average; most have either little or great benefit. A responder is defined as someone with at least a 50% reduction in pain intensity maintained for 12 weeks, without intolerable adverse events that mean stopping treatment. Patients want this outcome, and it is accompanied by major improvements in sleep, function, and quality of life. Cochrane reviews do not report effective non-drug alternatives.

Several large good quality trials provide good evidence that duloxetine is effective in painful diabetic neuropathy and fibromyalgia. The effect size is not massive—a number needed to treat of five tells you that—but it is comparable to other treatments in neuropathic and other chronic pain, where treatment failure is expected more often than treatment success.

We are not “steeped in conflicts of interest.” Some (but not all) make declarations of interest, not quite the same as conflicts.

Spence omits the most important, valid, criticisms. These might include the bias against older treatments because authorities require trials fundable only by industry or government (which neither fund). Effective drugs like amitriptyline are understudied so have less evidence of efficacy to support their use. In addition, inappropriate imputation methods in statistical analysis...
of results can mean that efficacy is sometimes hugely inflated; this is not the case for duloxetine.

Patients need options. Encouraging people to act against the evidence is the real “bad medicine.”

Competing interests: ML, RH, and PW wrote a Cochrane review on duloxetine. AM has received honorariums for consulting from Eli Lilly. AM, DA, and EK were authors of a BMJ article criticised by Spence. CE and AW declare no conflicts.

Full response at: www.bmj.com/content/348/bmj.g139/rr/685335.

Cite this as: BMJ 2014;348:g1490
© BMJ Publishing Group Ltd 2014