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LETTER TO THE EDITOR

Authors' responses to Letter to the Editor regarding 'Early digoxin-specific antibody fragments to treat patients at risk of life-threatening toxicity'

To the Editor:

We thank Drs. Megarbane and Baud for the letter discussing the role of prophylactic use of digoxin-Fab in patients with elevated digoxin levels. This is in response to our recent review article on digoxin-specific antibodies in the treatment of digoxin toxicity.¹ The prophylactic use of half equimolar dose of digoxin-Fab in patients with 'a poor prognosis', elevated digoxin levels, but without any indication of severe digoxin toxicity, is a very expensive strategy. Further, there are no convincing data to suggest that this will improve outcomes compared with observation and treatment of any deterioration with digoxin-Fab.

The suggestion that the group of patients have a poor prognosis is based on analysis of patients with acute digoxin/digitoxin poisoning.² These 6 prognostic factors have not been validated in chronic digoxin poisoning and some are extremely common. A majority of asymptomatic patients with an elevated digoxin level would meet three or more criteria, which they suggest should indicate prophylactic digoxin-Fab. In their own series, over 95% of patients met two of the three criteria (age > 55 and underlying heart disease). So simply being male, OR having a heart rate of less than 60 bpm after atropine OR having a K value of greater than 4.5mmol/L would on their own be an indication for digoxin-Fab, if the authors' prognostic criteria were used. The result would be a tenfold increase in the use of digoxin-Fab.^{3,4}

We agree that patients with high digoxin concentrations from 'chronic toxicity' have a higher mortality rate. These patients are frequently old, with deterioration in underlying heart, kidney and/or liver disease. The deterioration in organ function might have caused the high digoxin concentration. Many of these patients will still die if digoxin-Fab is given. In a prospective multi-centre study of 150 patients treated with equimolar dose of digoxin-Fab, the mortality rate was 29% (43/150).⁵ The death in majority of patients was thought to be caused by other medical illnesses. Even if Digoxin-Fab

is not given, it is likely that patients, who die, will die 'with' rather than 'from' chronic digoxin toxicity.

We are currently conducting a prospective non-randomised study on acute and chronic digoxin poisoning and have frequently found that digoxin-Fab does not improve symptoms of patients with chronic digoxin toxicity even when half molar neutralising doses are used. It is even harder to measure an effect of prophylactic digoxin-Fab in asymptomatic patients. Thus, we would endorse the merit of conducting a randomised clinical trial to test the benefits of prophylactic digoxin-Fab, as it is plausible (but not likely) that there is a small benefit, and it will be impossible to determine this with less stringent research designs.

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Declaration of interest

The authors report no declarations of interest. The authors alone are responsible for the content and writing of the paper.

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