

Contents lists available at ScienceDirect

EClinicalMedicine

journal homepage: https://www.journals.elsevier.com/ eclinicalmedicine

EClinicalMedicine

Published by THE LANCET

Improving the management of the paracetamol poisoned patient

Anselm Wong^{a,b,c,*}, Andis Graudins^{d,e}, Kennon Heard^f, Kim Dalhoff^g, Marco L.A. Sivilotti^h

^a Victorian Poisons Information Centre, Emergency Department and Austin Toxicology Unit, Austin Health, Heidelberg, Victoria 3084, Australia

^b School of Clinical Sciences at Monash Health, Monash University, Victoria, Australia

^c Centre for Integrated Critical Care, University of Melbourne, Victoria, Australia

^d Monash Toxicology Service and Monash Emergency Research Collaborative, Dandenong Hospital, Victoria, Australia

^e School of Clinical Sciences, Faculty of Medicine, Nursing and Health Sciences, Monash University, Victoria, Australia

^f University of Colorado School of Medicine, Denver, Colorado, USA

^g Department of Clinical Medicine, Bispebjerg-Frederiksberg Hospital -, Bispebjerg, Denmark

h Departments of Emergency and of Biomedical & Molecular Sciences, Queen's University, Kingston, Ontario, Canada

ARTICLE INFO

Article history: Received 14 May 2019 Accepted 18 July 2019 Available online 20 July 2019

Dear Editor,

We read with interest the article by Pettie et al. [1] and commend the group for aiming to improve the management of paracetamol overdose.

We note a high proportion of (25.9%, n = 294) RIE patients in the 12-h SNAP protocol group did not have a 20-h blood concentration, largely as the result of self-discharge (n = 164) or medical decision (n = 118). Of these, six had elevated and rising ALT activity after 10 h of acetylcysteine including presumably one with a 10-h ALT of 961 U/L. It is reassuring to see a low rate of hepatotoxicity and coagulopathy (and zero deaths in Scotland), presumably attributable to excluding those presenting with hepatotoxicity. However, with imbalanced loss to follow up by 20 h, low event rates, and by modeling only the highest measured ALT (rather than rise from baseline [2,3]), a subtler efficacy signal may have been masked.

One strategy that may help prevent early discharge, in patients with signs of early liver injury, is to treat all patients requiring acetylcysteine with a 2-bag 20-h regimen [4,5], only ceasing the infusion for patients at

low risk of developing hepatotoxicity (i.e. ALT <40 U/L and a low paracetamol concentration) after 12-h of treatment [3].

It should also be acknowledged that the SNAP studies were performed under the supervision of specialist toxicology units. Wider adoption at other hospitals needs to be monitored to ensure safety and efficacy.

We look forward to further studies optimizing the dose and duration of acetylcysteine treatment to improve treatment of the paracetamol poisoned patient.

References

- Pettie J, Caparrotta TM, Hunter RW, et al. Safety and efficacy of the SNAP 12-hour acetylcysteine regimen for the treatment of paracetamol overdose. EClinicalMedicine 2019. https://doi.org/10.1016/j.eclinm.2019.04.005.
- [2] Bateman DN, Dear J, Thanacoody HK, et al. Reduction of adverse effects from intravenous acetylcysteine treatment for paracetamol poisoning: a randomised controlled trial. Lancet 2014;383(9918):697–704.
- [3] Wong A, McNulty R, Taylor DM, et al. The NACSTOP trial: a multi-center, clustercontrolled trial of early cessation of acetylcysteine in acetaminophen overdose. Hepatology 2019;69(2):774–84.
- [4] Wong A, Isbister GK, McNulty R, et al. Efficacy of a 2 bag acetylcysteine regimen to treat paracetamol overdose (2NAC study). Abstracts from the "39th International Congress of European Association of Poison Centres and Clinical Toxicologists (EAPCCT) 21-24th May 2019," Naples, Italy. Clin Toxicol 2019. https://doi.org/10. 1080/15563650.2019.1598646.
- [5] Wong A, Sivilotti ML, Mcnulty R, Gunja N, Graudins A. Accuracy of the paracetamolaminotransferase product to predict hepatotoxicity in paracetamol overdose treated with a 2 bag acetylcysteine regimen. Clin Tox 2018;56(3):182–8.

* Corresponding author.

E-mail address: anselm.wong@austin.org.au (A. Wong).

https://doi.org/10.1016/j.eclinm.2019.07.009

2589-5370/© 2019 Published by Elsevier Ltd. This is an open access article under the CC BY-NC-ND license (http://creativecommons.org/licenses/by-nc-nd/4.0/).