

Review of: "Severe Cutaneous Adverse Drugs Reactions: From Causes To Mechanisms"

Tallulah Spina-Tensini

Potential competing interests: The author(s) declared that no potential competing interests exist.

The text should be reviewed by a native speaker.

The definition of "recent articles" should be mentioned, (e.g. from 20xx to 2022) in the abstract and in the body of the text. Authors should describe how many articles were found using those descriptors and how many were used. The last phrase about authors knowledge in methods should be suppressed.

References should be checked (eg table 1 is referred to a study of incidence and most probably not to the original article that designed those definitions).

Discussion

Antiseizure medication is the updated expression for the old Anticonvulsants or antiepileptic drugs (both expressions were used in the text), similarly, toxic epidermal necrolysis is the preferred expression, rather than Lyell syndrome.

Another well known expression is CAR (Cutaneous adverse reaction, as instituted by the RegiSCAR study) instead of ADR.

Erythema is a signal, such as morbilliform, roseoliform etc. The authors would be more specific if cited the presentation such as Urticaria (cited) or maculopapular exanthema.

I recommend presenting each CAR in a different topic, with description of the clinical manifestation and main drugs associated to a particular CAR.

Also, nowadays much more is known about physiopathology of CAR, specially about the interaction between drugs and immune system. Too little was mentioned about HLA role in CAR, and what was mentioned was incomplete (eg HLA-B*15:02 was described in strong association to SJS induced by CBZ in Han Chinese, not any severe CAR).

Rash characteristics in DRESS is well defined and used in diagnostic criteria suggested by Kardaun et al, 2007.

Are all images from the same article? Do the authors have the permission to use it? Which kind of CAR do they represent? A little explaining text about those pictures would further enlighten the reader.

The authors are engaged with the theme and much work have already been done. I hope this review will help to improve the text and that someday we can read an complete, organized and informative article born from this group.

Here are some references that may help:

1. Pichler WJ. Immune pathomechanism and classification of drug hypersensitivity. *Allergy Eur J Allergy Clin Immunol.* 2019;74(8):1457-1471. doi:10.1111/all.13765
2. Bellón T. Mechanisms of Severe Cutaneous Adverse Reactions: Recent Advances. *Drug Saf.* 2019;(Apr 24). doi:10.1007/s40264-019-00825-2
3. Hoetzenecker W, Nägeli M, Mehra ET, Jensen AN, et al. Adverse cutaneous drug eruptions: current understanding. *Semin Immunopathol.* 2016;38(1):75-86. doi:10.1007/s00281-015-0540-2
4. Kardaun SH, Sidoroff A, Valeyrie-Allanore L, Halevy S, et al. Variability in the clinical pattern of cutaneous side-effects of drugs with systemic symptoms: Does a DRESS syndrome really exist? *Br J Dermatol.* 2007;156(3):609-611. doi:10.1111/j.1365-2133.2006.07704.x