Title Goes Here

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**Keywords:** anaphylaxis, adrenaline (epinephrine), beta-blockers, insect venom allergy, yellow-jacket

**Highlights:** These are the highlights. **Document statistics:** Word count, figures, tables, references

## Loading venomanaphylaxiscompendium

## Loading required package: dplyr

##   
## Attaching package: 'dplyr'

## The following objects are masked from 'package:stats':  
##   
## filter, lag

## The following objects are masked from 'package:base':  
##   
## intersect, setdiff, setequal, union

## Loading required package: magrittr

## Loading required package: ggplot2

## Loading required package: forestplot

## Loading required package: grid

## Loading required package: checkmate

## Loading required package: summarytools

## Loading required package: DiagrammeR

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##   
## Attaching package: 'testthat'

## The following objects are masked from 'package:magrittr':  
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## equals, is\_less\_than, not

## The following object is masked from 'package:dplyr':  
##   
## matches

# Abstract

Insect-venom elicited anaphylaxis is a common hypersensitivity reaction which may be life-threatening. Using the data from the European Anaphylaxis Registry (11596 cases in total) we identified insect-venom elicited anaphylaxis cases (n = 4482) and analyzed these in comparison to anaphylaxis elicited by other elicitors (n = 7114).

The data show that 68.57% of all insect elicited cases were elicited by yellow jackets, followed by bees (21.86%). The insect venom elicited cases occurred mostly in outdoor places (44.65%) patients’ homes (12.87%) or urban places (9.616%).

Skin, gastrointestinal and respiratory symptoms occurred less frequently in insect elicited cases of anaphylaxis, whereas cardiologic symptoms (with hypotension, collapse, and loss of consciousness) were more frequent. Intramuscular adrenaline (as a first-line therapy) was administered significantly less often in insect venom elicited cases (4.04%, p < 0.0001). The mortality rate in insect anaphylaxis was comparable (0.156%) to other cases (0.295%, p = 0.174).

Patients who experienced insect-venom anaphylaxis were older (p < 0.0001), more often had concomitant mastocytosis (p < 0.0001) and cardiologic conditions (p < 0.0001) and females more often had concomitant thyroid diseases and less often suffered from a food allergy or atopic dermatitis.

Symptoms of insect venom anaphylaxis are distinctively different from other reactions, indicating that the therapy of insect elicited cases of anaphylaxis should be considered separately. Indeed we observed different therapeutic patterns in insect elicited cases of anaphylaxis (more antihistaminics but fewer corticosteroids, bronchodilators, and surprisingly - adrenaline). This indicates that the management of insect-venom induced anaphylaxis may be improved and is especially required in patients with concomitant cardiologic conditions and these with hyperreactive mast cells.

# Introduction

# Methods

The European Anaphylaxis Registry (**???**) database from March 2018 wassearched for anaphylaxis cases elicited by insect venom. The flowchart in figure ?? represents the detailed case-selection process.

The final database consisted of 3295 cases of insect elicited anaphylaxis from 11 countries. Severe reactions were identified based on the definition by NIAID/FAAN (**???**) and presented with significant hypoxia, hypotension, confusion, collapse and loss of consciousness, or incontinence. We compared the frequency of various elicitors, symptoms, and factors known to increase the risk of severe anaphylaxis (**???**) in both groups. We evaluated symptoms, managment and risk factors of insect elicited cases in comparison to other known triggers of anaphylaxis.

The statistical analysis was performed in the R Statistical Package (**???**). Simple comparison of categorical variables was performed using Fisher’s exact test, continuous variables were analyzed using Mann-Whitney U test. We defined statistical significance as α = 0.05. Data along with the analysis script can be accessed at online at….

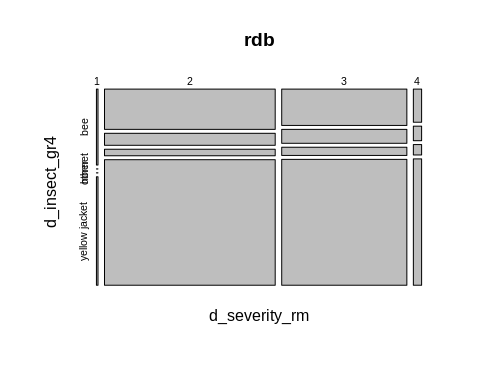
# Results

## Demographics

#> n Age Cardiologic DM Food allergy Mastocytosis Malignancy  
#> female 1520 46.9 22.763 2.368 1.842 2.961 1.842   
#> male 1775 44.4 26.423 3.831 1.408 3.042 1.183   
#> Atopic dermatitis tryptase [median]  
#> female 2.5 4.2   
#> male 1.634 4.63

#> ### Cross-Tabulation / Row Proportions   
#> \*\*Variables:\*\* d\_centres\_country \* q\_340\_insects   
#> \*\*Data Frame:\*\* rdb   
#>   
#> | | | | | | | | | | | | |  
#> |------------------:|--------------:|----------:|-------------:|----------:|-------------:|-----------:|----------:|--------------:|------------:|---------------:|---------------:|  
#> | | q\_340\_insects | no | bee | horse fly | hornet | bumble-bee | mosquito | yellow jacket | insects | \<NA\> | Total |  
#> | d\_centres\_country | | | | | | | | | | | |  
#> | Germany | | 0 (0.00%) | 408 ( 7.90%) | 3 (0.06%) | 118 ( 2.29%) | 7 (0.14%) | 4 (0.08%) | 1803 (34.92%) | 79 (1.53%) | 2741 ( 53.09%) | 5163 (100.00%) |  
#> | Austria | | 0 (0.00%) | 69 (14.11%) | 1 (0.20%) | 22 ( 4.50%) | 3 (0.61%) | 0 (0.00%) | 141 (28.83%) | 12 (2.45%) | 241 ( 49.28%) | 489 (100.00%) |  
#> | Switzerland | | 0 (0.00%) | 105 ( 9.08%) | 0 (0.00%) | 14 ( 1.21%) | 0 (0.00%) | 0 (0.00%) | 146 (12.62%) | 10 (0.86%) | 882 ( 76.23%) | 1157 (100.00%) |  
#> | Greece | | 0 (0.00%) | 3 ( 2.27%) | 0 (0.00%) | 4 ( 3.03%) | 0 (0.00%) | 0 (0.00%) | 2 ( 1.52%) | 0 (0.00%) | 123 ( 93.18%) | 132 (100.00%) |  
#> | Poland | | 0 (0.00%) | 35 (17.07%) | 0 (0.00%) | 6 ( 2.93%) | 0 (0.00%) | 0 (0.00%) | 33 (16.10%) | 0 (0.00%) | 131 ( 63.90%) | 205 (100.00%) |  
#> | Spain | | 0 (0.00%) | 6 ( 1.71%) | 0 (0.00%) | 3 ( 0.86%) | 1 (0.29%) | 0 (0.00%) | 7 ( 2.00%) | 0 (0.00%) | 333 ( 95.14%) | 350 (100.00%) |  
#> | France | | 0 (0.00%) | 21 ( 2.29%) | 0 (0.00%) | 12 ( 1.31%) | 0 (0.00%) | 0 (0.00%) | 29 ( 3.16%) | 2 (0.22%) | 854 ( 93.03%) | 918 (100.00%) |  
#> | Bulgaria | | 0 (0.00%) | 10 ( 7.25%) | 0 (0.00%) | 2 ( 1.45%) | 1 (0.72%) | 0 (0.00%) | 4 ( 2.90%) | 0 (0.00%) | 121 ( 87.68%) | 138 (100.00%) |  
#> | Netherlands | | 0 (0.00%) | 0 ( 0.00%) | 0 (0.00%) | 0 ( 0.00%) | 0 (0.00%) | 0 (0.00%) | 0 ( 0.00%) | 0 (0.00%) | 0 ( 0.00%) | 0 ( 0.00%) |  
#> | Denmark | | 0 (0.00%) | 0 ( 0.00%) | 0 (0.00%) | 0 ( 0.00%) | 0 (0.00%) | 0 (0.00%) | 0 ( 0.00%) | 0 (0.00%) | 0 ( 0.00%) | 0 ( 0.00%) |  
#> | Italy | | 0 (0.00%) | 33 (11.58%) | 0 (0.00%) | 47 (16.49%) | 0 (0.00%) | 0 (0.00%) | 80 (28.07%) | 0 (0.00%) | 125 ( 43.86%) | 285 (100.00%) |  
#> | UK | | 0 (0.00%) | 0 ( 0.00%) | 0 (0.00%) | 0 ( 0.00%) | 0 (0.00%) | 0 (0.00%) | 0 ( 0.00%) | 0 (0.00%) | 0 ( 0.00%) | 0 ( 0.00%) |  
#> | Russia | | 0 (0.00%) | 0 ( 0.00%) | 0 (0.00%) | 0 ( 0.00%) | 0 (0.00%) | 0 (0.00%) | 0 ( 0.00%) | 0 (0.00%) | 0 ( 0.00%) | 0 ( 0.00%) |  
#> | Ireland | | 0 (0.00%) | 0 ( 0.00%) | 0 (0.00%) | 1 ( 0.63%) | 0 (0.00%) | 0 (0.00%) | 1 ( 0.63%) | 1 (0.63%) | 155 ( 98.10%) | 158 (100.00%) |  
#> | Slovenia | | 0 (0.00%) | 0 ( 0.00%) | 0 (0.00%) | 0 ( 0.00%) | 0 (0.00%) | 0 (0.00%) | 0 ( 0.00%) | 0 (0.00%) | 1 (100.00%) | 1 (100.00%) |  
#> | not EU | | 0 (0.00%) | 1 ( 0.85%) | 0 (0.00%) | 0 ( 0.00%) | 0 (0.00%) | 0 (0.00%) | 0 ( 0.00%) | 6 (5.13%) | 110 ( 94.02%) | 117 (100.00%) |  
#> | Total | | 0 (0.00%) | 691 ( 7.58%) | 4 (0.04%) | 229 ( 2.51%) | 12 (0.13%) | 4 (0.04%) | 2246 (24.65%) | 110 (1.21%) | 5817 ( 63.83%) | 9113 (100.00%) |





## Symptoms

#> # A tibble: 10 x 7  
#> variableName counts\_1 counts\_2 fraq\_1 fraq\_2 pval section  
#> <chr> <int> <int> <dbl> <dbl> <dbl> <fct>   
#> 1 q\_114 2732 2804 0.840 0.545 5.24e-171 sympto…  
#> 2 q\_114\_dizziness 1512 1011 0.465 0.197 1.68e-151 sympto…  
#> 3 q\_120\_time\_between\_… NA NA NA NA 4.91e- 78 sympto…  
#> 4 q\_114\_loss\_of\_consc… 833 629 0.318 0.139 1.11e- 73 sympto…  
#> 5 q\_112\_abdominal\_pain 155 845 0.0488 0.165 5.54e- 56 sympto…  
#> 6 q\_114\_hypotension\_c… 1426 1427 0.438 0.277 4.77e- 53 sympto…  
#> 7 q\_113\_wheezing\_expi… 99 660 0.0437 0.157 7.55e- 41 sympto…  
#> 8 q\_112\_incontinence 223 77 0.0701 0.0150 7.27e- 39 sympto…  
#> 9 q\_113\_throat\_tightn… 639 621 0.270 0.147 1.30e- 33 sympto…  
#> 10 q\_112\_nausea 885 856 0.278 0.167 1.51e- 33 sympto…

Patients with Insect Venom Allergy had a lot more cardiologic symptoms than anaphylaxis cases that were elicited through other triggers.  
Dizziness, Loss of consciousness hypotension, collapse, Throat tightness and nausea were a lot more often associated with the insect cases.

Forest plot ! Risk that the symptom will occure given that the reaction was elicited by Insects or not

## Co-factors

## Managment

# Discussion

# Conclusion

# Acknowledgements

##### pagebreak

# References

##### pagebreak

### Colophon

This report was generated on 2019-02-05 21:57:21 using the following computational environment and dependencies:

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#> setting value   
#> version R version 3.5.1 (2018-07-02)  
#> os Ubuntu 18.10   
#> system x86\_64, linux-gnu   
#> ui X11   
#> language en\_US   
#> collate en\_US.UTF-8   
#> ctype en\_US.UTF-8   
#> tz Europe/Berlin   
#> date 2019-02-05   
#>   
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#> [1] /home/wolass/R/x86\_64-pc-linux-gnu-library/3.5  
#> [2] /usr/local/lib/R/site-library  
#> [3] /usr/lib/R/site-library  
#> [4] /usr/lib/R/library  
#>   
#> P ── Loaded and on-disk path mismatch.

The current Git commit details are:

#> Local: master /home/wolass/Documents/venomanaphylaxiscompendium  
#> Head: [f5c44f7] 2019-02-04: Add GitHub links to DESCRIPTION