

ANSWERING REVIEWERS

Name of journal: World Journal of Immunology

ESPS manuscript NO: 21054

Title: New insights on Chitinases immunologic activities

Reviewer's code: 00503062

Comments to authors

This manuscript reviewed chitinases immunologic activities. The review comprehensively reviewed the functions and roles of CHIT-1, CHI3L1 and CHI3L2 in inflammation and various diseases. Basically, the manuscript is well written and the references are new and informative. But, the functions of these proteins are so broad and it is hard to follow the review. It is better to set more sections in this review to organize the text. And many careless mistakes are found through manuscript. Authors should check the text carefully again before the submission. I give some suggestions below. "Chitinases" is not necessary to be capitalized through manuscript. Some references are duplicated. Check references carefully. For example, reference #7 and #68 are the same. Words in title of #7 are not necessary to be capitalized (page 12, lines 18, 19) Reference #8 and #40 are the same. Title of #8 (your paper!) is not correct. Page 2, line 7. Spell out "CLPs". Page 5, line 6. "malaria" should be "malaria parasite". Page 5, line 8. Spell out "NAFLD". Page 6, line 23. Remove a comma after "AMCase". Page 7, line 3. "an haplotype" reads "a haplotype". Page 7, lines 1-3 from bottom. Show a reference for the statement. Page 8, line 4. "is capable to induce" should be "is capable of inducing". Page 8, lines 8, 9. The positions of [59] and [60] are strange. Check them. Page 8, line 11. "INF" reads "IFN". Page 8, line 12. Add "diseases" after "infectious". Page 8, line 16. Add a period after "atherosclerosis". Page 8, line 20. Spell out "HMMs". Page 9, lines 3, 5. "CHIT3L1" reads "CHI3L1". Page 9, line 4. "Escherichia coli" should be italicized. Page 10, line 1. Spell out "IBL". Page 10, lines 2, 3. "IBL" is enough here. Page 10, line 4. Show a reference for the statement.

Response to reviewer comments

Through manuscript "Chitinases" has been corrected with chitinases

The references have been corrected.

All the duplicated references have been deleted.

Words in title of #7 has been changed (page 12, lines 18, 19)

Page 2, line 7. "CLPs" has been spelled out.

Page 5, line 6. "malaria" has been modified in "malaria parasite".

Page 5, line 8. "NAFLD" has been spelled out.

Page 6, line 23. Comma after "AMCase" has been removed

Page 7, line 3. "an haplotype" has been corrected in "a haplotype".

Page 7, the reference for the statement has been added

Page 8, line 4. "is capable to induce" has been corrected in "is capable of inducing".

Page 8, lines 8, 9. The positions of [59] and [60] has been corrected.

Page 8, line 11. "IFN has been corrected

Page 8, line 12. after "infectious" was added "diseases"

Page 8, line 16. after "atherosclerosis" a references has been added

Page 8, line 20. HMMs has been deleted and was corrected with "human macrophages".

Page 9, lines 3, 5. "CHIT3L1" has been modified "CHI3L1".

Page 9, line 4. "Escherichia coli" has been italicized.

Page 10, line 1. "IBD" has been spelled out.

Page 10, lines 2, 3. "IBL has been corrected with IBD.

Page 10, line 4. A reference for the statement has been provided

Reviewer's code: 00502947

Comments To Authors

This review focuses on new insights on chitinases in immune regulation. However there is very little direct evidence for a role being discussed and I think that the authors need to be careful in over emphasising views without providing evidence. There are a couple of major issues which authors' attention is required. I looked at the last 2 references and the content of those papers do not appear to say what author's say in the text. This raises the issue as to how accurate the authors have been in their references and needs to be addressed. The English expression is poor and in some cases I cannot understand what the authors' mean. I have attempted to list some of these below but there are many more. Please take care and spend more time and indeed get assistance with the English expression including sentence structure. Page 4 Line 6: Correct: that course with increased' change to 'that are caused by increased' Line 17: change 'which to act as immunological' to 'which acts as adjuvants'. Line 18: Change 'evidence indicate that' to 'evidence indicates that' Page 5 Line 3: Change to ' Saharan regions suggests that Line 7: Change to ' Other studies have been' Line 10: Delete 'is' Line 15: The word is 'context' Line 16 Word should be 'role' Line 18: Change 'detection' to 'expression' Line 18-19: 'infection-induced ' Line 19-23: Argument difficult to follow: Is IL-4 good or bad for atherosclerosis? Page 6 Line 5. I am not sure what you mean by prerogative in the context of this sentence? Line 13 : Change to contains Line 21: Change to demonstrated Line 21: change to 'role' Line 22: restructure sentence Line Line 26: 'precociously, and weakly'. Provide evidence to support this. Line 30: Difficult to know what this sentence means. Page 7 Line 5: data is plural. Should be either' The data or These data. Line 9: delete 'to' induce changes Line 15: explain that analog allosamidin is an inhibitor of chitinase. Line 15: change 'improved' to 'enhanced' Line 28: AMCase prevents Page 8 Line 2: add the 'in the absence' Line 4: change to capable of inducing. Line 30: Change to 'clearance' Page 10 Line 1: define IBL here and not in next paragraph. Line 6: what

does ..'prevented by the pre-storage leukocyte depletion by filtration of whole blood' mean? Line 11: suddenly another name appears for CHI3L1 ie YKL40 Page 11 Line 2-4: Appears to be incorrect reference. Ref 95 talks about gene signatures in EC and macrpphages in lymph node sinuses. Also ref 96. Line 11: 'conditions in which they' Lines 16, 17: Not sure if this can be substantiated as most of the discussion is correlative with expression or release from neutrophils or macrophages – not direct proof that these enzymes are causative.

Response to reviewer comments

Page 4 Line 6: Correct: 'that course with increased' has been changed to 'that are caused by increased'

Line 17: 'which to act as immunological' has been changed to 'which acts as adjuvants'.

Line 18: Change 'evidence indicate that' has been changed to 'evidence indicates that'

Page 5 Line 3: ' Saharan regions suggests that... has been corrected

Line 7: ' Other studies have been' has been corrected

Line 10: 'is' has been deleted

Line 15: The word 'contest' has been changed

Line 16: "rule" has been correct with 'role'

Line 18: 'detection' has been changed with 'expression'

Line 18-19: infection disease has been changed: 'infection-induced '

Line 19-23: the statements: "Additionally, IL-4 is well-appreciated antagonist of the M1 response and macrophage pro-inflammatory properties. In addition, IL-4 promotes a M2 phenotype in regressive" have been deleted

Page 6 Line 5. prerogative has been changed with "exclusively"

Line 13 : contains has been changed with has

Line 21: demonstrates has been corrected with demonstrated

Line 21: 'role' has been corrected

Line 26: 'precociously, and weakly'. The sentence was modified with "indicated AMCase is ineffective against infections"

Page 7 Line 5: data is plural. Ok These data.

Line 9: delete 'to' induce changes OK

Line 15: explain that analog allosamidin is an inhibitor of chitinase. Ok

Line 15: 'improved' was changed to 'enhanced'

Line 28: AMCase prevents Ok

Page 8 Line 2: add the 'in the absence' OK

Line 4: change to capable of inducing OK

Line 30: 'clearance' has been changed with removal

Page 10 Line 1: IBD" has been defined

Line 6: the sentence: 'prevented by the pre-storage leukocyte depletion by filtration of whole blood' has been modified "which were avoided depleting leukocytes by filtration of whole blood in order to inhibit extracellular CHI3L1 accumulation during storage of erythrocyte



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components”

Line 11: YKL40 has been corrected

Page 11 Line 2-4: reference has been corrected

Line 11: ‘conditions in which they’ Ok