

The Correlation between Laboratory Markers in Patients with Stimulants Dependence in Diwaniyah City

Noor Ali Gebur^{1*}

¹Department of Chemistry, College of Science, University of Al-Qadisiyah, Diwaniyah, Iraq

Abstract: Background and Aim: In stimulants dependence, the initial immune defense that play a role in response to central nervous system (CNS) inflammation. A novel nucleotide (TRB3 Q84) has been linked to a genetic variant with several metabolic disorders. In serum of individuals with stimulants dependence, assess of TRB3 Q84 levels and find associations with other markers, this was our aim in this study. **Results:** The *mean* of TRB3 Q84, IL-6 and CTX levels showed a significantly increased in stimulants dependence as compared with control. **Conclusions:** The present study demonstrated that individuals with stimulants dependence exhibited a significantly elevated TRB3 Q84 levels as compared to the group of control. In the turn, TRB3 Q84 had a significant strong direct relationship with (IL-6, CTX) these findings suggest that TRB3 Q84 levels may serve a protective function during the early stages of cellular dysfunction associated with stimulants dependence that contributing to stroke pathogenesis.

Keywords: Stimulants Dependence, Inflammatory markers.

Copyright © 2026 The Author(s): This is an open-access article distributed under the terms of the Creative Commons Attribution 4.0 International License (CC BY-NC 4.0) which permits unrestricted use, distribution, and reproduction in any medium for non-commercial use provided the original author and source are credited.

Research Paper

*Corresponding Author:

Noor Ali Gebur

Department of Chemistry, College of Science,
University of Al-Qadisiyah, Diwaniyah, Iraq

How to cite this paper:

Gebur, N. A. (2026). The Correlation between Laboratory Markers in Patients with Stimulants Dependence in Diwaniyah City. *Middle East Res J. Case Rep*, 6(4), 30-35.

<https://doi.org/10.36348/merjmcr.2026.v06i04.001>

Article History:

| Submit: 25.07.2026 |

| Accepted: 27.08.2026 |

| Published: 02.09.2026 |

1. INTRODUCTION

In stimulants dependence, the initial immune defense that play a role in response to central nervous system (CNS) inflammation. These immune cells secrete a range of cytokines of pro-inflammatory which is responsible for recruiting neutrophils to the site of inflammation [1-5]. Appropriate antimicrobial therapy often leads to significant improvement for CNS inflammation that can result in tissue damage and a variety of clinical manifestations. A well-coordinated immune response by the host is essential for pathogen clearance and symptomatic relief in affected [6-12].

A novel nucleotide the gene of human the tribbles* - of kinase 3 – by glutamine* at the position 84 (TRB3 Q84) has been linked to a genetic variant with several metabolic disorders [13-15], including inflammatory disease, nephropathy, chronic kidney disease and other metabolic syndromes [16-18].

Inflammatory bowel results in; destruction of; the alveolar - and other inflammatory illnesses bone that supporting the teeth compared to healthy individuals bone resorption of affected joints be characterized by arthritis of rheumatoid and systemic lupus erythematosus disease [19-20].

The preventing managing bone loss in inflammatory the pave of the way for novel therapeutic approaches of the osteoclasts are the cells of sole that responsible for; osteoporosis – and - across various diseases follows a similar pattern: surpasses that of osteoblasts bone resorption arises when osteoclast activity [21-22]. Inflammation influences this bone, the interaction between inflammation and osteoclast activity patient population by a deeper understanding. Bone loss first regulates osteoclast formation (osteoclastogenesis) by degradation primarily through two key pathways modulating stimulating pathway the factor of; macrophage colony [23-25].

Assess of TRB3 Q84 levels in serum of individuals with stimulants dependence in addition to find associations with the other markers, was the aim of our this study.

2. EXPERIMENTAL

The collection of blood samples process was at fasting <8–12> hours from patients and healthy, then blood was clotted in plain tubes at room temperature.

3. RESULTS

The results in Table 1 showed the mean of IL-6, also CTX and TRB3 Q84 levels were increased with

significantly in stimulants dependence; as compared to - control. These findings are depicted in Figure 1 (A, B, C).

Table 1: Data for stimulants dependence and control

Parameters	Groups		P-value
	Control Mean $\pm$ SD (n=60)	Stimulants Dependence Mean $\pm$ SD (n=60)	
IL-6 (pg/mL)	9 $\pm$ 0.6	22 $\pm$ 3	0.05
CTX (ng/mL)	0.5 $\pm$ 0.2	0.89 $\pm$ 0.2	0.03
TRB3 Q84 (ng/mL)	1.9 $\pm$ 0.2	3.2 $\pm$ 0.3	0.02

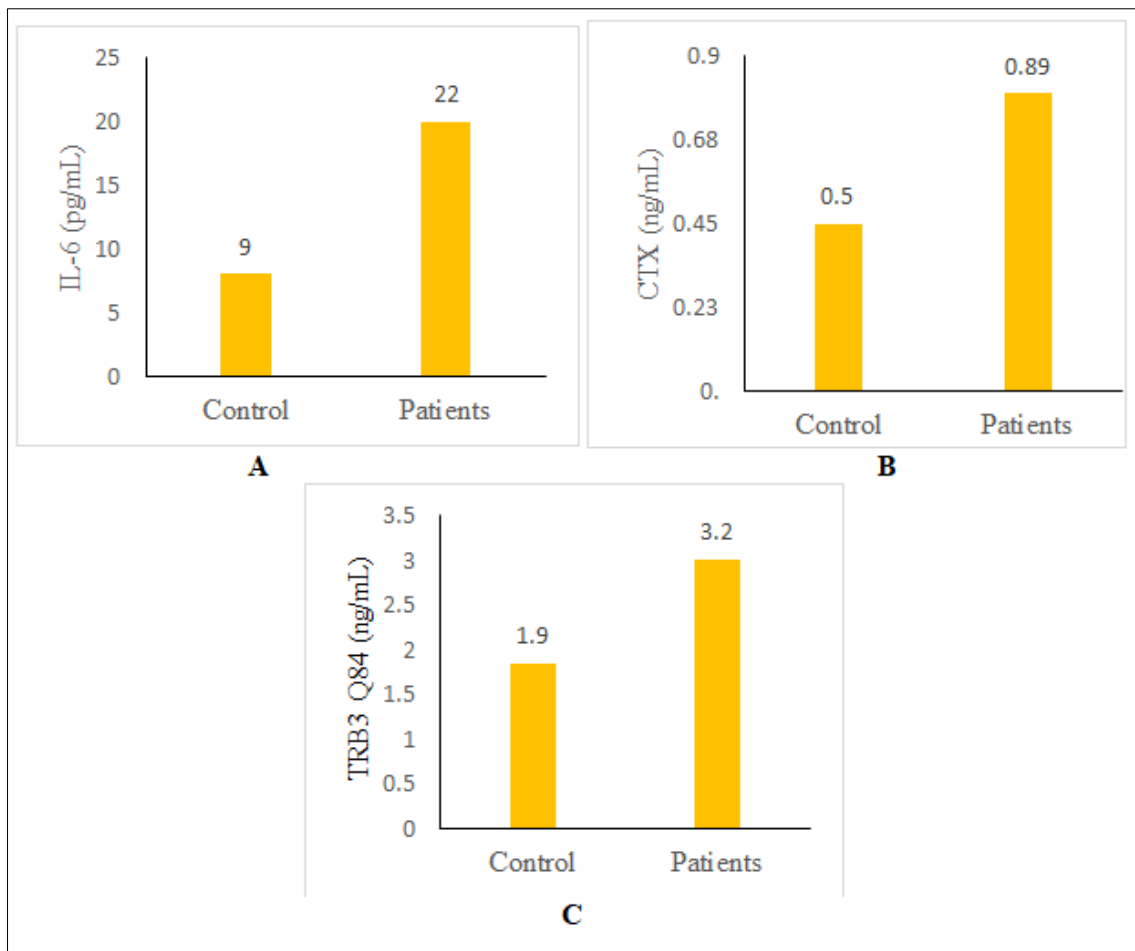


Figure 1: Comparison the levels of A: IL-6, B: CTX and C: TRB3 Q84 between stimulants dependence and control

Table 2 showed the results of a linear regression analysis for assessing the correlation of TRB3 Q84 levels with biomarkers in stimulants dependence individuals.

TRB3 Q84 levels showed a strong significant direct correlation with IL-6 and CTX levels, as represented in Figure 2 (A, B).

Table 2: TRB3 Q84 levels correlation with other biomarkers in stimulants dependence

Parameters	TRB3 Q84 (ng/mL)	
IL-6 (pg/mL)	r	0.98
	P-value	0.05
CTX (ng/mL)	r	0.97
	P-value	0.01

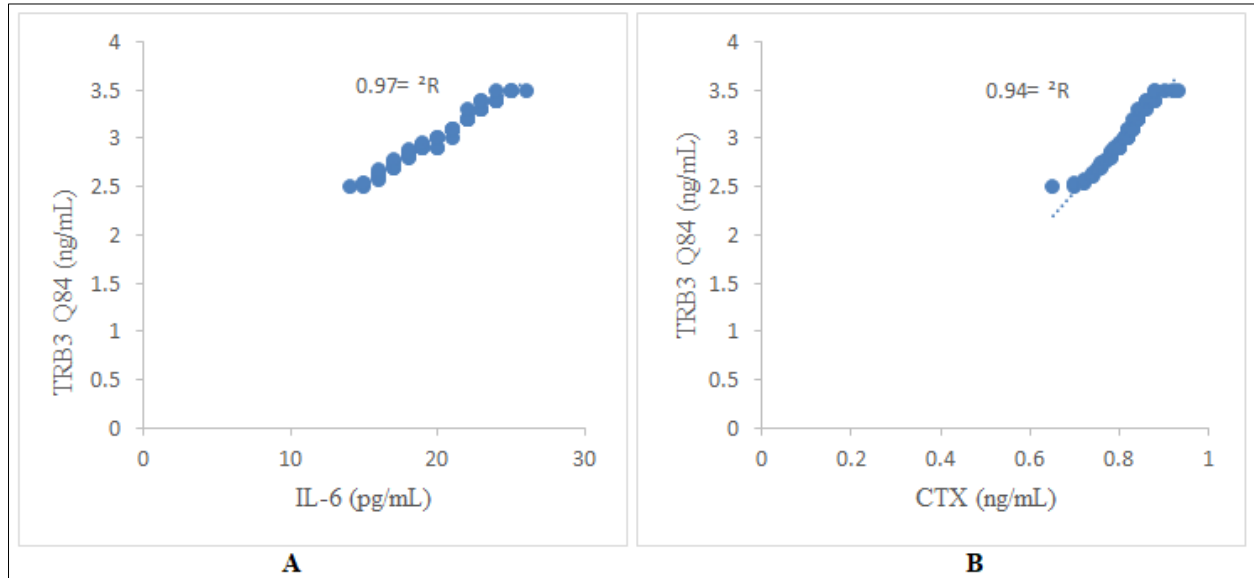


Figure 2: TRB3 Q84 levels correlation with A: IL-6 and B: CTX in stimulants dependence

The operating characteristic of receiver ROC curve for TRB3 Q84 was shown in Table 3, Figure 3, for detecting stimulants dependence a point of cut-off was

95%. The curve of - under area (AUC) was - 0.983, performance diagnostic was high. Sensitivity of TRB3 Q84 was 95% and a specificity was 100%.

Table 3: TRB3 Q84 analysis by (ROC) and (AUC) in diagnosing stimulants dependence

Variable	Group	Cut-off concentration %	Sensitivity %	Specificity %	AUC	Std. Error	95% CI of AUC	P-value
TRB3 Q84	Stimulants Dependence	95	95	100	0.983	0.015	0.953 - 1.000	0.001

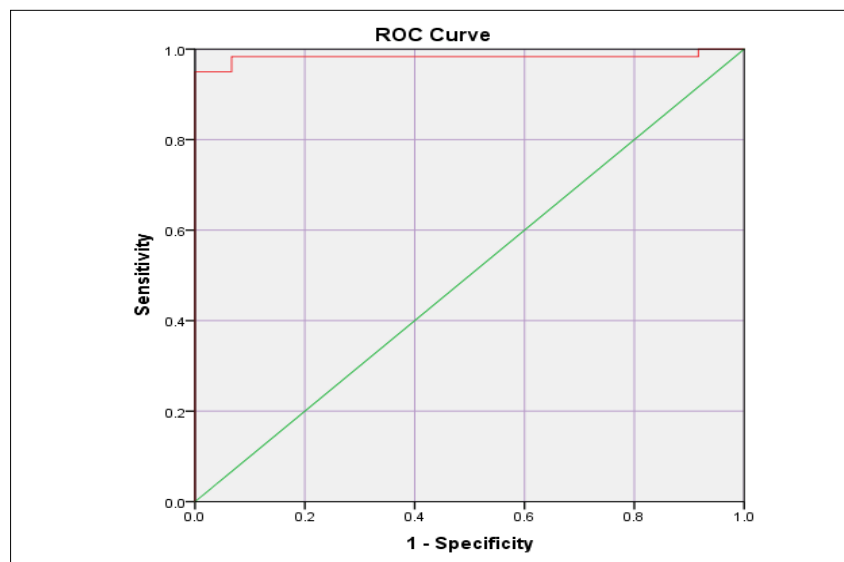


Figure 3: TRB3 Q84 analysis by (ROC) in stimulants dependence diagnosing

4. DISCUSSION

An increase with significant of TRB3 Q84, IL-6 and CTX levels among individuals with stimulants dependence. In these patients, levels of TRB3 Q84 was correlated a significant with IL-6 and CTX by a strong positively. Our results can be interpreted as following: in stimulants dependence, the infection triggers an acute

inflammatory response characterized by cytokines of pro-inflammatory elevation levels such as TNF- α and IL-6. This inflammatory environment stimulates immune cells to secrete large amounts of TRB3 Q84, both within the peripheral immune tissues - and - central nervous system. Additionally, chronic inflammation influences the balance of bone cells by promoting the activity of

osteoclasts, which break down bone, while inhibiting osteoblasts, the cells responsible for bone formation. This shift in cellular activity accelerates the progression of osteoporosis, contributes to increased production of factors that enhance bone resorption, leading to a loss of bone mass. Consequently, in stimulants dependence individuals, levels of TRB3 Q84 possible to use as early indicator for osteoporosis.

Liver graphic assessments indicated better liver function in individuals with lower plasma TRB3 Q84 levels [26, 27]. In a patients study involving infarction of liver, those who did not survive exhibited significantly higher plasma TRB3 Q84 levels compared to survivors. Furthermore, an experimental study using anti-TRB3 Q84 antibodies in mice demonstrated reduced infiltration of neutrophils and macrophages into the liver, along with a decrease in liver tissue damage [28, 29]. Its levels in the left liver tissue have been found to increase significantly within the first week post-infarction. Moreover, patients with reduced ejection had elevated TRB3 Q84 concentrations [30-31]. During liver infarction, TRB3 Q84, a glycoprotein secreted by neutrophils is released in large amounts. While this process may help limit the extent of tissue damage, excessive neutrophil accumulation can hinder proper liver repair. Elevated serum TRB3 Q84 levels in liver patients have been associated with worse clinical outcomes [32-35]. Elevated TRB3 Q84 concentrations have been observed in liver patients. Following an infarction, neutrophils infiltrate the damaged liver tissue and trigger inflammatory responses [36-38].

Additionally; inhibiting the activity by TRB3 Q84 that may exert anti-inflammatory effects in adipose tissue, thereby counteracting TNF-alpha-mediated inflammation. Previous research found that increased TRB3 Q84 mRNA expression, which in turn activated in obese mice [39, 40]. It is thought to be involved in the regulation of a key regulator of adipogenesis and lipogenesis [41-43]. Several studies suggest that the useful predictor for metabolic complications associated with obesity is measuring TRB3 Q84 concentrations in the bloodstream [44-46]. Furthermore, the risk factor for disturbances in lipid metabolism; hypertension - and insulin resistance - has been identified by elevated TRB3 Q84 [47, 48]. In individuals with; disease of nonalcoholic fatty liver - and - obesity have been observed elevated levels of TRB3 Q84 [49].

5. CONCLUSION

In this our study, levels of TRB3 Q84 was correlated a significant with IL-6 and CTX by a strong positively. These findings suggest that TRB3 Q84 levels can be used as indicator for the early diagnosis of osteoporosis in patients with stimulants dependence.

6. Acknowledgement: The staff at Diwaniyah Teaching Hospital in Al-Diwaniyah, Iraq, as well as the patients were sincerely thanked.

7. Conflict of Interests: None.

8. Funding: Self.

REFERENCES

1. Abio A, Neal K and Beck C. An epidemiological review of changes in stimulants dependence during the last 100 years. *Pathogens and Global Health* 2013; 10(5): 20-30.
2. Stankowski R, Kloner R and Rezkalla S, Cardiovascular consequences of stimulants dependence, *Trends Cardiovasc Med*. 2014, 25: 26-517.
3. Hyman S and Malenka R, Stimulants dependence and the brain: The neurobiology of compulsion and its persistence, *Nat Rev Neurosci*, 2001, 10: 695-703.
4. Riezzo I, Fiore C and Carlo D, Side effects of stimulants dependence: multiorgan toxicity and pathological consequences, *Curr Med Chem*, 2012, 19: 46-562.
5. Andreozzi F, Formoso G and Prudente S, TRIB3 R84 variant is associated with inflammatory production in human endothelial cells, *Arterioscler. Thromb. Vasc Biol*, 2008; 28:1355-1360.
6. Houghland J. Seizures in Adults with stimulants dependence. *The Journal of Emergency Medicine* 2008; 35(16): 13-36.
7. Brandt C. Experimental studies of pneumococcal stimulants dependence. *Danish medical bulletin* 2010; 57(19): 14-25.
8. Turner M, Nedjai B, Hurst T and Pennington D. Cytokines and chemokines: At the crossroads of cell signalling and inflammatory disease. *Biochimica et Biophysica Acta (BBA) - Molecular Cell Research* 2014; 43(25): 12-32.
9. Gong H, Wang Z and Jiang H, TRIB3 functional Q84R polymorphism is a risk factor for metabolic syndrome and inflammatory, *Diabetes Care*, 2009; 32:1311-1313.
10. Prudente S, Dallapiccola B and Pellegrini F, Genetic prediction of common diseases. Still no help for the clinical inflammatory, *Nutr. Metab. Cardiovasc. Dis*, 2012; 22:929-936.
11. Prudente S, Bailetti D and Mendonca C, Infrequent TRIB3 coding variants and inflammatory disease, *Atherosclerosis*, 2015; 242:334-339.
12. Yokoyama T, Toki T and Aoki Y, Identification of TRIB1 R107L gain- of-function mutation in human acute megakaryocytic leukemia, *Blood*, 2012; 119:2608-2611.
13. Koyama T, Mendes C and Mirth C, Mechanisms regulating nutrition-dependent developmental plasticity through organ-specific effects in insects,

- Front. Physiol*, 2013 ; 4:263-275.
14. Formoso G, Tomo P and Andreozzi F, The TRIB3 R84 variant is associated with increased inflammatory in vivo and with enhanced MAPK signalling in human endothelial cells, *Cardiovasc. Res*, 2011; 89:184-192.
 15. Prudente S, Hribal M and Flex E, The functional Q84R polymorphism of mammalian Tribbles homolog TRB3 is associated with inflammatory disorders, *Diabetes*, 2005; 54:2807-2811.
 16. Dobens L and Bouyain S, Developmental roles of tribbles protein family members, *Dev. Dyn*, 2012; 241: 1239-1248.
 17. Romas E, Gillespie M and Martin T. Involvement of receptor activator of NFκB ligand and tumor necrosis factor-α in bone destruction in rheumatoid arthritis. *Bone* 2002; 20(6): 14-20.
 18. Krishna G, Bir S and Kevil C, Endothelial Dysfunction and Inflammation: Effects on Angiogenesis, Vascular Remodeling, and Wound Healing, *Int. J. Vasc. Med*, 2012; 20:30-36.
 19. Das R and Dobens L, Conservation of gene and tissue networks regulating inflammatory signalling in flies and vertebrates, *Biochem. Soc. Trans*, 2015; 43:1057-1062.
 20. Mundy GR. Osteoporosis and inflammation. *Nutr Rev* 2007; 25(14): 11-35.
 21. Karaolanis G, Moris D and Palla V. TRB3 Q84 as a Biomarker- Does It Apply in Abdominal Aortic Aneurysms- A Review of Literature. *Indian J. Surg* 2015; 7(3): 17-29.
 22. Ramos P, Madrigal J and Vegade M. Increased plasma levels of TRB3 Q84, a marker of neutrophil activation- in patients with abdominal aortic aneurysm. *Atherosclerosis* 2012; 22(12):16-27.
 23. Teitelbaum S. Osteoclasts: what do they do and how do they do it. *Am J Pathol* 2007; 17(12): 15-31.
 24. Tarin C, Fernandez C, Burillo E and Pastor C. TRB3 Q84 or blockade protects against aortic abdominal aneurysm development in mice. *Cardiovasc. Res* 2016; 11(2): 23-36.
 25. Han X, Zhang S, Chen Z and Adhikari B. Cardiac biomarkers of heart failure in chronic kidney disease. *Clin. Chim. Acta* 2020; 10(2): 11-24.
 26. Bolignano D, Basile G, Parisi P and Coppolino G. Increased plasma TRB3 Q84 levels predict mortality in elderly patients with chronic heart failure. *Rejuvenation Res* 2009; 12(7): 14-27.
 27. Oikonomou E, Tsalamandris S and Karlis D. The association among biomarkers of renal and heart function in patients with heart failure: The role of TRB3 Q84. *Biomark. Med* 2018; 12(3): 13-25.
 28. Yndestad A, Landrø L, Ueland T, Dahl C and Flo T. Increased systemic and myocardial expression of TRB3 Q84 in clinical and experimental heart failure. *Eur. Heart J* 2009; 30(15): 12-27.
 29. TeBoekhorst B, Bovens S, Hellings W and vanderKraak P. Molecular MRI of murine atherosclerotic plaque targeting TRBR Q84: A protein associated with unstable human plaque characteristics. *Cardiovasc. Res* 2011; 19(8): 18-45.
 30. Martínez E, Buonafina M, Boukhalfa I, Ibarrola J and Fernández A. Aldosterone Target TRB3 Q84 Is Involved in Cardiac Remodeling After Myocardial Infarction Through NFκB Pathway. *Hypertension* 2017; 50(11): 11-31.
 31. Freitas I, Lima N and Silva G. Novel biomarkers in the prognosis of patients with atherosclerotic coronary artery disease. *Rev. Port. Cardiol* 2020; 39(6): 12-30.
 32. Cheng L, Xing H and Mao X. Lipocalin-2 promotes m1 macrophages polarization in a mouse cardiac ischaemia-reperfusion injury model. *Scand. J. Immunol* 2015; 21(11): 18-39.
 33. Libby P, Ridker P and Hansson G. Progress and challenges in translating the biology of atherosclerosis. *Nature* 2011; 43(17): 25-33.
 34. Oberoi R, Bogalle E, Matthes L and Schuett H. Atherosclerotic Processes and Is Elevated in Patients with Coronary Artery Disease. *PLoS ONE* 2015; 10(3): 14-42.
 35. Leclercq A, Houard X, Philippe M and Ollivier V. Involvement of intra plaque hemorrhage in atherothrombosis evolution via neutrophil protease enrichment. *J. Leukoc. Biol* 2007; 12(10): 14-33.
 36. Van der Wal A, Becker A, van der Loos C and Das P. Site of intimal rupture or erosion of thrombosed coronary atherosclerotic plaques is characterized by an inflammatory process irrespective of the dominant plaque morphology. *Circulation* 1994; 89(36): 24-50.
 37. Yan Q, Yang Q, Mody N and Graham T. The TRB3 Q84 is regulated by obesity and promotes insulin resistance. *Diabetes* 2007; 56(25): 35-40.
 38. Ricote M and Glass C. PPARs and molecular mechanisms of transrepression. *Biochim. Biophys. Acta* 2007; 71(26): 25-46.
 39. Lin Y, Rajala M, Berger J and Moller D. Hyperglycemia-induced production of acute phase reactants in adipose tissue. *J. Biol. Chem* 2001; 26(4): 20-32.
 40. Zhang J, Wu Y, Zhang Y, and Leroith D. The role of TRB3 Q84 in the regulation of inflammation in adipocytes and macrophages. *Mol. Endocrinol* 2008; 22(16): 14-39.
 41. Jang Y, Lee J, Wang Y, Sweeney G. Emerging clinical and experimental evidence for the role of TRB3 Q84 in metabolic syndrome. *Clin. Exp. Pharmacol. Physiol* 2012; 39(14): 19-33.
 42. Liu J, Song E, Xu A, Berger T and Mak T. TRB3 Q84 deficiency prevents endothelial dysfunction associated with dietary obesity: Role of cytochrome P450 2C inhibition. *Br. J. Pharmacol* 2012; 65(20): 21-35.
 43. Bhusal A, Rahman M, Lee W and Bae Y. Paradoxical role of TRB3 Q84 in metabolic disorders and neurological complications. *Biochem. Pharmacol* 2019; 16(11): 26-38.

44. Law I, Xu A and Lam K. TRB3 Q84 deficiency attenuates insulin resistance associated with aging and obesity. *Diabetes* 2010; 59(18): 12-47.
45. Çelik T, Altekin E, Kenesari Y, Duman M and Arslan N. Evaluation TRB3 Q84 in pediatric patients with acute rotavirus gastroenteritis and dehydration. *Ital. J. Pediatr* 2013; 39(22): 21-45.
46. Wu G, Li H, Zhou M, Fang Q, Bao Y, Xu A and Jia W. Mechanism and clinical evidence of TRB3 Q84 and adipocyte fatty acid-binding protein linking obesity and atherosclerosis. *Diabetes Metab. Res. Rev* 2014; 30(14): 15-52.
47. Wang Y, Lam K, Kraegen E, Sweeney G, Zhang J and Tso A. TRB3 Q84 is an inflammatory marker closely associated with obesity, insulin resistance, and hyperglycemia in humans. *Clin. Chem.* 2007; 53(34): 21-
48. Landrø L, Damås J, Flo T, Heggelund L, Ueland T, Tjønnfjord G, Espevik T, Aukrust P and Frøland S. Decreased serum TRB3 Q84 levels in human immunodeficiency virus-infected patients: Increase during highly active anti-retroviral therapy. *Clin. Exp. Immunol* 2008; 15(12): 13-28.
49. Vijay M, Gentsch J, Kaiser W, Borregaard N, Offermann M, Neish A and Gewirtz A. Protein kinase R mediates intestinal epithelial gene remodeling in response to double-stranded RNA and live rotavirus. *J. Immunol* 2005; 74(22): 13-25.