

THE EFFECT OF *Toxoplasma gondii* PROFILIN EXPOSURE ON HDL LEVEL IN *Rattus norvegicus* WISTAR STRAIN





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INTRODUCTION - BACKGROUND

- In Indonesia, the national prevalence of obesity aged over 15 years is 10.3% in 2011 and the prevalence in women and men aged > 18 years in 2013 is 32.9% and 19.7%, respectively. (Indonesian Basic Health Research, 2013)
- Increased prevalence of obesity has led to some hypothetical explanations because lifestyle and diet modification do not fully explain the phenomenon.
- Some studies suggest that obesity is triggered by the presence of chronic inflammation which probably is a response to particular infection. (Milovanovic *et al.*, 2009); (Portugal, 2008)

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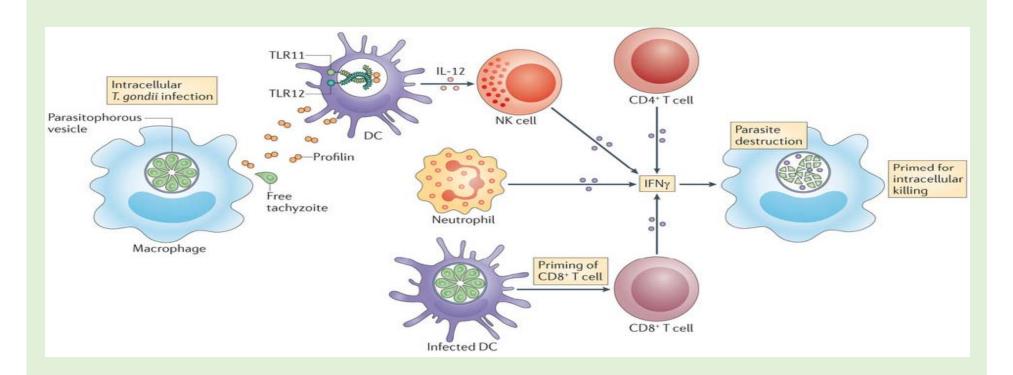
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The level of chemerin and adipocyte fatty acid binding protein in *Toxoplasma gondii* seropositive obese individuals



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T. gondii, lipid metabolism and obesity

- There is a possible association between *T. gondii* and obesity. (Reeves et al, 2013) Individuals with positive *T. gondii* serology had twice the odds of being obese compared to seronegative individuals.
- ➤ Host lipids have recently been implicated in the pathogenesis of *Toxoplasma gondii* infection. It has been shown that cholesterol uptake from the host plasma membrane is essential for parasite replication *in vitro*. (Milovanovic *et al.*,2009)

- Toxoplasma gondii has profilin-like protein which will be recognized by toll-like receptor (TLR-11) of the natural immune system and followed by inflammation of the host cell. (Hermes-Uliana et al, 2011)
- T gondii profilin have capability to bind with TLR-11 and cause an increase of inflammatory cytokine (TNF- α , IL-10, chemerin, aFABP) in adipocyte culture. (Sudjari et al, 2015)
- Toxoplasma gondii alters host cell metabolism for entry and replication and uses host metabolic products for its own metabolic pathways. (Portugal, 2008)
- A significant decrease in HDL and total Cholesterol was noted in infected *T* gondii mice at day 14 and persisted to day 42. Conversely, LDL was increased at day 42. (Milovanovic, 2009)

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Floral research

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The levels of *Toxoplasma gondii* profilin and adiponectin in obese patients complicated with or without metabolic syndrome as compared to non-obese patients

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Problem Statement

➤ Does *Toxoplasma gondii* profilin have effect on body weight and HDL level in *Rattus norvegicus* Wistar Strain?



General Objective

➤To reveal the effect of to *Toxoplasma* gondii profilin exposure on body weight and HDL level in *Rattus norvegicus* Wistar Strain

Specific Objectives

- ➤ To know the bodyweight and HDL level in normal *Rattus Norvegicus* Wistar Strain
- To know the bodyweight and HDL level in Rattus norvegicus Wistar Strain given normal diet that are exposed to Toxoplasma gondii profilin.
- To know the bodyweight and HDL level in Rattus norvegicus Wistar Strain given hypercaloric diet that are exposed to Toxoplasma gondii profilin.

Research Method

Research Design

The study was an experimental study

Site/Time of Research

 The research was conducted in Laboratory of Pharmacology and Clinical Pathology, Faculty of Medicine, Universitas Brawijaya from January to April 2017

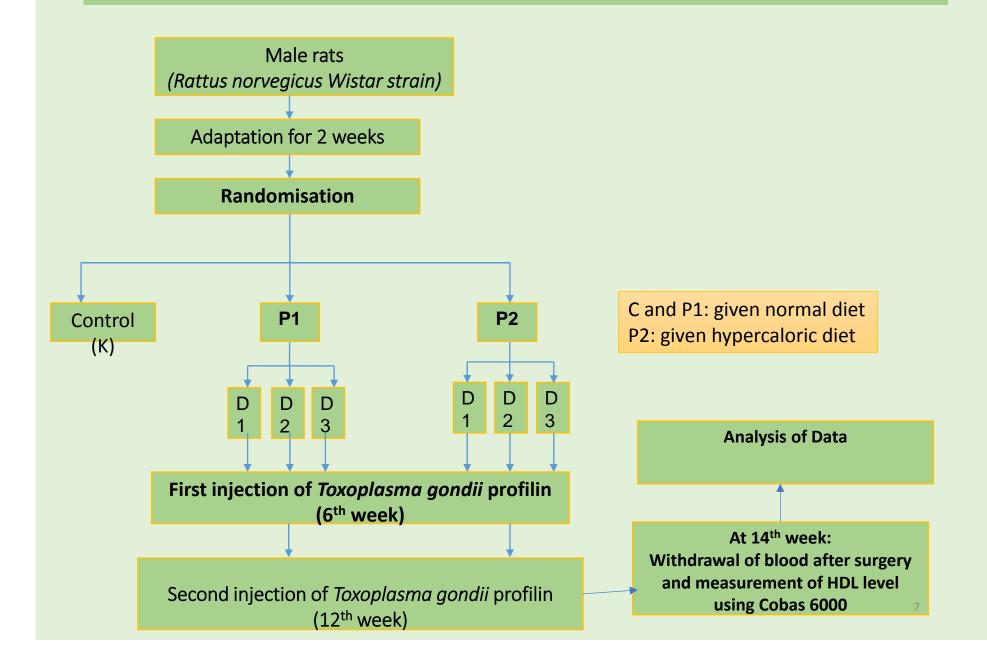
Population and Sample

 Male, white Rattus norvegicus Wistar strain obtained from Parasitology Laboratory Medical Faculty of Brawijaya University.

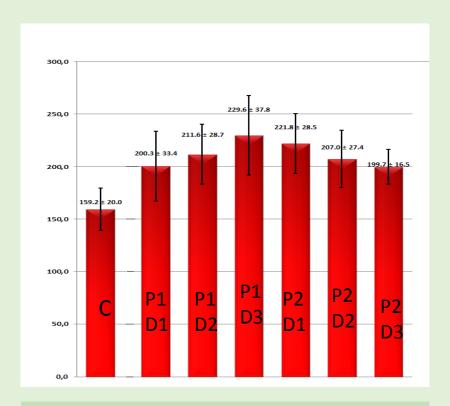
Working Definition

- Profilin dose given based on preliminary study
- HDL was determined by using enzymatic method with Cobas 6000

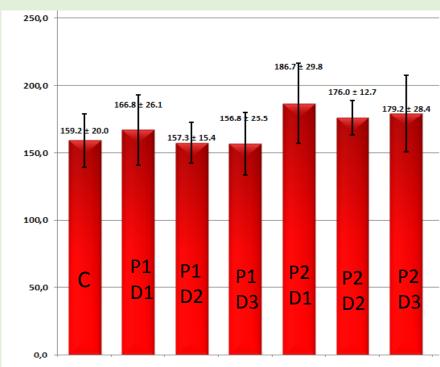
STUDY FLOWCHART



Results



Mean of body weight after once injection of *T gondii* profilin.



Mean of body weight after twice injection of *T gondii* profilin.

HDL Level

Mean of Body weight after once injection of *T gondii* profilin

Group	HDL	Standard Deviation	р
K	36.0000	9.38083	
D1	54.0000	21.57159	
D2	46.0000	10.13246	0.07
D3	49.0000	19.28730	0.07
D4	40.8333	4.07022	$(\alpha = 0.05)$
D5	46.1667	10.53407	
D6	40.1667	6.21021	

Mean od Body weight after twice injection of *T gondii* profilin

Group	HDL	Standard Deviation	р
K	36.0000	9.38083	
D1	46.4000	14.39792	
D2	37.0000	16.14001	0.252
D3	34.2000	5.44977	0.232
D4	43.2000	2.58844	(α=0.05)
D5	49.4000	4.77493	
D6	43.2000	20.77739	

Note:

K: normal group; D1: normal diet + profilin 15ug/mL; D2: normal diet + profilin 30 ug/mL; D3: normal diet + profilin 45ug/mL; D4: hypercaloric diet + profilin 15ug/mL; hypercaloric diet + profilin 30 ug/mL; D6:

hypercaloric diet + profilin 45ug/mL

Correlation test

Correlation test	р	r	r²	
Profilin dose and HDL	Normal Diet	0.259	0.338	0.114
level (Once injection)	Hypercaloric diet	0.257	-0.355	0.126
Profilin dose and HDL	Normal Diet	0.072	-0.411	0.169
level (twice injection)	Hypercaloric diet	0.341	-0.275	0.076
HDL level and body	Normal Diet	0.253	0.377	0.142
weight (Once injection)	Hypercaloric diet	0.246	-0.289	0.084
HDL level and body	Normal Diet	0.472	-0.201	0.040
weight (twice injection)	Hypercaloric diet	0.629	-0.136	0.185

Discussion

Effect of T gondii profilin exposure on body weight

- There is a significant difference of body weight after once injection of *T* gondii profilin, but no significant difference between 3 doses and also no significant difference after twice injection
- There was no significant correlation between profilin doses and body weight, the higher profilin doses the lower body weight
 - → *T gondii* profilin could increase body weight, but still need further research to optimize dose and time of profilin injection.
 - → upon infection with Toxoplasma, uptake of Chl from serum LDL by proliferating parasites triggers mobilization of Chl from the liver to the periphery.
 - → Due to the use of Chl in peripheral tissues by *Toxoplasma*, less remains to be returned to the liver and thus decreased serum HDL.
 - → The decrease at day 14 (but not at day 7) coincides with the time when the parasite burden has increased sufficiently to require significant LDL acquisition, measurable (in a host with low LDL and high HDL) only by HDL decrease, (Milovanović et al., 2009),

Effect of *T gondii* profilin on HDL level

- There is no significant difference of HDL level neither once nor twice injection of *T gondii* profilin.
- No significant correlation between *T gondii* profilin doses and HDL level neither in normal diet nor hypercaloric diet.
 - → *T gondii* altered lipid metabolism and decrease body weight, but it still need further research especially on the optimal dose for profilin injection and other factor?

Correlation between body weight and HDL after injection of T gondii profilin.

- There was no significant and very weak correlation between body weight and HDL level neither in once nor twice injection.
 - →The role of *T gondii* profilin on body weight need further research,
 - → It can be postulated that profilin cause obesity and adipocyte dysfunction but didn't altered lipid metabolism → other ways?

Conclusion & suggestions

CONCLUSION

- The normal diet + twice injection of T gondii profilin 45 μg/mL, has the lowest average weight.
- The normal diet + once injection of T gondii profilin 45 μg/mL, has the highest average weight
- The normal diet + twice injection of T gondii profilin 45 μg/mL, has the lowest average HDL level
- The normal diet + once injection of T gondii profilin 15 μg/mL, has the highest average HDL level
- No significant difference and correlation between doses and body weight also HDL level both once and twice injection of *T gondii* profilin.

SUGGESTION

 Further research by injection of Toxoplasma gondii (not only profilin of T gondii) to make sure that there was relationship between T gondii infection and obesity.



