

THE 46TH ANNUAL CONFERENCE OF THE MALAYSIAN SOCIETY FOR BIOCHEMISTRY AND MOLECULAR BIOLOGY

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#### THE AMELIORATIVE EFFECTS OF TUALANG HONEY MEDIATED SILVER NANOPARTICLES ON HIPPOCAMPAL DAMAGES FOLLOWING KAINIC ACID ADMINISTRATION IN MALE RATS

**Hidani Hasim**<sup>1</sup>, Sirajudeen Kuttulebbai Naina Mohamed Salam<sup>\*2</sup>, Pasupuleti Visweswara Rao<sup>3,4</sup>, Sangu Muthuraju<sup>5</sup> and Mohd Asnizam Asari<sup>\*6</sup>

<sup>1</sup>Department of Chemical Pathology, School of Medical Sciences, Universiti Sains Malaysia, Health Campus, Kota Bharu, Kelantan, Malaysia; <sup>2</sup>Department of Basic Medical Sciences, Kulliyyah of Medicine, International Islamic University Malaysia, Kuantan, Pahang, Malaysia; <sup>3</sup>Department of Biochemistry, Faculty of Medicine and Health Sciences, Abdurrab University, Riau, Indonesia; <sup>4</sup>Center for International Collaboration and Research, Reva University, Rukmini Knowledge Park, Bangalore, India; <sup>5</sup>Department of Pharmacological and Pharmaceutical Sciences, College of Pharmacy, University of Houston, Houston, TX, United States; <sup>6</sup>Department of Anatomy, School of Medical Sciences, Universiti Sains Malaysia, Health Campus, Kota Bharu, Kelantan, Malaysia

\*Corresponding authors: asnizam@usm.my; knssiraj@iium.edu.my

Kainic acid (KA) was shown to be associated in the mechanism of excitotoxicity-induced neurodegeneration in the brain. Tualang honey (TH) was reported to have protection against neurodegeneration but no study has explored on its silver nanoparticles (THSN). Therefore, present study aimed to investigate the effects of THSN on glutathione status and hippocampal histology in KA-induced rats. Sprague-Dawley rats (n=42) were divided into seven groups such as control, THSN 10 mg, THSN 50 mg, KA alone, THSN 10 mg + KA, THSN 50 mg + KA and Topiramate + KA, and each group were pretreated orally with either distilled water, THSN (10 mg/kg or 50 mg/kg) or Topiramate (40 mg/kg), respectively, five times at 12 h intervals. Saline or KA (15 mg/kg body weight) were injected subcutaneously 30 min after last oral treatment. All animals were sacrificed 24 h post KA injection and the hippocampus was harvested for histological examination using cresyl violet staining. The reduced glutathione (GSH) and oxidized glutathione (GSSG) was determined using commercially available ELISA kits. The significant (p<0.05) decrease in the level of cresyl violet-positive cells in hippocampal CA3 in KA alone group was ameliorated by THSN (10 mg/kg) pretreatment group. Meanwhile, the KAinduced reduction in GSH:GSSG ratio in KA alone group was significantly (p<0.05) increased by both doses of THSN pretreatments. In conclusion, THSN showed potential protective effects by improving the glutathione status and reduce hippocampal cells injury in the rats after KAinduced.

Keywords: Silver nanoparticles, rat hippocampus, kainic acid, glutathione, cresyl violet



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<sup>1</sup>Department of Chemical Pathology, School of Medical Sciences, Universiti Sains Malaysia, Health Campus, 16150 Kubang Kerian, Kota Bharu, Kelantan, Malaysia <sup>2</sup>Department of Basic Medical Sciences, Kulliyyah of Medicine, International Islamic University Malaysia, Bandar Indera Mahkota, 25200 Kuantan, Pahang, Malaysia <sup>3</sup>Department of Biochemistry, Faculty of Medicine and Health Sciences, Abdurrab University, JI Riau Ujung No. 73, Pekanbaru 28292, Riau, Indonesia <sup>4</sup>Center for International Collaboration and Research, Reva University, Rukmini Knowledge Park, Kattigenahalli, Yelahanka, Bangalore, Karnataka, 560064, India <sup>5</sup>Department of Pharmacological and Pharmaceutical Sciences, College of Pharmacy, University of Houston, Houston 77240, TX, United States <sup>5</sup>Department of Anatomy, School of Medical Sciences, Universiti Sains Malaysia, Health Campus, 16150 Kubang Kerian, Kota Bharu, Kelantan, Malaysia

#### INTRODUCTION

Green synthesis of nanoparticles using plant-mediated process has been used for therapeutic and diagnostic purposes [1]. Recently, researchers have develop nanoparticles that can cross the blood-brain barrier [2]. Glutathione (GSH) is the most abundant antioxidant intracellular thiol in the brain [3]. It reacts with free radicals and protects cells from singlet oxygen, hydroxyl radical, and superoxide radical damage [3]. Tualang honey (TH), a potential natural antioxidant agent, has been shown to protect against neurodegenerative disorders [4,5]. Therefore, the present study aimed to explore the ameliorative effects TH mediated silver nanoparticles (THSN) on glutathione status and hippocampal histology in kainic acid (KA)-induced rats.

#### METHODOLOGY

Sprague Dawley male rats (n=42) were randomly divided into seven groups:

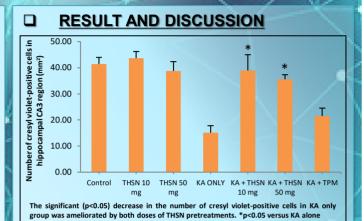
Group (1): control	Group (5): THSN 10 mg + KA
Group (2): THSN 10 mg	Group (6): THSN 50 mg + KA
Group (3): THSN 50 mg	Group (7): TPM + KA
Group (4): KA only	

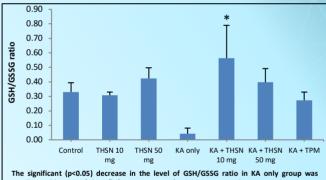
Each group were pretreated orally with either distilled water, AgNPs (10 mg/kg or 50 mg/kg) or Topiramate (TPM) (40 mg/kg), five times at 12 h intervals. Saline or KA (15 mg/kg body weight) were injected subcutaneously 30 min after last oral treatment.

All animals were sacrificed 24 h post KA injection and the hippocampus was harvested for histological examination using cresyl violet staining. The reduced glutathione (GSH) and oxidized glutathione (GSSG) was determined using commercially available ELISA kits.

#### CONCLUSION

THSN showed potential protective effects by improving the glutathione status and reduce hippocampal cells injury in the rats after KA-induced.





ameliorated by THSN (10 mg/kg) pretreatment. \*p<0.05 versus KA alone

It was reported that TH mediated silver nanoparticles exhibited remarkable antioxidant activity with 1,1-diphenyl-2-picrul hydrazyl and reducing antioxidant power values of 95.54  $\pm$  0.96 (%) and 1032.30  $\pm$  102.76  $\mu m$  Fe(II), respectively [6].

The improvement in GSH status and hippocampal CA3 morphology by THSN suggested that its antioxidant properties possibly increased the brain's endogenous defence against KA-induced oxidative stress.

#### ACKNOWLEDGEMENT

This research was financially supported by the Universiti Sains Malaysia under Research University (Individual) (RUI Grant No: 1001/PPSP/8012249).

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