





LEADING THE WORLD

INTERNATIONAL MULTI-AWARD WINNING INSTITUTION FOR SUSTAINABILITY







ROLE OF DBP & FGG PROTEINS IN OBESE SCHIZOPHRENIA: A PROTEOMIC STUDY

Siti Norain Mat Rasid¹, Nour El Huda Abd Rahim¹, Norbaiyah Mohamed Bakrim¹, Mohd Asyraf Abdull Jalil¹, Norlelawati A. Talib², Mohd Yusri Idorus³, Ahmad Nabil Md Rosli⁴, Mohd Arifin Kaderi, Norainin Sofiya Azman

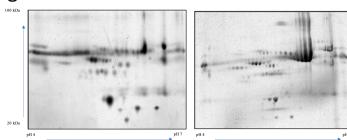
- ¹Department of Basic Medical Sciences, Kulliyyah of Medicine, IIUM, 25200 Kuantan, Pahang
- ² Department of Pathology and Laboratory Medicine, Kulliyyah of Medicine, IIUM, 25200 Kuantan, Pahang ³ Institute of Medical Molecular Biotechnology, Faculty of Medicine, UiTM, 47000 Sungai Buloh, Selangor
 - Department of Psychiatry, Kulliyyah of Medicine, IIUM, 25200 Kuantan, Pahang
- ⁵ Department of Biomedical Science, Kulliyyah of Allied Health Sciences, IIUM, 25200 Kuantan, Pahang

INTRODUCTION

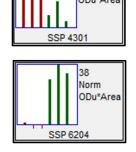
- Both schizophrenia and obesity are complicated conditions that can have a major negative impact on an individual's health and quality of life.
- Obesity heightens the health risks and contributes to elevated morbidity & mortality in schizophrenia [1,2].
- Even though each condition has been thoroughly researched on its own, the intersection of obesity & schizophrenia remains underexplored, particularly at the proteomic level [3,4].
- This study aims to compare the protein profiles of obese schizophrenia and obese non-schizophrenia subjects. Then, the role of the differently expressed proteins was determined using a publicly available protein database and literature search.

RESULTS

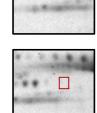
Representative of two-dimensional electrophoresis (2-DE) gel images of plasma proteomic profiling:



• Representative patterns of significantly differently expressed protein spots (pvalue < 0.05):







 The proteins identified through Liquid Chromatography with Tandem Mass Spectrometry (LC-MS/MS) were:

oury (20 100) 1000.					
ID	Protein	Accession number	Molecular Weight	Isoelectric Point	Sequence coverage
SSP 4301	Vitamin D binding protein (DBP)	P02774	55.1 kDa	5.74	45 %
SSP 6204	Fibrinogen gamma chain (FGG)	P02679	51.5 kDa	5.62	65 %

DBP was underexpressed, while FGG was overexpressed in obese schizophrenia compared to obese non-schizophrenia.

CONCLUSION

- DBP & FGG play significant roles in linking obesity and schizophrenia.
- The underexpression of DBP suggests reduced vitamin D bioavailability & exacerbates metabolic disturbances.
- The overexpression of FGG indicates increased inflammation & coagulation tendencies, heightening cardiovascular & metabolic risks.

ACKNOWLEDGEMENT

analysis. Vol. 121, Progress in Neuro-Psychopharmacology and Biological Psychiatry. 2023.

 We would like to acknowledge the Ministry of Higher Education Malaysia for a research grant (FRGS/1/2022/SKK05/UIAM/03/1), Kulliyyah of Medicine, International Islamic University Malaysia.

REFERENCES

- Mazereel V, Detraux J, Vancampfort D, van Winkel R, De Hert M. Impact of Psychotropic Medication Effects on Obesity and the Metabolic Syndrome in People With Serious Menta llness. Vol. 11, Frontiers in Endocrinology. Frontiers Media S.A.; 2020. Rodrigues JE, Martinho A, Santa C, Madeira N, Coroa M, Santos V, et al. Systematic Review and Meta-Analysis of Mass Spectrometry Proteomics Applied to Human
- Assess Potential Biomarkers of Schizophrenia. Vol. 23, International Journal of Molecular Sciences. MDPI; 2022. Aleksandrova K, Egea Rodrigues C, Floegel A, Ahrens W. Omics Biomarkers in Obesity: Novel Etiological Insights and Targets for Precision Prevention. Vol. 9. Current obesity reports
- Setayesh L, Casazza K, Moradi N, Mehranfar S, Yarizadeh H, Amini A, et al. Association of vitamin D-binding protein and vitamin D3 with insulin and homeostatic model assessment (HOMA-IR) in overweight and obese females. BMC Res Notes. 2021;14(1).
 6. Setayesh L, Amini A, Bagheri R, Moradi N, Yarizadeh H, Asbaghi O, et al. Elevated plasma concentrations of vitamin d-binding protein are associated with lower high-density lipoprotein
- and higher fat mass index in overweight and obese women. Nutrients. 2021;13(9). Albiñana C, Boelt SG, Cohen AS, Zhu Z, Musliner KL, Vilhjálmsson BJ, et al. Developmental exposure to vitamin D deficiency and subsequent risk of schizophrenia. Schizophr Res
- Wimalawansa SJ. Associations of vitamin D with insulin resistance, obesity, type 2 diabetes, and metabolic syndrome. Vol. 175, Journal of Steroid Biochemistry and Molecular Biology
- Purdy JC, Shatzel JJ. The hematologic consequences of obesity. Vol. 106, European Journal of Haematology. 2021
- Patlola SR, Donohoe G, McKernan DP. The relationship between inflammatory biomarkers and cognitive dysfunction in patients with schizophrenia: A systematic review and meta

METHODS

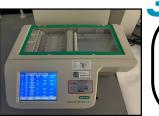
Sample Pooling:

Plasma from 10 subjects from each group were pooled.

Sample Preparation:

Protein extraction and preparation for 2-DE





1st Dimension of 2-DE:

Protein separation according isoelectric point using IPG strip for Isoelectric Focusing

> 2nd Dimension of 2-DE: Protein separation according molecular weight using SDS-PAGE





Protein Analysis:

Gel scanning to capture images of separated protein spots then analysed using PDQuest software

> Protein Identification: Protein spots of interest were excised from gel and identified by LC-MS/MS

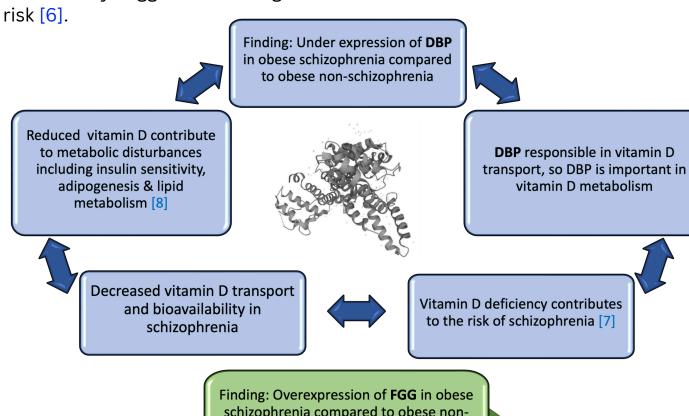


Role of Protein:

Verification of identified protein & its role using UniProtKB database

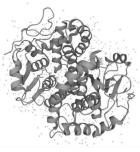
DISCUSSION

- DBP has been studied in relation to obesity, with a notable association between DBP levels and various metabolic factors in obesity.
- A study demonstrated the protective role of DBP in obesity specifically due to improved insulin sensitivity [5].
- Other study suggested that higher DBP level are associated with increase obesity



schizophrenia compared to obese nonschizophrenia

Obese schizophrenia exhibit more pronounced inflammation compared to those with obesity alone.



Obesity is linked to chronic low-grade inflammation, with elevated fibrinogen levels increasing inflammation [9]

Schizophrenia has been associated with hypercoagulable state and increased inflammatory markers [10,11]

- This suggests FGG as a potential biomarker for assessing the severity of inflammation and coagulation disturbances in this group.
- Therefore, it is important to target inflammation and coagulation pathways in managing the cardiovascular risks associated with obesity and schizophrenia.