

BAB V

KESIMPULAN DAN SARAN

A. KESIMPULAN

1. Ekstrak etanol daun sirsak dengan dosis 100, 200, dan 400 mg/KgBB dapat memperbaiki memori spasial dan menurunkan kadar malondialdehid, pada mencit model demensia yang diinduksi plumbum asetat
2. Pemberian ekstrak etanol daun sirsak pada dosis 400 mg/KgBB memberikan aktivitas yang lebih baik dari seluruh aspek pengujian.
3. Pemberian ekstrak etanol daun sirsak dapat menurunkan jumlah sel piramidal pada area CA1 dan CA2-CA3 yang mengalami kerusakan pada mencit model demensia yang diinduksi dengan plumbum asetat.

B. SARAN

1. Perlu dilakukan pengukuran aktivitas Glutathion Peroksidase (GPx) otak mencit.
2. Perlu dilakukan pengujian *moris water maze* dengan perhitungan panjang lintasan mencit.
3. Perlu dilakukan penelitian tentang efek ekstrak etanol daun sirsak terhadap total status antioksidan otak mencit *demensia* yang diinduksi plumbum asetat.
4. Perlu *Morris water maze* dengan n yang lebih besar (n = 10)
5. Perlu uji *behavior* lain

DAFTAR PUSTAKA

- Abbas, MA.,El-Haleem MRA, 2011, Evaluation of monosodium glutamate induced neurotoxicity in adult male albino rats. *Journal of American Science*, 7 (8), 264-76
- Alrawaiq N.S and Abdullah A.2014. A Review of Flavonoid Kuersetin: Metabolism, Bioactivity and Antioxidant Properties. *International Journal of PharmTech Research*. Vol.6 (3) : 933-941.
- Alvin, V., Terry, Jr. 2009. Methods of Behavior Analysis in Neuroscience, 2nd edition : Chapter13 Spatial Navigation (Water Mask) Tasks. *BocaRaton (FL) : CRC Press*
- Alzheimer's & Dementia Association.2018. Alzheimer's disease facts and figures. *Alzheimer's & Dementia* 14: 367-429
- Alzheimer's Disease International. 2015. *Dementia: a public health problem*. Geneva: World Health Organization
- Amelia, F.E., Angeline, K & Wahyu. 2012. Tablet salut enteric ekstrak etanol daun sirsak (*Annona muricata* L.) sebagai antikanker kolon yang potensial. Skripsi Sarjana. Yogyakarta: Universitas Gadjah Mada
- Anonim, 2012, *Alzheimer's Disease Facts and Figures 2012, Factsheet March 2012*, www.alz.org/facts, Alzheimer's Association diakses 20 Juli 2018.
- Asmoro, Feni Tri., 2011, Teh herbal lempuyang gajah sebagai penurun lipid peroksida darah tikus hiperkolesterolemia. *Jurnal Pangan dan Hasil Pertanian*.Institut Pertanian Bogor, Bogor.
- Arifuddin, Aswiyanti Asri, Elmantns, 2016, Efek Pemberian Vitamin C terhadap gambaran hispatolpgi hati tikus wistar yang terpapar timbal asetat, *Jurnal Kesehatan Andalas*, vol 1(1) : 227-228
- Arimura, N. dan Kaibuchi, K., 2005, *Key regulators in neuronal polarity*, *Neuron*,**48**, 881–884
- Arjadi F, Soejono SK, Maurits LS, Pangestu M. 2014. Jumlah Sel Piramidal CA3 Hipokampus Mencit Putih Jantan Pada Berbagai Model Stres Kerja Kronik. *MKB*. 46(4):197–202.
- Arkhaesi, N. 2008. Kadar Malaondialdehyde (MDA) Serum sebagai indikator Prognosis Keluaran pada Sepsis Neonatorum. Semarang: Thesis Universitas Diponegoro

- Asni, E., Harahap, I.P., dkk, 2009. Pengaruh Hipoksia Berkelanjutan Terhadap Kadar Malondialdehid, Glutathion Tereduksi, dan Aktivitas Enzim Katalase Ginjal Tikus. *Majalah Kedokteran Indonesia* Vol. 59 (1) : 23-24. Jakarta: IDI
- Baddeley A. 2003. Working memory: Looking back and looking forward. *Nature Reviews Neuroscience*. 4(10):829–39.
- Balaban RS, Nemoto S, Finkel T. 2005. Mitochondria, oxidants, and aging. *Cell* 120:483- 495
- Baroroh, H.N & Warsinah. 2009. Antiinflammatory activity of ethanolic extract of leaves of jarak pagar (*Jatropha curcas* L.) and neutrophils profile in rats foot induced Carrageenan. *Proceedings of International Conference On Medicinal Plants*. Surabaya, 22–23
- Christen, Y., 2000, Oxidative stress and Alzheimer disease, *Am. J. Clin. Nutr.*, **71**, 621S–9S.
- Chun Shi, Jun Liu, Fengming Wu and David T. Yew . Ginkgo biloba Extract in Alzheimer's Disease: From Action Mechanisms to Medical Practice. *International Journal of Molecular Sciences*. *Int. J. Mol. Sci.* 2010, *11*, 107-123
- Dalle-Done I, Rossi R, Colombo R, Giustarini D, Milzani A. *Biomarkers of Oxidative Damage in Human Disease*. *Clin Chem* 2006;52:601-23
- Departemen Kesehatan Republik Indonesia., 1979. *Materia Medika Indonesia*, Jakarta
- Depkes RI, 1995. *Farmakope Indonesia*, Edisi IV, 112, Departemen Kesehatan Republik Indonesia, Jakarta.
- Dogru, E., Gumusbas, U., Kara, F., 2003, Individual variation in the spatial reference and working memory assessed undreallothetic and idiothetic orientation cues in rat, *Acta Neurobiol. Exp.*, **63**, 17-23.
- Dringen, R., J. M. Gutterer and J. Hirrlinger. 2000. Glutathione Metabolism in Brain: Metabolic Interaction Between Astrocytes and Neurons in The Defense Against Reactive Oxygen Species. *Eur. J. Biochem* 267: 4912-4916
- Durtewitz, D., Seamans, J.K., Sejnowski, T.J., 2000, Neurocomputational models of working memory, *Nat. Neurosci.*, **3**, 1192–1198
- Endeoga, H. O., and A. Gomina. 2000. Nutritional values of some nonconventional leafy vegetables of Nigeria, *J. Econ, Taxon, Bot*, 24
- Ercal N, Gurer H, Aykin-Burns N. 2001. Toxic metals and oxidative stress. Part 1.

- Mechanisms involved in metal induced oxidative damage. *Curr Top Med Chem.* 1 (6): 529–39
- Eriksson J, Vogel EK, Lansner A, Bergster F, Nyberg L. 2015. *Neurocognitive Architecture of Working Memory*. *Neuron.* 88(1):33-46.
- Fajariyah, S., Utami, E.T & Arisandi, Y. 2010. Efek pemberian estrogen sintetis (Diethylstilbestrol) terhadap struktur hepar dan kadar SGOT dan SGPT pada mencit (*Mus musculus*) betina strain Balb/C. *J Ilmu dasar* 11(1): 76–82.
- Flora G, Gupta D, Tiwari A. 2012. Toxicity of Lead: A Review With Recent Updates. *Interdiscip Toxicol.* 5(2):47–58.
- Gajalakshmi,S., Vijayalakshmi.S.,dan Devi Rajeswari V. 2012. Phytochemical and Pharmacological Properties of *Annonamuricata* : A Review, *International Journal of Pharmacy and Pharmaceutical Sciences*, 4 (2): 5
- Gavamukulya.Y.,Wamunyokoli.F.,and El-Shemy,H.,A., 2017. *Anona muricata*.Is the natural therapy to most disease conditions including cancer growing in our backyard ? A systematic review of its research history and future prospect : A Review, *Asian Pasific Journal of Tropical Medicine*,10 (9): 835-848
- Gracia R., Snodgrass W. 2007. Lead Toxicity And Chelation Therapy. *Am J Health Syst Pharm.* 64(1), pp.45–53.
- Halliwell B & Gutteridge JMC. 2007.Cellular response to oxidative stress : adaptation, damage repair, senescence and death. In *Free Radical in Biology and Medicine*. 4th ed. London, Oxford : University Press ; 187 – 267
- Hariyatmi. 2004. Kemampuan Vitamin E sebagai Antioksidan terhadap Radikal Bebas pada Usia Lanjut. *MIPA* 14(1): 52–60
- H.J. Heo, S. J. Choi, S.-G. Choi, D.-H. Shin, J. M. Lee, and C. Y. Lee, “*Effects of banana, orange, and apple on oxidative stress-induced neurotoxicity in PC12 cells*,” *Journal of Food Science*, vol. 73, no. 2, pp. H28–H32, 2008. [View at Publisher](#) · [View at Google Scholar](#) · [View at Scopus](#)
- Holscher, C., 2001. *Neuronal mechanisms of memory formation*. Long-term potentiation as a model for memory mechanisms.The story so far, Cambridge University Press,1-37
- HölscherC, O'Mara SM. 1997. *Model Learning And Memory Systems In Neurobiological Research: Conditioning And Associative Learning Procedures And Spatial Learning Paradigms*. Dalam: Lynch MA & O'Mara

- Jones, M., 2013. Neural Pathway. *University of Bristol*. Available at: <http://www.bristol.ac.uk> [Accessed January 1, 2016].
- Juananda D, Astari R.V, 2018. Stres Imobilisasi Mengganggu Memori Spasial Mencit Putih (*Mus musculus*) galur Swiss Webster Jantan, *Anatomica Medical Journal*, vol.1.no.1, Universitas Riau, Pekanbaru
- Juliandi A, Boediono B, Sasai A, and Noriaki, 2018. Uji Simplisia Daun Sirsak (*Annona muricata* L.) terhadap Kompleksitas Dendrit Hipokampus dan Kinerja Memori Spasial Mencit (*Mus musculus*) UT Biologi IPB Bogor
- Kristianingrum, YP, Widyarini S, Kurniasih, Sutrisno B, Tabba C R, Sugiyono. 2016. Gambaran hispatologi otak mencit akibat injeksi Trimetyltin sebagai model penyakit Alzheimer (abstrak). Di dalam: *Jurnal Sain Veteriner*, hal 34
- Konig D, Berg A. Exercise and Oxidative Stress : is there a need for additioal antioxidant. *Osterreichisches J Fur Sportmedizin* 2002;3:6-15.
- Kujoth GC, Bradshaw PC, Haroon S, Prolla TA. 2007. The role of mitochondrial DNA mutations in mammals aging. *Plos Genet* 23:e24.
- Kuntjojo. 2009. *Metode Penelitian*. Kediri: Universitas Nusantara PGRI.Lungan, R. 2006. *Aplikasi Statistika dan Hitung Peluang* . Yogyakarta: Penerbit Graha Ilmu
- Lieberman M & Marks A. 2009. Mark's basic medical biochemistry : a clinical approach. Philadelphia : Wolter Kluwer / Lippincot Williams & Wilkins
- Liu, J., A. Wang, L. Li, Y. Huang, P. Xue and A. Hao. 2010. Oxidative Stress Mediates Hippocampal Neuron Death in Rats After Lithium–Pilocarpine-Induced StatusEpilepticus. *Seizure* 19: 165-172.
- Liu JT, Chen BY, Zhang JQ, Kuang F, Chen LW. 2015. Lead Exposure Induced Microgliosis And Astrogliosis In Hipokampus of Young Mice Potentially By Triggering TLR4-MyD88-NFκB Signaling Cascades. *Elsevier*. 239(2):97–107.
- Lubis B, Rosdiana N, Nafianti S, Rasyianti O, Panjaitan FM. 2013. Hubungan Keracunan Timbal dengan Anemia Defisiensi Besi pada Anak. *CDK-200*. 40(1):17–21
- Luciana, A.R. 2010. Acetogenins from *Annonacornifolia* and their antioxidant capacity. Departamento de Química, Instituto de CiênciasExatas, Universidade Federal de Minas Gerais. MG, Brazil. Page 2
- Lynch,2004. Long-term potentiation and memory. *Physiol.Rev*,**84**,87-136

- Maalik A, Farhan A., Amara M, Adeem M, Saira A, Muhammad Atif, Sabiha K, Yasir A and Imran Tariq. Pharmacological Applications of Kuersetin and its Derivatives: A Short Review. *Tropical Journal of Pharmaceutical Research*. September 2014; 13 (9): 1561-1566.
- Mello CF, Kraemer CK, Filippin A, Morsch VM, Rodrigues ALS, Martins AF, et al. 1998. Effect of lead acetate on neurobehavioral development of rats. *Braz J Med Biol Res*. 31(7):943–50.
- Middleton, E. J. R., Kandaswami, C and Theoharides, T.C. 2000. The Effect of Plant Flavonoids on Mammalian Cells: Implications for Inflammation, Heart Diseases, and Cancer. *Pharmacol Rev*. 52 (4): 673-751.
- Moghadamtousi, S.Z. , Fadaeinasab, M., Nikzad, S., Mohan, G., Ali, H.M., Kadir, H.A. 2015. *Annona muricata* (Annonaceae): A Review of its traditional uses, isolated acetogenins and biological activities. *International Journal of Molecular Sciences*. 16:15625-58.
- Nasrun MWS. 2013. *Demensia*. Dalam: S. D. Elvira & G. Hadisukanto, penyunting. *Buku Ajar Psikiatri*. Jakarta: Badan Penerbit Fakultas Kedokteran Universitas Indonesia. hlm. 537–48.
- Ngabekti, S & Isnaeni, W. 2000. Pemanfaatan Kurkumin Untuk Mengeliminir Pengaruh Diazonin terhadap Kerusakan Hati Mencit (*Mus musculus* L). Semarang: Fakultas MIPA Universitas Negeri Semarang
- Ohkawa H, Ohishi N, Yagi K. 1979. Assay for lipid peroxide in animal tissue by thiobarbituric acid reaction. *Anal Biochem*. 95: 351-358.
- Pagnussat Ads, Faccioni MC, Netto CA, Achavall M. 2007. An ultrastructural study of cell death in the CA1 pyramidal field of the hippocampus in rats submitted to transient global ischemia followed by reperfusion, *J. Anat*, 211, 589-99
- Palak M, Vishu P, and Gayatri R. Kuersetin – A Flavonoid :A Systematic Review. *Journal of Pharmaceutical Sciences and Research*. Vol. 8(8), 2016, 878-880.
- Paulsen F, Waschke J. 2012. Sobotta : Atlas Anatomi Manusia : Kepala, Leher, dan Neuroanatomi. Jakarta: EGC
- Perez, A.L., Mendoza, M.M.L., Cortes, Z.R., 2006, *Cytotoxic effect of curcumin on Giardia lamblia trophozoites*, *Acta Trop.*, **98**, 152 -161.
- Piccini A., Russo C., Gliozzi A., Relini A., Borghi R., Giliberto L., Armirotti A., D'Arrigo C., Bachi A., Cattaneo A., Canale C., Torrassa S., Saido T.C.,

- Markesbery W., Gamberti P., Tabaton M., 2005, Beta amyloid is different in normal aging and in Alzheimer disease, *J Biol Chem* **280**: 34186 – 34192
- Puspitasari M L.Wulansari TV, WidyaningsihTD,Maligan JM, Nugrahini NIP, 2016.Antioksidan suplemen herbal daun sirsak (*Annona muricata* L) dan kulit manggis (*Garcinia mangostana* L.: Kajian Pustaka. Di dalam: Jurnal Pangan Agroindustri vol 4 No 1 p 293-290
- Prasetya, D.Y., 2014. Efek Ekstrak Etanol Temulawak pada Tikus Model Demensia :Kajian Memori Spasial, Status Antioksidan, dan Gambaran hispatologi hipokampus, UAD Yogyakarta
- Prihatiningsih,T.,Hasniastuti,T.,and Agustina, D., 2017. Efek antigenotoksik ekstrak etanolik daun sirsak (*Annona muricata* Linn) terhadap frekuensi mikronukleus mukosa bukal tikus Sprague Dawley.Majalah Kedokteran Gigi Indonesia Vol 1 (3)
- Ratu, A.P, Setyahadi S.,Simanjutak, 2017. Aktivitas dan Kinetika inhibisi asetilkolinesterase obat fraksi etanol daun sirsak.the 5th Urecol Proceeding, UAD Yogyakarta
- Ren, W., Qiao, Z., Wang, H., Zhu, L & Zhang, L. 2003. Flavonoids: promising anticancer agent. *Medicinal Research Review* 23(4): 519–534
- Sadek, K. M.2012. Antioxidant and immunostimulant effect of *Carica papaya* Linn. Aqueous extract in acrylamide intoxicated rats. *Acta Informatica Medica*, 20(3), 180–185. <https://doi.org/10.5455/aim.2012.20.180-185>
- Sampaio, C.G., Trevisan, M.T.S., Brito, E.S.,Santiago, G.M.P., Feitosa, C.M.,Carvalho, J.I.X., et.al. 2008. Screening for acetylcholinesterase inhibitor from seeds to treat Alzheimer's. *Proceeding 4th Brazilian Symposium on Medicinal Chemistry Disease, Division of Medicinal Chemistry, Brazilian Chemical Society (SBQ). BrazMedChem.*
- SantaCruz K., Lewis J., Spires T., Paulson J., Kollinek L., Ingelsson M., Guimaraes A., DeTure M., Ramsden M., McGowan E., Forster C., Yue M., Orne J., Janus C., Mariash A., Kuskowski M., Hyman B., Hutton M., Ashe K.H., 2005, *Tau Suppression in a Neurodegenerative Mouse Model Improves Memory Function*, *Science* 309: 476-481
- Sari, D.M,Wijaya S.,Setiawan H.K., 2015. Fraksinasi dan Identifikasi senyawa antioksidan pada Ekstrak Etanol daun sirsak secara Kromatografi Kolom, Fakultas Farmasi, Universitas Katolik Widya Mandala, Surabaya
- Seeley WW, Miller BL. *Alzheimer disease and other dementias*.2015. In: Kasper DL, Fauci SA, Hasuer SL, Longo DL, Jameson JL, Loscalzo J, et al.

- Harrison's principles of internal medicine. 19th New York: McGraw-Hill Education
- Septiana, S.I., and Puruhita, N., 2015. Pengaruh Pemberian Ikan Teri pada memori spasial tikus Sprague Dawley usia satu bulan, *Journal of Nutrition College*, vol 1, no 1, Universitas Diponegoro, Semarang
- Serra V, Zglinicki T, Lorenz M, Saretzki G. 2003. Extracellular superoxide dismutase is a major antioxidant in human fibroblasts and slows telomere shortening. *J Biol Chem* 278:6824-6830.
- Sherwood L. 2014. *Fisiologi Manusia dari Sel ke Sistem*. Jakarta: Buku Kedokteran EGC.
- Shuto, M., Higuchi, K., Sugiyama, C., Yoneyama, M., Kuramoto, N., Nagashima, R., Kawada, K., Ogita, K., 2009, Endogenous and exogenous glucocorticoids prevent trimethyltin from causing neuronal degeneration of the mouse brain in vivo: involvement of oxidative stress pathways, *J. Pharmacol. Sci.*, 110, 424-36
- Speakman JR, Talbot DA, Selman C, Snart S, McLaren JS, Redman P, Krol E, Jackson DM, Johnson MS, Brand MD. 2004. Uncoupled and surviving: individual mice with high metabolism have greater mitochondrial uncoupling and live longer. *Aging Cell* 3:87-94
- Siswarni, M., Z., Nuryani, Sinaga, S., D., 2016. *Ekstraksi Acetogenin dari daun dan biji sirsak (Annona muricata L) dengan pelarut aceton*. *Jurnal Teknik Kimia USU*, vol 5, no. 2 SM, penyunting. Neuroscience Labfax. London: Academic Press. hlm. 10-24
- Siswonoto, S., 2008. Hubungan kadar malondialdehid plasma dengan keluaran klinis stroke iskemik akut, Program Pasca Sarjana, Universitas Diponegoro
- Stenberg RJ, Sternberg K. 2009. *Cognitive psychology*. Belmont: Wadsworth.
- Sreelatha, S & Padma, P.R. 2009. Antioxidant activity and total phenolic content of *Moringa oleifera* leaves in two stages of maturity. *Plant foods hum nutr* 64(4):302–311
- Sunarjono, H. 2005. *Sirsak dan Srikaya*. Cetakan pertama. Jakarta: Penebar Swadaya. Hal. 14-15, 22-25
- Sunarno, 2009, Profil Antioksidan Copper Zink Superoxide Dismutase (Cu, Zn, SOD) pada Sel-sel Ginjal Mencit Sprague dawley melalui Pewarnaan Imunohistokimia Polimer Peroksidase. *BIOMA*, Juni Vol 11, No. 1, hal 33-39

- Taupin, P., 2008. *The Hipokampus – Neurotransmission and Plasticity in TheNervous System*, New York: Nova Biomedical Books.
- TonniesE,Trushina E.2017.Journal Alzheimer’s Disease : Oxidative Stress ,Synaptic Dysfunction, and Alzheimer’s Disease..*Journal of Alzheimer Disease* 57 : 1105-1121
- Vianandra R. 2011. Pengaruh suplementasi ekstrak daun *Annona muricata* terhadap kejadian displasia epitel kelenjar payudara tikus Sprague dawley yang diinduksi 7, 12 Dimethylbenz[α]anthracene. [Skripsi]. Semarang: Universitas Diponegoro.
- Vorhees,Charles V. dan Williams, Michael T., 2006, Morris water maze: procedures for assessing spatial and related forms of learning and memory, *Nat. Protoc.*,2, 848-858
- Vorhees, Charles V. dan Williams, Michael T.,2010, Morris water maze : procedures for assessing spatial and related form of learning and memory, US National Library of medicine National Institute of Health
- Wagner, H., 1984. *Plant Drug analysis a Thin Layer Chromatography Atlas*. Springer-Verlag. Berlin
- Waji, R,A & Sugrami, A. 2009. Makalah kimia organic bahan alam flavonoid (quercetin). Makassar:Universitas Hasanuddin 8–9.
- Waxman, S.G., 2009. *Clinical Neuroanatomy* 26th ed., New York: McGraw-Hill.
- Wijaya. A, 1996, *Radikal Bebas dan Parameter status antioksidan*, Forum diagnosyicum no.1.
- Yustika, A.R.,Aulanni, Prasetyawan.,B.,2013, Kadar Malondialdehid (MDA) & Gambaran hispatologi pada ginjal tikus putih pasca induksi Cylosperine, *Kimia Student Journal*, vol.1(1), pp 222-228
- Zakiah, N.,Yanuarman Frengky,& Munazar, 2017. Aktivitas hepatoprotektif ekstrak etanol daun sirsak terhadap kerusakan hati tikus yang diinduksi dengan paracetamol, *Action Journal*, vol 2 (1)

LAMPIRAN

Lampiran 1. Surat Keterangan Ekstrak Daun Sirsak

 BOROBUDUR NATURAL HERBAL INDUSTRY		CPOTB/GMP Certified ISO 9001:2015	ISO 14001:2015	ISO 22000:2018
BOROBUDUR INDUSTRI JAMU, PT. Head Office : Jl. Madukoro Blok A/26 Semarang 50141 T. +62-24-7606888 F. +62-24-7605553 E-mail : office@borobudurherbal.com		www.borobudurherbal.com		
Safety Data Sheet				
1. Identification				
<i>Identification of Product</i>				
Product Name	: Soursof P.E			
Local Name	: Sirsak			
Plant species	: <i>Annona muricata</i>			
Indication	: Cancer			
<i>Identification of Company</i>				
Company Name	: Borobudur Natural Herbal Industry Semarang – Indonesia			
Emergency Telephone	: +62 24 3510 785			
2. Hazards Identification				
This Product is not considered to be hazardous				
3. Information of Ingredients				
Annonae Folium Extract				
Maltodextrine				
4. First Aid Measures				
Inhalation	: Remove individual to fresh air			
Skin Contact	: Flush skin with water. Call a physician if irritation develops			
Eye Contact	: Flush eye with water. Call a physician if irritation develops			
5. Fire Fighting Measures				
Extinguishing media	: Carbon dioxide, dry chemicals, foam, and water spray			
6. Accidental Release Measures				
Personal precautions	: Use personal protective equipment			
Environmental precautions	: No special environmental precautions required			
Methods for cleaning up	: Sweep up and flush with water			



BOROBUDUR
NATURAL HERBAL INDUSTRY

CPOTB/GMP Certified
Good Manufacturing Practice

ISO 9001:2015

SAP
Integrated System

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9. Physical and Chemical Properties

Physical Properties

Appearance	Granule
Color	Dark Brown
Odor	Aromatic
Taste	Bitter
Mesh Size	70 % pass mesh 12
Loss on Drying	5.0 % max.

Chemical Properties

Heavy Metal:	
Arsenic (As)	5 ppm max
Lead (Pb)	10 ppm max

10. Stability and Reactivity

Stability and Reactivity Summary : Stable under normal conditions

11. Toxicological Data

Human experience : no toxic effect in normal use

12. Ecological Information

No significant environmental hazard or adverse effect from human exposure resulting from the accidental release of this material is anticipated

13. Disposal Advice

No special disposal method required, except that it be in accordance with current local authority regulations

14. Transport Information

Not applicable

15. Regulation Information

Labelling according to NADFC guidelines.

Observe the general safety regulations when handling chemicals

16. Further Information

This statements are based on our present knowledge and are meant to describe our product in regards to our safety regulations.

Therefore they are not meant to assure special characteristics. Existing laws and regulations have to be noticed by consignee himself. Since the conditions of use are beyond the control of our company, it is



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Certificate of Analysis
Dry Extract

MANUFACTURING DATA		GENERAL DATA	
Product Name	Soursop P.E	Plant Species	Annona muricata
Local Name	Sirsak	Botanical part used	Leaves
Batch Number	030PQ01.5	Ratio Botanical Extract	10 : 1
Manufacture Date	June 12, 2017	Excipients	Maltodextrin
Testing Date	June 13, 2017	Preservatives	N/A
Expire Date	June 12, 2021	Extraction Solvent	Ethanol 70%
Shelf Life	4 years	Storage	store in cool and dry place, keep away from strong light and heat
ITEM	SPECIFICATION	TEST RESULT	TEST METHOD
IDENTIFICATION TEST			
Appearance	Granule	Complies	Visual
Color	Dark Brown	Complies	Visual
Odor	Aromatic	Complies	Organoleptic
Taste	Bitter	Complies	Organoleptic
Mesh Size	70 % pass mesh 12	Complies	12 mesh screen
Loss On Drying	5.0 % max	3.92 %	2g/105°C/15 minutes
HEAVY METALS			
Arsenic (As)	5 ppm max	Complies	AAS
Lead (Pb)	10 ppm max	Complies	AAS
MICROBIOLOGICAL TEST			
Total Plate Count	Not more than 1000 cfu/gram	< 100 cfu/gram	Dilution Plating
Fungi/Yeast and molds	Not more than 100 cfu/gram	< 100 cfu/gram	Dilution Plating
E. coli	Should be absent	Complies	MPN Method
Salmonella	Should be absent	Complies	Dilution Plating
S. aureus	Should be absent	Complies	Dilution Plating
P. aeruginosa	Should be absent	Complies	Dilution Plating

Lampiran 2. Surat Keterangan *Ethical Clearance*

**KOMISI ETIK PENELITIAN KESEHATAN
HEALTH RESEARCH ETHICS COMMITTEE
SEKOLAH TINGGI ILMU KESEHATAN SURYA MITRA HUSADA
INSTITUTE OF HEALTH SCIENCE SURYA MITRA HUSADA**

**KETERANGAN LOLOS UJI ETIK
DESCRIPTION OF ETHICAL APPROVAL
"ETHICAL APPROVAL"**

NOMOR : 930/KEPK/XI/2018

Komite Etik Penelitian Kesehatan STIKes Surya Mitra Husada dalam upaya melindungi hak asasi dan kesejahteraan subyek penelitian kesehatan, telah mengkaji dengan teliti protokol berjudul :

Health Research Ethics Committee STIKes Surya Mitra Husada in the effort to protect the rights and welfare of research subjects of health, has reviewed carefully the protocol entitled:

**EFEK EKSTRAK ETANOL DAUN SIRSAK (*Annona muricata* L.) TERHADAP MEMORI SPASIAL,
KADAR MALONDIALDEHID, DAN JUMLAH SEL PYRAMIDAL HIPOKAMPUS
AREA CA1 DAN CA2-CA3 PADA MENCIT MODEL DEMENSIA**

Peneliti Utama : Hengky Dwi Cahyono / SBF121710147
Principal Investigator

Lampiran 3. Surat Keterangan Uji MDA

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LAPORAN HASIL UJI <i>(Analysys Certificate)</i> No.PSPG/009/I/2019		
Nomor Pengujian <i>(Analysis Report Number)</i> Nama Pelanggan <i>(Name of client)</i> Alamat Pelanggan <i>(Address of client)</i> No. Telepon Pelanggan <i>(Phone No. of client)</i> Contoh Uji <i>(Type of sample)</i> Tanggal Penerimaan Contoh Uji <i>(Date of analysis)</i> Tanggal Pengujian <i>(Date of analysis)</i> Metode Uji <i>(Analysis Method)</i> Hasil Uji <i>(Analysis Result)</i>	: PS/456/XII/2018 : Hengky Dwi Cahyono : : : Jaringan : 26 Desember 2018 : 26 Desember 2018 : : MDA	

Lampiran 4. Gambar Ekstrak Daun Sirsak



Lampiran 5. Gambar Alat Uji Memori *Morris Water Maze*



Lampiran 6. Penentuan Dosis Ekstrak Etanol Daun Sirsak

Ekstrak daun sirsak kandungan flavonoid, terpenoid, alkaloid, minyak atsiri, polifenol, saponin, dan tanin yang berperan sebagai antitumor, antimikroba, antiparasit, dan antivirus (Amelia *et al.* 2012).

a.EEDS 100 mg/KgBB

Dikonversikan kemencit

Dosis untuk mencit 20 g = $0,14 \times 100 \text{ mg} / \text{KgBB}$

$$= 14 \text{ mg} / \text{KgBBmencit}$$

$$= 14 \text{ mg} / 1000 \text{ gBBmencit}$$

$$= 0,28 \text{ mg} / 20 \text{ gBBmencit}$$

volume pemberian untuk masing-masing mencit 20 g adalah 0,5 ml

$$\text{Larutan stok} = \frac{\text{volumelaru tan stok}}{\text{volumepemberian}} \times \text{dosis}$$

$$= \frac{100ml}{0,5ml} \times 0,28mg$$

$$= 56mg$$

Jadi sebanyak 56 mg EEDS dilarutkan dalam 100 ml CMC-Na 0,5%

Berat badan mencit (g)	Volume penyuntikkan (ml)
28	$\frac{28mg}{20mg} \times 0,5ml = 0,7$
30	$\frac{30mg}{20mg} \times 0,5ml = 0,75$
26	$\frac{26mg}{20mg} \times 0,5ml = 0,65$
32	$\frac{32mg}{20mg} \times 0,5ml = 0,8$
25	$\frac{25mg}{20mg} \times 0,5ml = 0,625$

b. EEDS 200 mg/KgBB

Dikonversikan ke mencit

$$\text{Dosis untuk mencit } 20 \text{ g} = 0,14 \times 200mg / KgBB$$

$$= 28mg / KgBB_{mencit}$$

$$= 28mg / 1000gBB_{mencit}$$

$$= 0,56mg / 20gBB_{mencit}$$

volume pemberian untuk masing-masing mencit 20 g adalah 0,5 ml

$$\text{Larutan stok} = \frac{\text{volumelarutan stok}}{\text{volume pemberian}} \times \text{dosis}$$

$$= \frac{100ml}{0,5ml} \times 0,56mg$$

$$= 112mg$$

Jadi sebanyak 112 mg EEDS dilarutkan dalam 100 ml CMC-Na 0,5%

Berat badan mencit (g)	Volume penyuntikkan (ml)
------------------------	--------------------------

28	$\frac{28mg}{20mg} \times 0,5ml = 0,7$
30	$\frac{30mg}{20mg} \times 0,5ml = 0,75$
26	$\frac{26mg}{20mg} \times 0,5ml = 0,65$
32	$\frac{32mg}{20mg} \times 0,5ml = 0,8$
25	$\frac{25mg}{20mg} \times 0,5ml = 0,625$

c. EEDS 400 mg/KgBB

Dikonversikan ke mencit

Dosis untuk mencit 20 g = $0,14 \times 400mg / KgBB$

$$= 56mg / KgBBmencit$$

$$= 56mg / 1000gBBmencit$$

$$= 1,12mg / 20gBBmencit$$

volume pemberian untuk masing-masing mencit 20 g adalah 0,5 ml

$$\text{Larutan stok} = \frac{\text{volumelarutan stok}}{\text{volume pemberian}} \times \text{dosis}$$

$$= \frac{100ml}{0,5ml} \times 1,12mg$$

$$= 224mg$$

Jadi sebanyak 224 mg EEDS dilarutkan dalam 100 ml CMC-Na 0,5%

Berat badan mencit (g)	Volume penyuntikkan (ml)
28	$\frac{28mg}{20mg} \times 0,5ml = 0,7$
30	$\frac{30mg}{20mg} \times 0,5ml = 0,75$
26	$\frac{26mg}{20mg} \times 0,5ml = 0,65$
32	$\frac{32mg}{20mg} \times 0,5ml = 0,8$

$$\frac{25mg}{20mg} \times 0,5ml = 0,625$$

Lampiran 7. Pemberian Sediaan Uji Ekstrak Etanol Daun Sirsak



Lampiran 8. Pembuatan Larutan Kuersetin dan Plumbum Acetat

1. Larutan CMC-Na 0,5%

Larutan CMC Na 0,5% = $0,5g/100ml$

$$= 500mg/100ml$$

Jadi sebanyak 500 mg CMC-Na dilarutkan dalam 100 ml air panas sedikit demi sedikit lalu aduk sampai larut dan homogen

Berat badan mencit (g)	Volume penyuntikkan (ml)
28	$\frac{28mg}{20mg} \times 0,5ml = 0,70$
30	$\frac{30mg}{20mg} \times 0,5ml = 0,75$
26	$\frac{26mg}{20mg} \times 0,5ml = 0,65$

$$\begin{array}{rcl} 32 & \frac{32mg}{20mg} \times 0,5ml & = 0,80 \\ 25 & \frac{25mg}{20mg} \times 0,5ml & = 0,625 \end{array}$$

2. Dosis kuersetin

Menurut penelitian (Kale et al, 2018) dosis kuersetin 50 mg/KgBB dapat melindungi saraf terhadap cedera otak setelah pengion radiasi

$$\begin{aligned} \text{Dosis untuk mencit } 20 \text{ g} &= 50mg / KgBB \\ &= 50mg / 1000KgBBmencit \\ &= 42mg / 1000gBBmencit \\ &= 1mg / 20gBBmencit \end{aligned}$$

volume pemberian untuk masing-masing mencit 20 g adalah 0,5 ml

$$\begin{aligned} \text{Larutan stok} &= \frac{\text{volumelarutan stok}}{\text{volume pemberian}} \times \text{dosis} \\ &= \frac{100ml}{0,5ml} \times 0,84mg \\ &= 168mg \end{aligned}$$

Jadi sebanyak 168 mg kuersetin dilarutkan dalam 100 ml CMC-Na 0,5%

Berat badan mencit (g)	Volume penyuntikkan (ml)
28	$\frac{28mg}{20mg} \times 0,5ml = 0,7$
30	$\frac{30mg}{20mg} \times 0,5ml = 0,75$
26	$\frac{26mg}{20mg} \times 0,5ml = 0,65$
32	$\frac{32mg}{20mg} \times 0,5ml = 0,8$
25	$\frac{25mg}{20mg} \times 0,5ml = 0,625$

3. Larutan Pb asetat

Larutan timbal asetat diberikan secara intra peritoneal dengan dosis 50 mg/kgBB kepad atikus (Owolabiet al, 2014).

Dikonversikan ke mencit

Dosis untuk mencit 20 g = $0,14 \times 50 \text{ mg} / \text{KgBB}$

$$= 7 \text{ mg} / \text{KgBBmencit}$$

$$= 7 \text{ mg} / 1000 \text{ gBBmencit}$$

$$= 0,14 \text{ mg} / 20 \text{ gBBmencit}$$

volume pemberian untuk masing-masing mencit 20 g adalah 0,2 ml

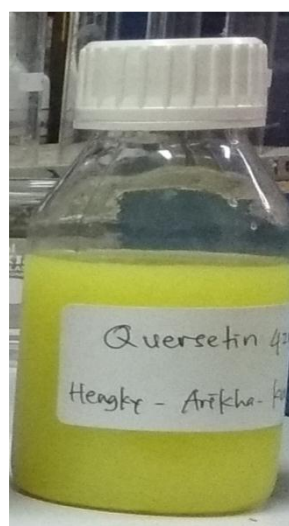
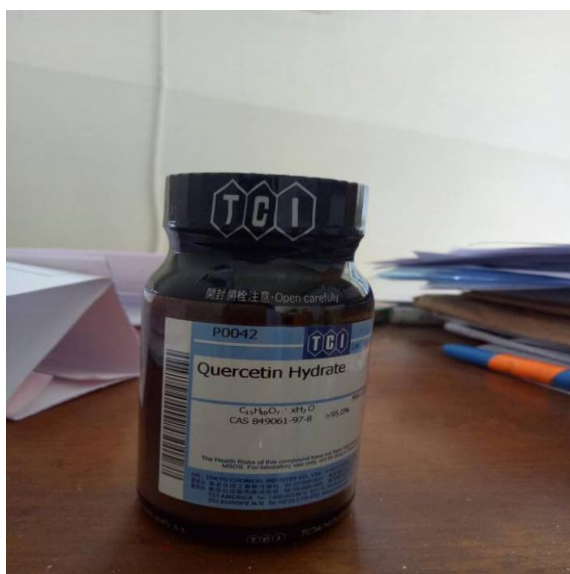
$$\text{Larutan stok} = \frac{\text{volumelarutan stok}}{\text{volume pemberian}} \times \text{dosis}$$

$$= \frac{10 \text{ ml}}{0,2 \text{ ml}} \times 0,14 \text{ mg}$$

$$= 7 \text{ mg}$$

Jadi sebanyak 7 mg Pb asetat dilarutkan dalam aquabidestilata 10 ml

Berat badan mencit (g)	Volume penyuntikkan (ml)
28	$\frac{28 \text{ mg}}{20 \text{ mg}} \times 0,2 \text{ ml} = 0,28$
30	$\frac{30 \text{ mg}}{20 \text{ mg}} \times 0,2 \text{ ml} = 0,30$
26	$\frac{26 \text{ mg}}{20 \text{ mg}} \times 0,2 \text{ ml} = 0,26$
32	$\frac{32 \text{ mg}}{20 \text{ mg}} \times 0,2 \text{ ml} = 0,32$
25	$\frac{25 \text{ mg}}{20 \text{ mg}} \times 0,2 \text{ ml} = 0,25$



Lampiran 9. Gambar Pengambilan Otak Pada Mencit



Lampiran 10. Data Hasil Pengujian

1. DATA AQCUISITION TRIAL

KELOMPOK	MENCIT	LATIHAN		T0	T1	T2	T1-T2	%
NORMAL	1	26.79	27.83	27.31	26.88	25.59	1.29	4.80
	2	27.35	27.71	27.53	27.07	25.33	1.74	4.38
	3	26.81	26.89	26.85	26.22	25.33	0.89	4.27
	4	27.84	26.39	27.12	26.09	25.08	1.01	4.21
	5	28.04	26.03	27.04	26.08	25.13	0.95	4.53
	RERATA	27.37	26.97	27.17	26.47	25.29	1.18	4.44
	SD						0.14	0.17
KONTROL POSITIF	1	26.94	24.32	25.63	23.18	22.62	0.56	2.42
	2	26.17	24.94	25.56	22.86	23.47	-0.61	-2.67
	3	25.63	24.25	24.94	22.93	22.62	0.31	1.35

	4	26.76	24.92	25.84	23.17	22.64	0.53	2.29
	5	26.65	24.87	25.74	22.79	22.17	0.21	1.04
	RERATA	26.43	24.66	25.54	22.90	22.70	0.20	0.89
	SD						0.07	0.42
KONTROL NEGATIF	1	26.78	24.71	25.75	25.38	29.14	-3.76	37.92
	2	27.18	26.07	26.63	25.13	28.75	-3.62	-34.66
	3	27.25	26.43	26.84	24.74	28.83	-4.09	27.83
	4	26.61	25.63	26.12	24.27	28.73	-4.46	27.40
	5	27.12	26.11	26.62	23.93	29.16	-5.23	28.08
	RERATA	26.99	25.79	26.39	24.69	28.92	-4.23	-17.31
	SD						0.31	2.30
EEDS 100	1	30.02	27.64	29.21	26.14	25.63	0.51	1.95
	2	29.18	28.06	29.25	25.95	24.95	1.00	3.85
	3	28.94	28.04	28.57	25.18	25.52	-0.34	-1.35
	4	29.19	27.93	28.05	25.09	24.98	0.11	2.94
	5	29.07	27.93	27.91	25.82	24.43	1.42	1.52
	RERATA	29.28	27.92	28.60	25.64	25.10	0.54	2.10
	SD						0.39	1.52
EEDS 200	1	28.52	27.96	28.34	25.67	25.21	0.29	1.45
	2	28.54	27.61	27.61	25.74	25.06	0.63	2.43
	3	28.31	26.37	27.98	25.85	25.41	0.39	1.51
	4	28.38	26.72	27.46	25.72	25.16	0.56	2.14
	5	28.32	26.75	27.35	25.86	25.17	0.51	1.83
	RERATA	28.41	27.08	27.75	25.77	25.29	0.48	1.87
	SD						0.43	1.69
EEDS 400	1	27.67	25.16	27.05	25.65	25.83	-0.18	-0.68
	2	28.14	25.32	26.55	24.39	25.96	-1.57	-6.44
	3	28.04	25.61	26.32	24.85	24.13	0.72	2.90
	4	27.12	25.17	26.48	25.13	24.27	0.84	3.08
	5	27.07	25.08	25.78	25.04	24.39	0.65	3.03
	RERATA	27.61	25.27	26.44	25.01	24.92	0.09	0.38
	SD						0.07	0.30

2. DATA PROBE TRIAL

KELOMPOK	MENCIT	LATIHAN		T0	T1	T2	T2-T1	%
NORMAL	1	6.46	6.37	6.42	6.18	7.17	0.99	13.25
	2	6.12	5.82	5.97	7.01	7.12	0.11	1.54
	3	5.3	5.47	5.39	6.46	7.74	1.28	15.84
	4	6.19	5.43	5.81	6.18	6.85	0.71	10.63
	5	5.72	5.74	5.73	6.14	6.95	0.81	11.65
	RERATA	5.96	5.77	5.86	6.39	7.17	0.78	10.58
	SD						0.77	9.29
KONTROL POSITIF	1	2.82	2.61	2.72	4.61	6.64	2.03	30.57
	2	2.43	3.45	2.94	4.91	5.68	0.77	13.56
	3	3.45	2.71	3.08	3.28	5.52	2.24	40.58
	4	2.86	2.95	2.91	3.96	5.17	1.21	23.40
	5	2.94	2.56	2.75	4.12	5.14	1.02	21.72
	RERATA	2.90	2.86	2.88	4.18	5.63	1.45	25.97
	SD						0.64	10.87
KONTROL NEGATIF	1	5.63	4.31	4.97	6.71	3.87	-2.84	-73.39
	2	5.65	4.29	4.97	6.72	4.12	-2.60	-63.11
	3	6.45	4.36	5.41	7.45	4.15	-3.30	-79.52
	4	5.73	4.56	5.18	6.83	4.16	-2.67	-64.18
	5	5.29	4.07	4.68	7.09	4.03	-3.06	-75.58
	RERATA	5.75	4.32	5.04	6.96	4.07	-2.89	-71.15
	SD						0.32	7.19
EEDS 100	1	4.12	4.29	4.21	3.52	5.12	1.60	31.25
	2	4.27	4.17	4.22	3.71	5.82	2.11	36.25
	3	4.12	3.28	3.70	5.51	7.58	2.07	27.31
	4	4.28	4.14	4.21	4.31	6.29	1.98	31.48
	5	4.06	3.69	3.88	4.21	6.02	1.81	31.72
	RERATA	4.17	3.91	4.04	4.25	6.17	1.93	31.60
	SD						0.28	4.48
EEDS 200	1	4.28	4.23	4.26	5.77	7.83	2.06	26.31

	2	4.44	3.12	3.78	4.84	6.28	1.44	22.93
	3	4.68	3.14	3.91	5.48	7.72	2.24	29.02
	4	4.32	3.52	3.92	5.62	7.42	1.80	24.26
	5	4.62	3.49	4.06	5.11	7.15	2.04	27.88
	RERATA	4.47	3.50	3.98	5.36	7.28	1.91	26.08
	SD						0.42	3.05
EEDS 400	1	4.18	4.23	4.21	5.82	6.76	0.94	13.91
	2	4.67	4.13	4.40	4.49	6.83	2.34	34.26
	3	4.72	3.09	3.91	4.46	6.97	2.51	36.01
	4	4.82	4.12	4.47	5.12	6.81	1.69	24.82
	5	4.19	3.52	3.86	4.78	6.87	2.09	31.32
	RERATA	4.52	3.82	4.17	4.92	6.85	1.93	28.06
	SD						0.86	12.29

3. DATA UJI SENSORIS

KELOMPOK	MENCIT	LATIHAN		T0	T1	T2	T1-T2	%
NORMAL	1	18.03	17.92	17.98	17.62	17.21	0.41	2.33
	2	17.3	16.04	16.67	16.33	15.89	0.44	2.69
	3	17.78	16.99	17.39	17.09	16.64	0.45	2.63
	4	17.86	17.14	17.50	16.83	16.73	0.10	0.59
	5	17.54	16.82	17.18	17.17	16.42	0.75	4.48
	RERATA	17.70	16.98	17.34	17.01	16.58	0.43	2.55
	SD						0.02	0.20
KONTROL POSITIF	1	17.83	17.18	17.51	17.10	16.92	0.18	1.05
	2	17.24	16.78	17.01	16.65	16.13	0.52	3.12
	3	18.56	18.74	18.65	18.32	18.11	0.21	1.15
	4	17.78	17.63	17.71	17.62	16.98	0.64	3.63
	5	17.98	17.52	17.75	17.12	17.09	0.03	0.18

	RERATA	17.88	17.57	17.72	17.36	17.05	0.30	1.77
	SD						0.19	1.17
KONTROL NEGATIF	1	18.43	18.95	18.69	17.75	21.37	-3.62	-20.39
	2	19.62	18.87	19.25	16.97	21.67	-4.70	-27.70
	3	19.92	19.62	19.77	17.89	21.69	-3.80	-21.24
	4	19.54	19.25	19.40	17.73	21.78	-4.05	-22.84
	5	19.08	19.06	19.08	17.36	21.39	-4.03	-23.21
	RERATA	19.32	19.15	19.24	17.54	21.58	-4.04	-23.11
	SD						0.58	3.99
EEDS 100	1	18.26	18.38	18.32	17.78	16.52	1.26	7.09
	2	19.11	18.95	19.03	18.01	16.33	1.68	9.33
	3	17.98	17.36	17.7	17.29	16.35	0.94	5.44
	4	18.53	18.32	18.4	17.82	16.73	1.09	6.12
	5	18.35	18.14	18.2	17.56	16.07	1.49	8.49
	RERATA	18.45	18.23	18.34	17.69	16.40	1.29	7.28
	SD						0.37	1.95
EEDS 200	1	18.86	17.96	18.41	18.09	17.74	0.35	1.93
	2	19.63	18.64	19.135	17.82	16.64	1.18	6.62
	3	18.74	17.59	18.165	17.09	16.24	0.85	4.97
	4	19.32	18.21	18.765	17.92	16.72	1.2	6.70
	5	18.84	17.91	18.375	17.43	17.02	0.41	2.35
	RERATA	19.08	18.06	18.57	17.67	16.87	0.79	4.51
	SD						0.42	2.38
EEDS 400	1	16.72	17.58	17.15	17.54	17.38	0.16	0.91
	2	19.82	17.88	18.85	17.91	17.25	0.66	3.69
	3	19.17	18.6	18.885	16.95	16.52	0.43	2.54
	4	18.83	17.98	18.405	17.62	16.89	0.73	3.89
	5	18.29	18.05	18.17	17.35	17.19	0.16	0.92
	RERATA	18.57	18.02	18.30	17.47	17.05	0.42	2.38
	SD						0.25	1.39

4. DATA CA

KELOMPOK	CA1	CA2	CA3
NORMAL	2	2	2
	2	1	1
	1	4	4
	2	2	2
	4	3	3
KONTROL POSITIF	2	3	3
	3	3	3
	2	2	2
	1	2	2
	2	1	1
KONTROL NEGATIF	23	17	17
	19	16	16
	16	14	14
	20	15	15
	22	15	15
EEDS_100	14	12	14
	14	12	12

	18	17	18
	11	10	15
	12	15	10
EEDS_200	7	5	35
	9	10	11
	9	8	8
	7	7	6
	8	9	7
EEDS_400	3	5	1
	4	1	3
	2	4	6
	1	2	5
	3	3	1

5. DATA MDA

	KELOMPOK	MDA (mmol/ml)	RATA-RATA
1	KONTROL SEHAT	0.76	
2		0.68	
3		0.52	
4		0.66	
5		0.79	0.682
1	KONTROL POSITIF (QUERSETIN)	0.34	
2		0.28	
3		0.37	
4		0.24	
5		0.36	0.318
1	KONTROL SAKIT/NEGATIF	1.92	
2		1.74	
3		1.70	
4		1.88	
5		1.85	1.818
1	EEDS 100MG	0.97	
2		0.82	

3		0.83	
4		1.12	
5		0.74	0.896
1	EEDS 200MG	0.73	
2		0.98	
3		0.65	
4		0.87	
5		0.56	0.758
1	EEDS 400MG	0.49	
2		0.24	
3		0.20	
4		0.31	
5		0.42	0.332

Lampiran 11. Hasil Signifikasi Data Uji Memori Spasial dan Gambaran Hispatologik Hipokampus

Uji	Normalitas	Homogenitas	ANOVA
<i>Aqcuisation trial</i>	0,125	0,064	0,000
<i>Probe trial</i>	0,175	0,325	0,000
Sensoris memoris	0,162	0,086	0,000
MDA	0,282	0,203	0,000
CA1	0,114	0,138	0,000
CA2	0,207	0,064	0,000

Lampiran 12. Hasil Uji LSD – TUKEY

1. ACQUISITION TRIAL

ANOVA

SELISIH_AQ

	Sum of Squares	df	Mean Square	F	Sig.
Between Groups	37.058	5	7.412	97.250	.000
Within Groups	.915	24	.076		
Total	37.972	29			

Post Hoc Tests

Multiple Comparisons

Dependent Variable: SELISIH_AQ

	(I) PERLAKUA N	(J) PERLAKUAN	Mean Difference (I- J)	Std. Error	Sig.	95% Confidence Interval	
						Lower Bound	Upper Bound
Tukey HSD	KONTROL NORMAL	KONTROL POSITIF	.98000 [*]	.22540	.009	.2229	1.7371
		KONTROL NEGATIF	-3.05667 [*]	.22540	.000	-3.8138	-2.2995
		EEDS_100	.64000	.22540	.117	-.1171	1.3971
		EEDS_200	.69667	.22540	.078	-.0605	1.4538
		EEDS_400	1.08333 [*]	.22540	.004	.3262	1.8405
	KONTROL POSITIF	KONTROL NORMAL	-.98000 [*]	.22540	.009	-1.7371	-.2229

	KONTROL NEGATIF	-4.03667 [*]	.22540	.000	-4.7938	-3.2795
	EEDS_100	-.34000	.22540	.666	-1.0971	.4171
	EEDS_200	-.28333	.22540	.802	-1.0405	.4738
	EEDS_400	.10333	.22540	.997	-.6538	.8605
KONTROL NEGATIF	KONTROL NORMAL	3.05667 [*]	.22540	.000	2.2995	3.8138
	KONTROL POSITIF	4.03667 [*]	.22540	.000	3.2795	4.7938
	EEDS_100	3.69667 [*]	.22540	.000	2.9395	4.4538
	EEDS_200	3.75333 [*]	.22540	.000	2.9962	4.5105
	EEDS_400	4.14000 [*]	.22540	.000	3.3829	4.8971
EEDS_100	KONTROL NORMAL	-.64000	.22540	.117	-1.3971	.1171
	KONTROL POSITIF	.34000	.22540	.666	-.4171	1.0971
	KONTROL NEGATIF	-3.69667 [*]	.22540	.000	-4.4538	-2.9395
	EEDS_200	.05667	.22540	1.000	-.7005	.8138
	EEDS_400	.44333	.22540	.412	-.3138	1.2005
EEDS_200	KONTROL NORMAL	-.69667	.22540	.078	-1.4538	.0605
	KONTROL POSITIF	.28333	.22540	.802	-.4738	1.0405
	KONTROL NEGATIF	-3.75333 [*]	.22540	.000	-4.5105	-2.9962
	EEDS_100	-.05667	.22540	1.000	-.8138	.7005
	EEDS_400	.38667	.22540	.547	-.3705	1.1438
EEDS_400	KONTROL NORMAL	-1.08333 [*]	.22540	.004	-1.8405	-.3262
	KONTROL POSITIF	-.10333	.22540	.997	-.8605	.6538
	KONTROL NEGATIF	-4.14000 [*]	.22540	.000	-4.8971	-3.3829
	EEDS_100	-.44333	.22540	.412	-1.2005	.3138
	EEDS_200	-.38667	.22540	.547	-1.1438	.3705

Homogeneous Subsets

SELISIH_AQ

PERLAKUAN		N	Subset for alpha = 0.05		
			1	2	3
Tukey HSD ^a	EEDS_400	5	.0933		
	KONTROL POSITIF	5	.1967		
	EEDS_200	5	.4800	.4800	
	EEDS_100	5	.5367	.5367	

KONTROL NORMAL	5	1.1767	
KONTROL NEGATIF	5		4.2333
Sig.		.412	.078 1.000

Means for groups in homogeneous subsets are displayed.

a. Uses Harmonic Mean Sample Size = 3.000.

2. PROBE TES

ANOVA

PROBE

	Sum of Squares	df	Mean Square	F	Sig.
Between Groups	53.488	5	10.698	30.666	.000
Within Groups	4.186	24	.349		
Total	57.674	29			

Post Hoc Tests

Multiple Comparisons

Dependent Variable: PROBE

	(I) PERLAKUAN	(J) PERLAKUAN	Mean Difference (I-J)	Std. Error	Sig.	95% Confidence Interval	
						Lower Bound	Upper Bound
Tukey HSD	KONTROL NORMAL	KONTROL POSITIF	-.66333	.48225	.740	-2.2832	.9565
		KONTROL NEGATIF	3.67667*	.48225	.000	2.0568	5.2965
		EEDS_100	-1.14333	.48225	.240	-2.7632	.4765
		EEDS_200	-1.13000	.48225	.250	-2.7498	.4898
		EEDS_400	-1.14667	.48225	.238	-2.7665	.4732
	KONTROL POSITIF	KONTROL NORMAL	.66333	.48225	.740	-.9565	2.2832
		KONTROL NEGATIF	4.34000*	.48225	.000	2.7202	5.9598
		EEDS_100	-.48000	.48225	.911	-2.0998	1.1398
		EEDS_200	-.46667	.48225	.920	-2.0865	1.1532
		EEDS_400	-.48333	.48225	.909	-2.1032	1.1365
	KONTROL NEGATIF	KONTROL NORMAL	-3.67667*	.48225	.000	-5.2965	-2.0568

	KONTROL POSITIF	-4.34000*	.48225	.000	-5.9598	-2.7202
	EEDS_100	-4.82000*	.48225	.000	-6.4398	-3.2002
	EEDS_200	-4.80667*	.48225	.000	-6.4265	-3.1868
	EEDS_400	-4.82333*	.48225	.000	-6.4432	-3.2035
EEDS_100	KONTROL NORMAL	1.14333	.48225	.240	-.4765	2.7632
	KONTROL POSITIF	.48000	.48225	.911	-1.1398	2.0998
	KONTROL NEGATIF	4.82000*	.48225	.000	3.2002	6.4398
	EEDS_200	.01333	.48225	1.000	-1.6065	1.6332
	EEDS_400	-.00333	.48225	1.000	-1.6232	1.6165
EEDS_200	KONTROL NORMAL	1.13000	.48225	.250	-.4898	2.7498
	KONTROL POSITIF	.46667	.48225	.920	-1.1532	2.0865
	KONTROL NEGATIF	4.80667*	.48225	.000	3.1868	6.4265
	EEDS_100	-.01333	.48225	1.000	-1.6332	1.6065
	EEDS_400	-.01667	.48225	1.000	-1.6365	1.6032
EEDS_400	KONTROL NORMAL	1.14667	.48225	.238	-.4732	2.7665
	KONTROL POSITIF	.48333	.48225	.909	-1.1365	2.1032
	KONTROL NEGATIF	4.82333*	.48225	.000	3.2035	6.4432
	EEDS_100	.00333	.48225	1.000	-1.6165	1.6232
	EEDS_200	.01667	.48225	1.000	-1.6032	1.6365

*. The mean difference is significant at the 0.05 level.

Homogeneous Subsets

PROBE

PERLAKUAN		N	Subset for alpha = 0.05	
			1	2
Tukey HSD ^a	KONTROL NEGATIF	5	-2.8933	
	KONTROL NORMAL	5		.7833
	KONTROL POSITIF	5		1.4467
	EEDS_200	5		1.9133
	EEDS_100	5		1.9267
	EEDS_400	5		1.9300
	Sig.		1.000	.238

Means for groups in homogeneous subsets are displayed.

a. Uses Harmonic Mean Sample Size = 3.000.

3. UJI SENSORIS MOTORIS

ANOVA

SENSORIS

	Sum of Squares	df	Mean Square	F	Sig.
Between Groups	30.732	5	6.146	49.457	.000
Within Groups	1.491	24	.124		
Total	32.223	29			

Post Hoc Tests

Multiple Comparisons

Dependent Variable: SENSORIS

	(I) PERLAK UAN	(J) PERLAKU AN	Mean Difference (I-J)	Std. Error	Sig.	95% Confidence Interval	
						Lower Bound	Upper Bound
Tukey HSD	NORMAL	POSITIF	.13000	.28784	.997	-.8368	1.0968
		NEGATIF	-3.60667*	.28784	.000	-4.5735	-2.6398
		EEDS100	-.86000	.28784	.092	-1.8268	.1068
		EEDS200	-.36000	.28784	.805	-1.3268	.6068
		EEDS400	.01667	.28784	1.000	-.9502	.9835
	POSITIF	NORMAL	-.13000	.28784	.997	-1.0968	.8368
		NEGATIF	-3.73667*	.28784	.000	-4.7035	-2.7698
		EEDS100	-.99000*	.28784	.044	-1.9568	-.0232
		EEDS200	-.49000	.28784	.555	-1.4568	.4768
		EEDS400	-.11333	.28784	.998	-1.0802	.8535
	NEGATIF	NORMAL	3.60667*	.28784	.000	2.6398	4.5735
		POSITIF	3.73667*	.28784	.000	2.7698	4.7035
		EEDS100	2.74667*	.28784	.000	1.7798	3.7135
		EEDS200	3.24667*	.28784	.000	2.2798	4.2135
		EEDS400	3.62333*	.28784	.000	2.6565	4.5902
	EEDS100	NORMAL	.86000	.28784	.092	-.1068	1.8268

	POSITIF	.99000	.28784	.044	.0232	1.9568
	NEGATIF	-2.74667*	.28784	.000	-3.7135	-1.7798
	EEDS200	.50000	.28784	.535	-.4668	1.4668
	EEDS400	.87667	.28784	.084	-.0902	1.8435
EEDS200	NORMAL	.36000	.28784	.805	-.6068	1.3268
	POSITIF	.49000	.28784	.555	-.4768	1.4568
	NEGATIF	-3.24667*	.28784	.000	-4.2135	-2.2798
	EEDS100	-.50000	.28784	.535	-1.4668	.4668
	EEDS400	.37667	.28784	.775	-.5902	1.3435
EEDS400	NORMAL	-.01667	.28784	1.000	-.9835	.9502
	POSITIF	.11333	.28784	.998	-.8535	1.0802
	NEGATIF	-3.62333*	.28784	.000	-4.5902	-2.6565
	EEDS100	-.87667	.28784	.084	-1.8435	.0902
	EEDS200	-.37667	.28784	.775	-1.3435	.5902

*. The mean difference is significant at the 0.05 level.

Homogeneous Subsets

SENSORIS

	PERLAKUAN	N	Subset for alpha = 0.05		
			1	2	3
Tukey HSD ^a	POSITIF	5	.3033		
	EEDS400	5	.4167	.4167	
	NORMAL	5	.4333	.4333	
	EEDS200	5	.7933	.7933	
	EEDS100	5		1.2933	
	NEGATIF	5			4.0400
	Sig.		.555	.084	1.000

Means for groups in homogeneous subsets are displayed.

a. Uses Harmonic Mean Sample Size = 3.000.

4. KADAR MDA

ANOVA

KADAR_MDA

	Sum of Squares	df	Mean Square	F	Sig.
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Between Groups	7.563	5	1.513	102.282	.000
Within Groups	.355	24	.015		
Total	7.918	29			

Post Hoc Tests

Multiple Comparisons

Dependent Variable: KADAR_MDA

	(I) PERLAKUAN	(J) PERLAKUAN	Mean Difference (I-J)	Std. Error	Sig.	95% Confidence Interval	
						Lower Bound	Upper Bound
Tukey HSD	KONTROL NORMAL	KONTROL POSITIF	.36400 [*]	.07691	.001	.1262	.6018
		KONTROL NEGATIF	-1.13600 [*]	.07691	.000	-1.3738	-.8982
		EEDS 100	-.21400	.07691	.095	-.4518	.0238
		EEDS 200	-.07600	.07691	.917	-.3138	.1618
		EEDS 400	.35000 [*]	.07691	.002	.1122	.5878
	KONTROL POSITIF	KONTROL NORMAL	-.36400 [*]	.07691	.001	-.6018	-.1262
		KONTROL NEGATIF	-1.50000 [*]	.07691	.000	-1.7378	-1.2622
		EEDS 100	-.57800 [*]	.07691	.000	-.8158	-.3402
		EEDS 200	-.44000 [*]	.07691	.000	-.6778	-.2022
		EEDS 400	-.01400	.07691	1.000	-.2518	.2238
	KONTROL NEGATIF	KONTROL NORMAL	1.13600 [*]	.07691	.000	.8982	1.3738
		KONTROL POSITIF	1.50000 [*]	.07691	.000	1.2622	1.7378
		EEDS 100	.92200 [*]	.07691	.000	.6842	1.1598
		EEDS 200	1.06000 [*]	.07691	.000	.8222	1.2978
		EEDS 400	1.48600 [*]	.07691	.000	1.2482	1.7238
	EEDS 100	KONTROL NORMAL	.21400	.07691	.095	-.0238	.4518
		KONTROL POSITIF	.57800 [*]	.07691	.000	.3402	.8158
		KONTROL NEGATIF	-.92200 [*]	.07691	.000	-1.1598	-.6842
		EEDS 200	.13800	.07691	.488	-.0998	.3758
		EEDS 400	.56400 [*]	.07691	.000	.3262	.8018
	EEDS 200	KONTROL NORMAL	.07600	.07691	.917	-.1618	.3138
		KONTROL POSITIF	.44000 [*]	.07691	.000	.2022	.6778

	KONTROL NEGATIF	-1.0600 [*]	.07691	.000	-1.2978	-.8222
	EEDS 100	-.13800	.07691	.488	-.3758	.0998
	EEDS 400	.42600 [*]	.07691	.000	.1882	.6638
EEDS 400	KONTROL NORMAL	-.35000 [*]	.07691	.002	-.5878	-.1122
	KONTROL POSITIF	.01400	.07691	1.000	-.2238	.2518
	KONTROL NEGATIF	-1.48600 [*]	.07691	.000	-1.7238	-1.2482
	EEDS 100	-.56400 [*]	.07691	.000	-.8018	-.3262
	EEDS 200	-.42600 [*]	.07691	.000	-.6638	-.1882

*. The mean difference is significant at the 0.05 level.

Homogeneous Subsets

KADAR_MDA

PERLAKUAN		N	Subset for alpha = 0.05		
			1	2	3
Tukey HSD ^a	KONTROL POSITIF	5	.3180		
	EEDS 400	5	.3320		
	KONTROL NORMAL	5		.6820	
	EEDS 200	5		.7580	
	EEDS 100	5		.8960	
	KONTROL NEGATIF	5			1.8180
	Sig.		1.000	.095	1.000

5. KADAR CA1

ANOVA

KADAR_CA1

	Sum of Squares	df	Mean Square	F	Sig.
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Between Groups	1381.900	5	276.380	88.678	.000
Within Groups	74.800	24	3.117		
Total	1456.700	29			

Post Hoc Tests

Multiple Comparisons

Dependent Variable: KADAR_CA1

		Mean Difference (I-J)	Std. Error	Sig.	95% Confidence Interval	
(I) PERLAKUAN	(J) PERLAKUAN				Lower Bound	Upper Bound
Tukey HSD NORMAL	POSITIF	.20000	1.11654	1.000	-3.2523	3.6523
	NEGATIF	-17.80000	1.11654	.000	-21.2523	-14.3477
	EEDS_100	-11.60000	1.11654	.000	-15.0523	-8.1477
	EES_200	-5.80000	1.11654	.000	-9.2523	-2.3477
	EEDS_400	-.40000	1.11654	.999	-3.8523	3.0523
	POSITIF	-.20000	1.11654	1.000	-3.6523	3.2523
	NEGATIF	-18.00000	1.11654	.000	-21.4523	-14.5477
	EEDS_100	-11.80000	1.11654	.000	-15.2523	-8.3477
	EES_200	-6.00000	1.11654	.000	-9.4523	-2.5477
	EEDS_400	-.60000	1.11654	.994	-4.0523	2.8523
	NEGATIF	17.80000	1.11654	.000	14.3477	21.2523
	POSITIF	18.00000	1.11654	.000	14.5477	21.4523
	EEDS_100	6.20000	1.11654	.000	2.7477	9.6523
	EES_200	12.00000	1.11654	.000	8.5477	15.4523
	EEDS_400	17.40000	1.11654	.000	13.9477	20.8523
	EEDS_100	11.60000	1.11654	.000	8.1477	15.0523
	POSITIF	11.80000	1.11654	.000	8.3477	15.2523
	NEGATIF	-6.20000	1.11654	.000	-9.6523	-2.7477
	EES_200	5.80000	1.11654	.000	2.3477	9.2523
	EEDS_400	11.20000	1.11654	.000	7.7477	14.6523
	EES_200	5.80000	1.11654	.000	2.3477	9.2523
	POSITIF	6.00000	1.11654	.000	2.5477	9.4523
	NEGATIF	-12.00000	1.11654	.000	-15.4523	-8.5477
	EEDS_100	-5.80000	1.11654	.000	-9.2523	-2.3477
	EEDS_400	5.40000	1.11654	.001	1.9477	8.8523
	EEDS_400	.40000	1.11654	.999	-3.0523	3.8523
	POSITIF	.60000	1.11654	.994	-2.8523	4.0523
	NEGATIF	-17.40000	1.11654	.000	-20.8523	-13.9477

EEDS_100	-11.20000*	1.11654	.000	-14.6523	-7.7477
EES_200	-5.40000*	1.11654	.001	-8.8523	-1.9477

*. The mean difference is significant at the 0.05 level.

Homogeneous Subsets

KADAR_CA1

PERLAKUAN		N	Subset for alpha = 0.05			
			1	2	3	4
Tukey HSD ^a	POSITIF	5	2.0000			
	NORMAL	5	2.2000			
	EEDS_400	5	2.6000			
	EES_200	5		8.0000		
	EEDS_100	5			13.8000	
	NEGATIF	5				20.0000
	Sig.		.994	1.000	1.000	1.000

Means for groups in homogeneous subsets are displayed.

a. Uses Harmonic Mean Sample Size = 5.000.

6. KADAR CA2-CA3

ANOVA

KADAR_CA2

	Sum of Squares	df	Mean Square	F	Sig.
Between Groups	845.867	5	169.173	59.014	.000
Within Groups	68.800	24	2.867		
Total	914.667	29			

Post Hoc Tests

Multiple Comparisons

Dependent Variable: KADAR_CA2

(I) PERLAKUAN	(J) PERLAKUAN	Mean Difference (I-J)	Std. Error	Sig.	95% Confidence Interval	
					Lower Bound	Upper Bound
Tukey HSD	NORMAL POSITIF	.20000	1.07083	1.000	-3.1109	3.5109
	NEGATIF	-13.00000*	1.07083	.000	-16.3109	-9.6891
	EEDS_100	-10.80000*	1.07083	.000	-14.1109	-7.4891

	EES_200	-5.40000 [*]	1.07083	.000	-8.7109	-2.0891
	EEDS_400	-.60000	1.07083	.993	-3.9109	2.7109
POSITIF	NORMAL	-.20000	1.07083	1.000	-3.5109	3.1109
	NEGATIF	-13.20000 [*]	1.07083	.000	-16.5109	-9.8891
	EEDS_100	-11.00000 [*]	1.07083	.000	-14.3109	-7.6891
	EES_200	-5.60000 [*]	1.07083	.000	-8.9109	-2.2891
	EEDS_400	-.80000	1.07083	.974	-4.1109	2.5109
NEGATIF	NORMAL	13.00000 [*]	1.07083	.000	9.6891	16.3109
	POSITIF	13.20000 [*]	1.07083	.000	9.8891	16.5109
	EEDS_100	2.20000	1.07083	.343	-1.1109	5.5109
	EES_200	7.60000 [*]	1.07083	.000	4.2891	10.9109
	EEDS_400	12.40000 [*]	1.07083	.000	9.0891	15.7109
EEDS_100	NORMAL	10.80000 [*]	1.07083	.000	7.4891	14.1109
	POSITIF	11.00000 [*]	1.07083	.000	7.6891	14.3109
	NEGATIF	-2.20000	1.07083	.343	-5.5109	1.1109
	EES_200	5.40000 [*]	1.07083	.000	2.0891	8.7109
	EEDS_400	10.20000 [*]	1.07083	.000	6.8891	13.5109
EES_200	NORMAL	5.40000 [*]	1.07083	.000	2.0891	8.7109
	POSITIF	5.60000 [*]	1.07083	.000	2.2891	8.9109
	NEGATIF	-7.60000 [*]	1.07083	.000	-10.9109	-4.2891
	EEDS_100	-5.40000 [*]	1.07083	.000	-8.7109	-2.0891
	EEDS_400	4.80000 [*]	1.07083	.002	1.4891	8.1109
EEDS_400	NORMAL	.60000	1.07083	.993	-2.7109	3.9109
	POSITIF	.80000	1.07083	.974	-2.5109	4.1109
	NEGATIF	-12.40000 [*]	1.07083	.000	-15.7109	-9.0891
	EEDS_100	-10.20000 [*]	1.07083	.000	-13.5109	-6.8891
	EES_200	-4.80000 [*]	1.07083	.002	-8.1109	-1.4891

Homogeneous Subsets

KADAR_CA2

PERLAKUAN		N	Subset for alpha = 0.05		
			1	2	3
Tukey HSD ^a	POSITIF	5	2.2000		
	NORMAL	5	2.4000		
	EEDS_400	5	3.0000		
	EES_200	5		7.8000	
	EEDS_100	5			13.2000
	NEGATIF	5			15.4000
	Sig.		.974	1.000	.343

Means for groups in homogeneous subsets are displayed.

- a. Uses Harmonic Mean Sample Size = 5.000.