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LAMPIRAN

Lampiran 1. Ethical Clearance

KEMENTERIAN PENDIDIKAN, KEBUDAYAAN, RISET DAN TEKNOLOGI
UNIVERSITAS HASANUDDIN FAKULTAS KEDOKTERAN
KOMITE ETIK PENELITIAN UNIVERSITAS HASANUDDIN
RSPTN UNIVERSITAS HASANUDDIN
RSUP Dr. WAHIDIN SUDIROHUSODO MAKASSAR
Sekretariat : Lantai 2 Gedung Laboratorium Terpadu
JL.PERINTIS KEMERDEKAAN KAMPUS TAMALANREA KM.10 MAKASSAR 90245.
Contact Person: dr. Agussalim Bukhari.,MMed,PhD, SpGK TELP. 081241850858, 0411 5780103, Fax : 0411-581431

REKOMENDASI PERSETUJUAN ETIK

Nomor : 483/UN4.6.4.5.31/ PP36/ 2023

Tanggal: 13 Juli 2023

Dengan ini Menyatakan bahwa Protokol dan Dokumen yang Berhubungan Dengan Protokol berikut ini telah mendapatkan Persetujuan Etik :

No Protokol	UH23050359	No Sponsor	
Peneliti Utama	**dr. Anak Agung Ayu Niti, M.Kes**	Sponsor	
Judul Peneliti	Analisis Ekspresi Hubungan Protein Leuchine-Rich Repetas And Immunoglobulin Like Domains-1 (LRIG1) dan Ekspresi Epidermal Growth Factor (EGFR) serta Gambaran Histopatologi pada Tikus Wistar Dengan Displasia Nasofaring yang diberikan Isolate Betanin		
No Versi Protokol	**2**	Tanggal Versi	**12 Juli 2023**
No Versi PSP		Tanggal Versi	
Tempat Penelitian	Laboratorium Farmakologi Universitas Gadjah Mada, Yogyakarta		
Jenis Review	☐ Exempted ☒ Expedited ☐ Fullboard Tanggal	Masa Berlaku **13 Juli 2023** sampai **13 Juli 2024**	Frekuensi review lanjutan
Ketua KEP Universitas Hasanuddin	Nama **Prof.Dr.dr. Suryani As'ad, M.Sc.,Sp.GK (K)**	Tanda tangan	
Sekretaris KEP Universitas Hasanuddin	Nama **dr. Agussalim Bukhari, M.Med.,Ph.D.,Sp.GK (K)**	Tanda tangan	

Kewajiban Peneliti Utama:

- Menyerahkan Amandemen Protokol untuk persetujuan sebelum di implementasikan
- Menyerahkan Laporan SAE ke Komisi Etik dalam 24 Jam dan dilengkapi dalam 7 hari dan Lapor SUSAR dalam 72 Jam setelah Peneliti Utama menerima laporan
- Menyerahkan Laporan Kemajuan (progress report) setiap 6 bulan untuk penelitian resiko tinggi dan setiap setahun untuk penelitian resiko rendah
- Menyerahkan laporan akhir setelah Penelitian berakhir
- Melaporkan penyimpangan dari prokol yang disetujui (protocol deviation / violation)
- Mematuhi semua peraturan yang ditentukan

Lampiran 2. Case Report Form

Case Report Form

Research Title: Expression Analysis of Leuchine-Rich Repeats and Immunoglobulin-Like Domains-1 (LRIG1) protein and Epidermal Growth Factor (EGFR) Expression with Histopatological Features in Wistar Rats With Nasopharyngeal displasia Given Betanin.
Researcher: dr. Anak Agung Ayu Niti Wedayani, M.Sc
Date Complecated: (Month/date/years) **: …/…/….**

Animal Information									
Type of animal:	(…) ***Rattus norvegicus***	(…) **strain wistar**							
Number of animal	(…) One	(…) Multiple ……	Species ……	Breed ……	Color ……..	Weight ……..	Sex ……	Age …….	Name ……..
Anatomical abnormalities	Eyes: Dextra ……. Sinistra …….	Nose:	Mouth	Feet Dextra …….. …….. Sinistra ……. …….	Tail				
Animal Owner									
First name ……..	Last Name ……								
Address	City: ……..				Zip:				
Phone	Email:								
Animal Location									
First name ……..	Last Name ……								
Address	City: ……..				Zip:				
Phone	Email:								
Animal Origin									
Name of veterinarian ……..	Facility name: ……								
Address	City: ……..				Zip:				
Phone	Email:								
History (please provide any details about this animal, including travel history or vaccine history if applicable)									
Was the animal imported from outside? (…) Yes (…) No If Yes, from where?									
Clinical Signs/Physical Examination Finding									

Onset date:	Presentation Date:			Date of death:
General	(…) Normal	(…) Abnormal	Comments	
Skin	(…) Normal	(…) Abnormal	Comments	
Eyes	(…) Normal	(…) Abnormal	Comments	
Ears	(…) Normal	(…) Abnormal	Comments	
Nose	(…) Normal	(…) Abnormal	Comments	
Throat	(…) Normal	(…) Abnormal	Comments	
Respiratory	(…) Normal	(…) Abnormal	Comments	
Cardiovascular	(…) Normal	(…) Abnormal	Comments	
Abdomen/ Digestive	(…) Normal	(…) Abnormal	Comments	
Urogenital	(…) Normal	(…) Abnormal	Comments	
Musculoskeletal	(…) Normal	(…) Abnormal	Comments	
Nervous system	(…) Normal	(…) Abnormal	Comments	
Lymph nodes	(…) Normal	(…) Abnormal	Comments	
Other	(…) Normal	(…) Abnormal	Comments	
ELISA (LRIG1) Pre induction formaldehyde:				
Presentation date:				
ELISA (LRIG1) Post induction formaldehyde:				
Presentation date:				
ELISA (Egf) Pre induction formaldehyde:				
Presentation date:				
ELISA (Egf) Post induction formaldehyde:				
Presentation date:				
IHC (LRIG1) :				
Presentation date:				

IHC (EGFR) :
Presentation date:
Histopathology features of Nasopharyng:
Presentation date:
Treatment (please describe any treatments or medications given)
Date: ……. Treatment (name, strength/dose, duration given): ……
Additional Comments

Lampiran 3. Hasil Analisis Data Displasia

```
Your temporary usage period for IBM SPSS Statistics will expire in
4443 days.

NEW FILE.
DATASET NAME DataSet1 WINDOW=FRONT.
CROSSTABS
  /TABLES=KelompokPerlakuan BY Nasofaring Paru
  /FORMAT=AVALUE TABLES
  /STATISTICS=CHISQ
  /CELLS=COUNT
  /COUNT ROUND CELL.
```

Crosstabs

Notes

Output Created		01-NOV-2023 22:35:31
Comments		
Input	Active Dataset	DataSet1
	Filter	<none>
	Weight	<none>
	Split File	<none>
	N of Rows in Working Data File	30
Missing Value Handling	Definition of Missing	User-defined missing values are treated as missing.
	Cases Used	Statistics for each table are based on all the cases with valid data in the specified range(s) for all variables in each table.
Syntax		CROSSTABS /TABLES=KelompokPerlakuan BY Nasofaring Paru /FORMAT=AVALUE TABLES /STATISTICS=CHISQ /CELLS=COUNT /COUNT ROUND CELL.
Resources	Processor Time	00:00:00.00

	Elapsed Time	00:00:00.00
	Dimensions Requested	2
	Cells Available	524245

[DataSet1]

Case Processing Summary

	Cases					
	Valid		Missing		Total	
	N	Percent	N	Percent	N	Percent
Kelompok perlakuan * Derajat displasia nasofaring	30	100.0%	0	0.0%	30	100.0%
Kelompok perlakuan * Derajat displasia paru-paru	30	100.0%	0	0.0%	30	100.0%

Kelompok perlakuan * Derajat displasia nasofaring

Crosstab

Count

		Derajat displasia nasofaring			
		.00	2.00	3.00	Total
Kelompok perlakuan	Sehat	5	0	0	5
	Kontrol negatif	0	0	5	5
	Kontrol positif	0	0	5	5
	Perlakuan 1	0	0	5	5
	Perlakuan 2	1	0	4	5
	Perlakuan 3	2	2	1	5
Total		8	2	20	30

Chi-Square Tests

	Value	df	Asymptotic Significance (2-sided)
Pearson Chi-Square	32.100[a]	10	.000
Likelihood Ratio	32.646	10	.000
Linear-by-Linear Association	1.088	1	.297
N of Valid Cases	30		

a. 18 cells (100.0%) have expected count less than 5. The minimum expected count is .33.

Lampiran 4. Hasil Analisis LRIG1 dan EGFR

Descriptives

			Statistic	Std. Error
Lapisan mukosa LRIG1	Mean		.6613	.05725
	95% Confidence Interval for Mean	Lower Bound	.5443	
		Upper Bound	.7784	
	5% Trimmed Mean		.6756	
	Median		.8400	
	Variance		.098	
	Std. Deviation		.31355	
	Minimum		.11	
	Maximum		.96	
	Range		.85	
	Interquartile Range		.62	
	Skewness		-.749	.427
	Kurtosis		-1.194	.833
Lapisan mukosa EFGR	Mean		.3537	.06408
	95% Confidence Interval for Mean	Lower Bound	.2226	
		Upper Bound	.4847	
	5% Trimmed Mean		.3411	
	Median		.2150	
	Variance		.123	
	Std. Deviation		.35096	
	Minimum		.00	
	Maximum		.94	
	Range		.94	
	Interquartile Range		.74	
	Skewness		.665	.427
	Kurtosis		-1.287	.833

Tests of Normality

	Kolmogorov-Smirnov[a]	Shapiro-Wilk

Descriptives

	Kelompok perlakuan			Statistic	Std. Error
Lapisan mukosa LRIG1	Sehat	Mean		.9140	.01600
		95% Confidence Interval for Mean	Lower Bound	.8696	
			Upper Bound	.9584	
		5% Trimmed Mean		.9150	
		Median		.9200	
		Variance		.001	
		Std. Deviation		.03578	
		Minimum		.86	
		Maximum		.95	
		Range		.09	
		Interquartile Range		.06	
		Skewness		-.871	.913
		Kurtosis		.148	2.000
	Kontrol negatif	Mean		.1820	.02267
		95% Confidence Interval for Mean	Lower Bound	.1191	
			Upper Bound	.2449	
		5% Trimmed Mean		.1833	
		Median		.2000	
		Variance		.003	
		Std. Deviation		.05070	
		Minimum		.11	
		Maximum		.23	
		Range		.12	
		Interquartile Range		.10	
		Skewness		-.751	.913
		Kurtosis		-1.296	2.000
	Kontrol positif	Mean		.3240	.07534
		95% Confidence Interval for Mean	Lower Bound	.1148	
			Upper Bound	.5332	
		5% Trimmed Mean		.3244	
		Median		.3000	
		Variance		.028	
		Std. Deviation		.16846	
		Minimum		.11	
		Maximum		.53	
		Range		.42	
		Interquartile Range		.32	
		Skewness		.008	.913

		Kurtosis		-1.437	2.000
	Perlakuan 1	Mean		.7440	.05391
		95% Confidence Interval for Mean	Lower Bound	.5943	
			Upper Bound	.8937	
		5% Trimmed Mean		.7467	
		Median		.7600	
		Variance		.015	
		Std. Deviation		.12054	
		Minimum		.56	
		Maximum		.88	
		Range		.32	
		Interquartile Range		.21	
		Skewness		-.824	.913
		Kurtosis		.933	2.000
	Perlakuan 2	Mean		.8780	.01562
		95% Confidence Interval for Mean	Lower Bound	.8346	
			Upper Bound	.9214	
		5% Trimmed Mean		.8800	
		Median		.9000	
		Variance		.001	
		Std. Deviation		.03493	
		Minimum		.82	
		Maximum		.90	
		Range		.08	
		Interquartile Range		.06	
		Skewness		-1.600	.913
		Kurtosis		2.098	2.000
	Perlakuan 3	Mean		.9260	.01249
		95% Confidence Interval for Mean	Lower Bound	.8913	
			Upper Bound	.9607	
		5% Trimmed Mean		.9256	
		Median		.9200	
		Variance		.001	
		Std. Deviation		.02793	
		Minimum		.90	
		Maximum		.96	
		Range		.06	

		Interquartile Range		.05	
		Skewness		.340	.913
		Kurtosis		-2.692	2.000
Lapisan mukosa EFGR	Sehat	Mean		.0000	.00000
		95% Confidence Interval for Mean	Lower Bound	.0000	
			Upper Bound	.0000	
		5% Trimmed Mean		.0000	
		Median		.0000	
		Variance		.000	
		Std. Deviation		.00000	
		Minimum		.00	
		Maximum		.00	
		Range		.00	
		Interquartile Range		.00	
		Skewness		.	.
		Kurtosis		.	.
	Kontrol negatif	Mean		.8760	.03124
		95% Confidence Interval for Mean	Lower Bound	.7893	
			Upper Bound	.9627	
		5% Trimmed Mean		.8767	
		Median		.9200	
		Variance		.005	
		Std. Deviation		.06986	
		Minimum		.80	
		Maximum		.94	
		Range		.14	
		Interquartile Range		.13	
		Skewness		-.544	.913
		Kurtosis		-3.224	2.000
	Kontrol positif	Mean		.7580	.06629
		95% Confidence Interval for Mean	Lower Bound	.5740	
			Upper Bound	.9420	
		5% Trimmed Mean		.7633	
		Median		.7900	
		Variance		.022	
		Std. Deviation		.14822	
		Minimum		.51	

	Maximum		.91	
	Range		.40	
	Interquartile Range		.21	
	Skewness		-1.487	.913
	Kurtosis		3.190	2.000
Perlakuan 1	Mean		.2220	.03470
	95% Confidence Interval for Mean	Lower Bound	.1257	
		Upper Bound	.3183	
	5% Trimmed Mean		.2244	
	Median		.2400	
	Variance		.006	
	Std. Deviation		.07759	
	Minimum		.10	
	Maximum		.30	
	Range		.20	
	Interquartile Range		.14	
	Skewness		-1.102	.913
	Kurtosis		1.113	2.000
Perlakuan 2	Mean		.2040	.01720
	95% Confidence Interval for Mean	Lower Bound	.1562	
		Upper Bound	.2518	
	5% Trimmed Mean		.2044	
	Median		.2000	
	Variance		.001	
	Std. Deviation		.03847	
	Minimum		.15	
	Maximum		.25	
	Range		.10	
	Interquartile Range		.07	
	Skewness		-.332	.913
	Kurtosis		-.310	2.000
Perlakuan 3	Mean		.0620	.01356
	95% Confidence Interval for Mean	Lower Bound	.0243	
		Upper Bound	.0997	
	5% Trimmed Mean		.0622	
	Median		.0600	
	Variance		.001	

	Std. Deviation	.03033	
	Minimum	.02	
	Maximum	.10	
	Range	.08	
	Interquartile Range	.05	
	Skewness	-.226	.913
	Kurtosis	-.139	2.000

	Statistic	df	Sig.	Statistic	df	Sig.
Lapisan mukosa LRIG1	.249	30	.000	.791	30	.000
Lapisan mukosa EFGR	.228	30	.000	.813	30	.000

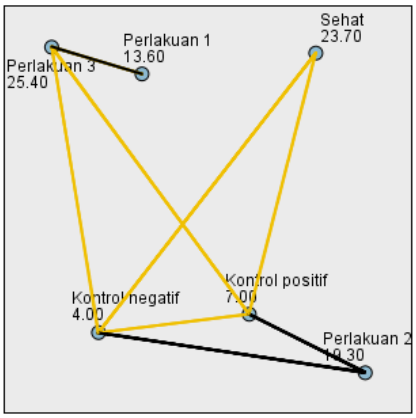
a. Lilliefors Significance Correction

Tes Normalitas

	Lapisan mukosa LRIG1	Lapisan mukosa EFGR
Kruskal-Wallis H	25.238	27.219
df	5	5
Asymp. Sig.	.000	.000

LM LGIR1

Pairwise Comparisons of Kelompok perlakuan



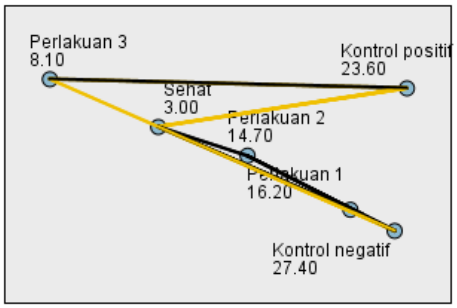
Each node shows the sample average rank of Kelompok perlakuan.

Sample1-Sample2	Test Statistic	Std. Error	Std. Test Statistic	Sig.	Adj.Sig.
Kontrol negatif-Kontrol positif	-3.000	5.544	-.541	.588	1.000
Kontrol negatif-Perlakuan 1	-9.600	5.544	-1.732	.083	1.000
Kontrol negatif-Perlakuan 2	-15.300	5.544	-2.760	.006	.087
Kontrol negatif-Sehat	19.700	5.544	3.554	.000	.006
Kontrol negatif-Perlakuan 3	-21.400	5.544	-3.860	.000	.002
Kontrol positif-Perlakuan 1	-6.600	5.544	-1.191	.234	1.000
Kontrol positif-Perlakuan 2	-12.300	5.544	-2.219	.027	.398
Kontrol positif-Sehat	16.700	5.544	3.013	.003	.039
Kontrol positif-Perlakuan 3	-18.400	5.544	-3.319	.001	.014
Perlakuan 1-Perlakuan 2	-5.700	5.544	-1.028	.304	1.000
Perlakuan 1-Sehat	10.100	5.544	1.822	.068	1.000
Perlakuan 1-Perlakuan 3	-11.800	5.544	-2.129	.033	.499
Perlakuan 2-Sehat	4.400	5.544	.794	.427	1.000
Perlakuan 2-Perlakuan 3	-6.100	5.544	-1.100	.271	1.000
Sehat-Perlakuan 3	-1.700	5.544	-.307	.759	1.000

Each row tests the null hypothesis that the Sample 1 and Sample 2 distributions are the same.
Asymptotic significances (2-sided tests) are displayed. The significance level is .05.
Significance values have been adjusted by the Bonferroni correction for multiple tests.

LM EFGR

Pairwise Comparisons of Kelompok perlakuan



Each node shows the sample average rank of Kelompok perlakuan.

Sample1-Sample2	Test Statistic	Std. Error	Std. Test Statistic	Sig.	Adj.Sig.
Sehat-Perlakuan 3	-5.100	5.551	-.919	.358	1.000
Sehat-Perlakuan 2	-11.700	5.551	-2.108	.035	.526
Sehat-Perlakuan 1	-13.200	5.551	-2.378	.017	.261
Sehat-Kontrol positif	-20.600	5.551	-3.711	.000	.003
Sehat-Kontrol negatif	-24.400	5.551	-4.396	.000	.000
Perlakuan 3-Perlakuan 2	6.600	5.551	1.189	.234	1.000
Perlakuan 3-Perlakuan 1	8.100	5.551	1.459	.145	1.000
Perlakuan 3-Kontrol positif	15.500	5.551	2.792	.005	.079
Perlakuan 3-Kontrol negatif	19.300	5.551	3.477	.001	.008
Perlakuan 2-Perlakuan 1	1.500	5.551	.270	.787	1.000
Perlakuan 2-Kontrol positif	8.900	5.551	1.603	.109	1.000
Perlakuan 2-Kontrol negatif	12.700	5.551	2.288	.022	.332
Perlakuan 1-Kontrol positif	7.400	5.551	1.333	.183	1.000
Perlakuan 1-Kontrol negatif	11.200	5.551	2.018	.044	.654
Kontrol positif-Kontrol negatif	3.800	5.551	.685	.494	1.000

Each row tests the null hypothesis that the Sample 1 and Sample 2 distributions are the same.
Asymptotic significances (2-sided tests) are displayed. The significance level is .05.
Significance values have been adjusted by the Bonferroni correction for multiple tests.