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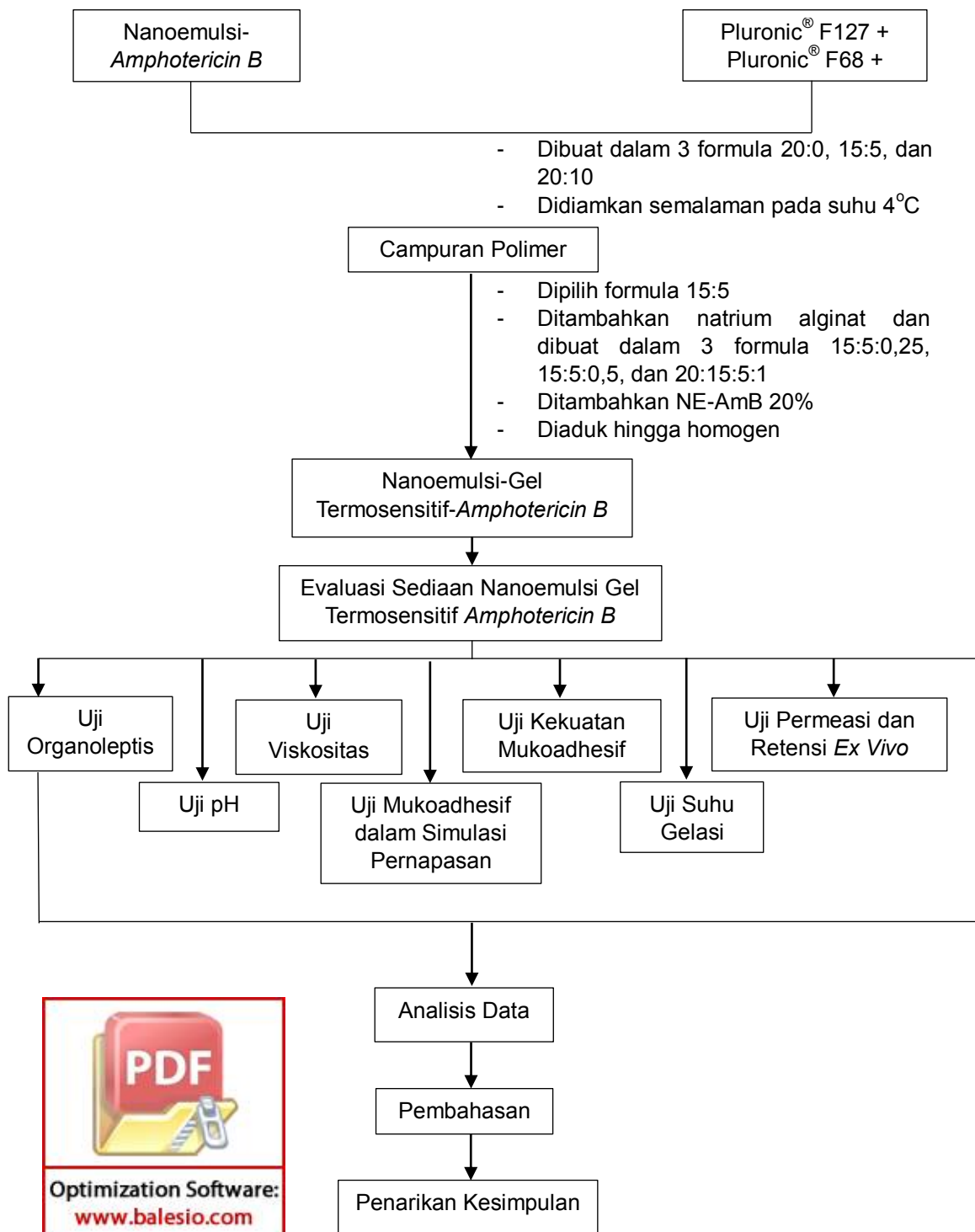
Zamir, A., Hussain, I., ur Rehman, A., Ashraf, W., Imran, I., Saeed, H., Majeed, A., Alqahtani, F., & Rasool, M. F. 2022. Clinical Pharmacokinetics of Metoprolol: A Systematic Review. *Clinical Pharmacokinetics*. 61(8), 1095–1114. <https://doi.org/10.1007/s40262-022-01145-y>



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LAMPIRAN

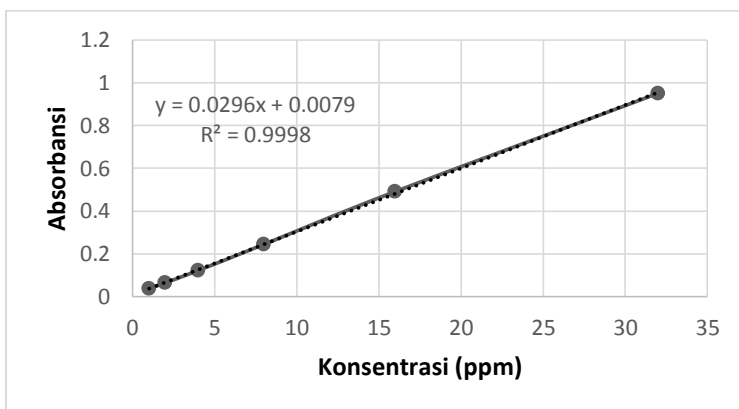
Lampiran 1. Skema Kerja Penelitian



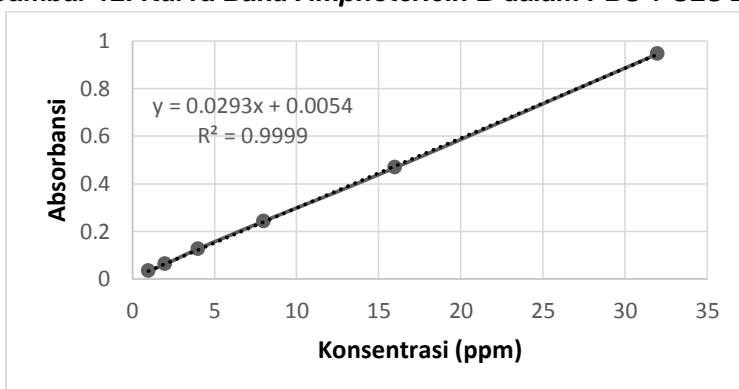
Lampiran 2. Komposisi Formula Nanoemulsi-*Amphotericin B*

Formula	Komposisi					
	AmB	Virgin Coconut Oil	Transcutol	Tween 80	Gliserin	Aquadest
NE1	10 mg	15 mL	5 mL	5 mL	5 mL	20 mL
NE2	10 mg	15 mL	10 mL	5 mL	5 mL	15 mL
NE3	10 mg	15 mL	15 mL	5 mL	10 mL	10 mL

Lampiran 3. Kurva Baku *Amphotericin B* dalam PBS + SLS 2% dan Mukosa Hidung Babi



Gambar 12. Kurva Baku *Amphotericin B* dalam PBS + SLS 2%



Gambar 13. Kurva Baku *Amphotericin B* dalam Mukosa Hidung Babi

Lampiran 4. Perhitungan Data

Lampiran 4.1 Perhitungan Data Uji Kekuatan Mukoadhesif

NE-TG-AmB 6, Replikasi 1

Dik : Bobot yang dibutuhkan = 7 gram
 Luas permukaan = 71,02 cm²
 Percepatan gravitasi = 980 cm/s²
 Kekuatan mukoadhesif = ?



$$\text{Kekuatan mukoadhesif (dyne/cm}^2\text{)} = \frac{m \times g}{A}$$

$$= \frac{7 \text{ gram} \times 980 \text{ cm/s}^2}{71,02 \text{ cm}^2}$$

$$= 96,5925 \text{ dyne/cm}^2$$

Lampiran 4.2 Perhitungan Data Uji Permeasi secara *Ex Vivo*

Keterangan:

x = konsentrasi

y = absorbansi

Untuk NE-TG-AmB replikasi 1 jam ke-7 diperoleh absorbansi = 0,686 dengan faktor pengenceran = 8

$$y = 0,0296x + 0,0079$$

$$0,686 = 0,0296x + 0,0079$$

$$x = \frac{(0,686 - 0,0079)}{0,0296}$$

$$x = 22,90878 \times \text{faktor pengenceran}$$

$$x = 22,90878 \times 8 = 183,27027 \mu\text{g/mL}$$

$$\begin{aligned} \text{Konsentrasi dalam 1 mL} &= 183,27027 \mu\text{g/mL} \times 1 \text{ mL} \\ &= 183,27027 \mu\text{g} \\ &= 0,18327 \text{ mg} \end{aligned}$$

$$\begin{aligned} \text{Konsentrasi dalam 28 mL} &= 0,18327 \text{ mg} \times 28 \text{ mL} \\ &= 5,13156 \text{ mg} \end{aligned}$$

$$\begin{aligned} \text{Faktor koreksi} &= \frac{\text{konsentrasi jam sebelumnya}}{1000} + \text{faktor koreksi jam sebelumnya} \\ &= \frac{15,67905}{1000} + 0,02964 \\ &= 0,04532 \end{aligned}$$

$$\begin{aligned} \text{Jumlah terpermeasi} &= \text{konsentrasi dalam 28 mL} + \text{faktor koreksi} \\ &= 5,13156 + 0,04532 \\ &= 5,17689 \text{ mg} \end{aligned}$$

Lampiran 4.3 Perhitungan Data Uji Retensi secara *Ex Vivo*

Persamaan: $y = 0,0293x + 0,0054$

Keterangan:

x = konsentrasi

y = absorbansi

Untuk NE-TG-AmB replikasi 1 diperoleh absorbansi = 1,382



Jumlah AmB yang terdeposisi = $\frac{46,9833}{1000} \times 5 = 0,2349165 \text{ mg}$

Lampiran 5. Tabel Hasil Evaluasi

Lampiran 5.1 Hasil Uji pH

Replikasi	Formula					
	NE-TG- AmB 1	NE-TG- AmB 2	NE-TG- AmB 3	NE-TG- AmB 4	NE-TG- AmB 5	NE-TG- AmB 6
1	6,57	6,52	6,65	6,69	6,69	6,41
2	7,09	7,13	7,21	7,18	7,22	7,07
3	7,61	7,74	7,77	7,67	7,75	7,73
Rata-Rata	7,09	7,13	7,21	7,18	7,22	7,07
SD	0,52	0,61	0,56	0,49	0,53	0,66

Lampiran 5.2 Hasil Uji Viskositas

Lampiran 5.2.1 Hasil Uji Viskositas (Suhu 4°C)

Replikasi	Formula					
	NE-TG- AmB 1	NE-TG- AmB 2	NE-TG- AmB 3	NE-TG- AmB 4	NE-TG- AmB 5	NE-TG- AmB 6
1	185,8635	185,6369	156,3025	187,7896	194,7833	230,1638
2	207,4935	204,1769	173,8125	209,4196	217,7729	251,0728
3	229,1235	222,7169	191,3225	231,0496	240,7625	271,9818
Rata-Rata	207,4935	204,1769	173,8125	209,4196	217,7729	251,0728
SD	21,63	18,54	17,51	21,63	22,9896	20,909

Lampiran 5.2.2 Hasil Uji Viskositas (Suhu 25°C)

Replikasi	Formula					
	NE-TG- AmB 1	NE-TG- AmB 2	NE-TG- AmB 3	NE-TG- AmB 4	NE-TG- AmB 5	NE-TG- AmB 6
1	20445,5	503,67	404,79	511,91	566,50	17901,4
2	22042	536,63	432,6	557,23	605,64	19158
3	23638,5	569,59	460,41	602,55	644,78	20414,6
Rata-Rata	22042	536,63	432,6	557,23	605,64	19158
SD	1596,5	32,96	27,81	45,32	39,14	1256,6

Lampiran 5.2.3 Hasil Uji Viskositas (Suhu 37°C)

Replikasi	Formula					
	NE-TG- AmB 1	NE-TG- AmB 2	NE-TG- AmB 3	NE-TG- AmB 4	NE-TG- AmB 5	NE-TG- AmB 6
1	23854,8	20826,6	490,28	24060,8	25729,4	27768,8
2	25544	22454	535,6	25338	27089	29252
3	27223,2	240814	580,92	26615,2	28448,6	30735,2
Rata-Rata	25544	22454	535,6	25338	27089	29252
SD	27223,2	1627,4	45,32	1277,2	1359,6	1483,2



Lampiran 5.3 Hasil Uji Kekuatan Mukoadhesif

Formula	Bobot yang dibutuhkan (g)			Luas Permukaan Mukosa (cm ²)	Percepatan gravitasi (cm/s ²)	Kekuatan Mukoadhesif			Rata-rata	SD
	Replikasi 1	Replikasi 2	Replikasi 3			Replikasi 1	Replikasi 2	Replikasi 3		
NE-TG-AmB 1	2	2	2	71,02	980	20,8196	23,09	25,3604	23,09	2,2704
NE-TG-AmB 2	2	2	2	71,02	980	22,3072	24,34	26,3728	24,34	2,0328
NE-TG-AmB 3	1	2	2	71,02	980	20,5984	22,09	23,5816	22,09	1,4916
NE-TG-AmB 4	2	2	2	71,02	980	28,4104	31,09	33,7696	31,09	2,6796
NE-TG-AmB 5	4	4	5	71,02	980	56,0124	60,54	65,0676	60,54	4,5276
NE-TG-AmB 6	7	7	7	71,02	980	92,1524	99,32	92,1524	99,32	7,1676



Lampiran 5.4 Hasil Uji Mukoadhesif dalam Simulasi Pernapasan

Replikasi	Formula					
	NE-TG- AmB 1	NE-TG- AmB 2	NE-TG- AmB 3	NE-TG- AmB 4	NE-TG- AmB 5	NE-TG- AmB 6
1	65,66	65,26	54,58	63,45	89,50	90,16
2	69,98	70,35	59,45	68,54	98,04	98,94
3	74,30	75,44	64,32	73,63	106,58	107,72
Rata-Rata	69,98	70,35	59,45	68,54	98,04	98,94
SD	4,32	5,09	4,87	5,09	8,54	8,78

Lampiran 5.5 Hasil Uji Suhu Gelasi

Replikasi	Formula					
	NE-TG- AmB 1	NE-TG- AmB 2	NE-TG- AmB 3	NE-TG- AmB 4	NE-TG- AmB 5	NE-TG- AmB 6
1	21,00	32,22	41,11	32,42	31,67	21,82
2	23,09	35,09	44,56	35,43	34,56	24,01
3	25,18	37,96	47,01	38,44	37,45	26,02
Rata-Rata	23,09	35,09	44,56	35,43	34,56	24,01
SD	2,09	2,87	3,45	3,01	2,89	2,19



Lampiran 5.6 Hasil Uji Permeasi secara *Ex Vivo*

NE-TG-AmB

Waktu	Faktor pengenceran	Absorbansi	Konsentrasi (µg/mL)	28 mL (mg)	Faktor koreksi	AmB yang terpermeasi (%)	Rata-rata (%)	SD
0,5	2	0,040	1,08	0,06	0,000	0,60	0,65	0,04
		0,043	1,19	0,07	0,000	0,66		
		0,045	1,25	0,07	0,000	0,70		
1	4	0,037	0,98	0,11	0,001	1,11	1,23	0,10
		0,040	1,08	0,12	0,001	1,23		
		0,043	1,19	0,13	0,001	1,34		
2	8	0,086	2,64	0,59	0,002	5,93	6,59	0,65
		0,094	2,91	0,65	0,002	6,54		
		0,104	3,25	0,73	0,002	7,30		
3	8	0,143	4,56	1,02	0,005	10,27	11,33	1,05
		0,157	5,04	1,13	0,005	11,34		
		0,170	5,48	1,23	0,006	12,33		
4	8	0,265	8,69	1,95	0,009	19,55	21,39	1,87
		0,289	9,50	2,13	0,010	21,37		
		0,315	10,38	2,32	0,011	23,35		
		0,354	11,69	2,62	0,018	26,37	29,27	2,87
		0,392	12,98	2,91	0,020	29,26		
		0,430	14,26	3,20	0,022	32,17		



Waktu	Faktor pengenceran	Absorbansi	Konsentrasi ($\mu\text{g/mL}$)	28 mL (mg)	Faktor koreksi	AmB yang terpermeasi (%)	Rata-rata (%)	SD
6	8	0,472	15,68	3,51	0,030	35,42	39,33	3,90
		0,523	17,40	3,90	0,033	39,31		
		0,575	19,16	4,29	0,036	43,27		
7	8	0,686	22,91	5,13	0,045	51,77	57,26	5,49
		0,758	25,34	5,68	0,050	57,27		
		0,83	27,77	6,22	0,055	62,76		
8	12	0,559	18,62	6,26	0,068	63,24	69,45	6,18
		0,613	20,44	6,87	0,075	69,44		
		0,667	22,27	7,48	0,083	75,64		
12	12	0,780	26,08	8,76	0,087	88,51	97,84	9,35
		0,862	28,85	9,70	0,096	97,91		
		0,943	31,59	10,61	0,105	107,20		
24	16	0,892	29,87	13,38	0,113	134,94	147,26	12,32
		0,973	32,60	14,61	0,125	147,32		
		0,995	33,35	14,94	0,137	150,76		



TG-AmB

Waktu	Faktor pengenceran	Absorbansi	Konsentrasi (µg/mL)	28 mL (mg)	Faktor koreksi	AmB yang terpermeasi (mg)	Rata-rata (mg)	SD
1	1	0,025	0,58	0,02	0,000	0,16	2,44	0,24
		0,026	0,61	0,02	0,000	0,17		
		0,028	0,68	0,02	0,000	0,19		
2	1	0,036	0,95	0,03	0,001	0,27	4,96	0,37
		0,038	1,02	0,03	0,001	0,29		
		0,042	1,15	0,03	0,001	0,33		
3	1	0,095	2,94	0,08	0,002	0,84	6,33	0,56
		0,103	3,21	0,09	0,002	0,92		
		0,115	3,62	0,10	0,002	1,03		
4	1	0,176	5,68	0,16	0,004	1,63	10,86	0,98
		0,187	6,05	0,17	0,005	1,74		
		0,209	6,79	0,19	0,005	1,96		
5	1	0,252	8,25	0,23	0,010	2,41	20,61	1,77
		0,275	9,02	0,25	0,011	2,64		
		0,309	10,17	0,28	0,012	2,97		
		0,315	10,38	0,29	0,018	3,09	27,65	2,61
		0,339	11,19	0,31	0,020	3,33		
		0,340	11,22	0,31	0,022	3,37		



Waktu	Faktor pengenceran	Absorbansi	Konsentrasi ($\mu\text{g/mL}$)	28 mL (mg)	Faktor koreksi	AmB yang terpermeasi (%)	Rata-rata (%)	SD
7	2	0,473	15,71	3,52	0,029	5,72	6,25	0,51
		0,518	17,23	3,86	0,031	6,22		
		0,562	18,72	4,19	0,034	6,81		
8	2	0,689	23,01	5,15	0,038	7,44	8,1	0,72
		0,743	24,83	5,56	0,042	8,23		
		0,798	26,69	5,98	0,045	8,97		
12	2	0,785	26,26	5,88	0,051	9,84	10,69	0,95
		0,877	29,36	6,58	0,056	10,79		
		0,960	32,17	7,21	0,060	11,75		
24	2	1,201	40,31	9,03	0,068	10,80	12,72	2,04
		1,099	36,86	8,26	0,074	12,83		
		1,047	35,10	7,86	0,080	14,88		



Lampiran 5.7 Hasil Uji Retensi secara *Ex Vivo*

NE-TG-AmB

Waktu	Replikasi	Serapan	Konsentrasi (µg/mL)	Faktor Koreksi	Jumlah AmB Terdeposisi Setelah 24 Jam	Rata-rata (mg)	SD
0	1	0	0	30	0	0	0
	2	0	0	30	0		
	3	0	0	30	0		
1	1	0,623	632,03	30	3,16015	3,22336667	0,571453533
	2	0,530	537,23	30	2,68615		
	3	0,752	764,76	30	3,8238		
2	1	0,923	939,94	30	4,6997	4,79368333	0,849831604
	2	0,786	798,95	30	3,99475		
	3	1,116	1137,32	30	5,6866		
3	1	1,087	1106,94	30	5,5347	5,64538333	1,000825834
	2	0,924	940,9	30	4,7045		
	3	1,314	1339,39	30	6,69695		
4	1	1,295	1320,37	30	6,60185	6,7339	1,19381498
	2	1,102	1122,32	30	5,6116		
	3	1,566	1597,65	30	7,98825		
5	1	1,451	1480,04	30	7,4002	7,54818333	1,33817596
	2	1,234	1258,03	30	6,29015		
	3	1,754	1790,84	30	8,9542		



Optimization Software:
www.balesio.com

6	1	0,791	804,76	30	4,0238	4,10425	0,727618325
	2	0,673	684,04	30	3,4202		
	3	0,956	973,75	30	4,86875		
7	1	0,672	682,68	30	3,4134	3,48165	0,617211623
	2	0,572	580,28	30	2,9014		
	3	0,812	826,03	30	4,13015		
8	1	0,669	679,33	30	3,39665	3,46457833	0,614216153
	2	0,569	577,43	30	2,88715		
	3	0,808	821,987	30	4,109935		
12	1	0,561	569,17	30	2,84585	2,90275	0,514614681
	2	0,478	483,79	30	2,41895		
	3	0,678	688,69	30	3,44345		
24	1	0,464	469,83	30	2,34915	2,39611667	0,424801746
	2	0,395	399,35	30	1,99675		
	3	0,561	568,49	30	2,84245		



TG-AmB

Waktu	Replikasi	Serapan	Konsentrasi (µg/mL)	Faktor Koreksi	Jumlah AmB Terdeposisi Setelah 24 Jam	Rata-rata (mg)	SD
0	1	0	0	15	0	0	0
	2	0	0	15	0		
	3	0	0	15	0		
1	1	0,739	375,51	15	1,87755	1,91508333	0,339509596
	2	0,629	319,18	15	1,5959		
	3	0,893	454,36	15	2,2718		
2	1	1,039	529,03	15	2,64515	2,69805	0,478299088
	2	0,884	449,68	15	2,2484		
	3	1,256	640,12	15	3,2006		
3	1	1,297	661,34	15	3,3067	3,37283333	0,597949206
	2	1,103	562,14	15	2,8107		
	3	1,568	800,22	15	4,0011		
4	1	1,585	808,42	15	4,0421	4,12291667	0,730933568
	2	1,348	687,15	15	3,43575		
	3	1,916	978,18	15	4,8909		
	1	1,127	574,26	15	2,8713	2,92871667	0,519211504
	2	0,959	488,12	15	2,4406		
	3	1,363	694,85	15	3,47425		



Optimization Software:
www.balesio.com

6	1	0,746	379,33	15	1,89665	1,93456667	0,342950646
	2	0,635	322,43	15	1,61215		
	3	0,902	458,98	15	2,2949		
7	1	0,364	183,52	15	0,9176	0,93593333	0,16591144
	2	0,310	155,99	15	0,77995		
	3	0,439	222,05	15	1,11025		
8	1	0,055	25,39	15	0,12695	0,12948333	0,022955083
	2	0,048	21,58	15	0,1079		
	3	0,065	30,72	15	0,1536		
12	1	0,014	4,42	15	0,0221	0,02251667	0,003991345
	2	0,013	3,75	15	0,01875		
	3	0,016	5,34	15	0,0267		
24	1	0,012	3,21	15	0,01605	0,01793333	0,008434206
	2	0,010	2,12	15	0,0106		
	3	0,016	5,43	15	0,02715		



Lampiran 6. Data Hasil Uji Analisis Statistik

Lampiran 6.1 Analisis Statistik Uji pH

Tests of Normality							
Formula	Kolmogorov-Smirnov ^a			Shapiro-Wilk			
	Statistic	df	Sig.	Statistic	df	Sig.	
Uji pH	F1	,175	3	.	1,000	3	1,000
	F2	,175	3	.	1,000	3	1,000
	F3	,175	3	.	1,000	3	1,000
	F4	,175	3	.	1,000	3	1,000
	F5	,175	3	.	1,000	3	1,000
	F6	,175	3	.	1,000	3	1,000

a. Lilliefors Significance Correction

ANOVA

Uji pH

	Sum of Squares	df	Mean Square	F	Sig.
Between Groups	,059	5	,012	,037	,999
Within Groups	3,825	12	,319		
Total	3,885	17			



Multiple Comparisons

Dependent Variable: Uji pH

	(I) Formula	(J) Formula	Mean Difference		Sig.	95% Confidence Interval		
			(I-J)	Std. Error		Lower Bound	Upper Bound	
Tukey HSD	F1	F2	-,04000	,46100	1,000	-1,5885	1,5085	
		F3	-,12000	,46100	1,000	-1,6685	1,4285	
		F4	-,09000	,46100	1,000	-1,6385	1,4585	
		F5	-,13000	,46100	1,000	-1,6785	1,4185	
		F6	,02000	,46100	1,000	-1,5285	1,5685	
		F2	F1	,04000	,46100	1,000	-1,5085	1,5885
	F2	F3	-,08000	,46100	1,000	-1,6285	1,4685	
		F4	-,05000	,46100	1,000	-1,5985	1,4985	
		F5	-,09000	,46100	1,000	-1,6385	1,4585	
		F6	,06000	,46100	1,000	-1,4885	1,6085	
		F3	F1	,12000	,46100	1,000	-1,4285	1,6685
		F3	F2	,08000	,46100	1,000	-1,4685	1,6285
	F4		,03000	,46100	1,000	-1,5185	1,5785	
	F5		-,01000	,46100	1,000	-1,5585	1,5385	
	F6		,14000	,46100	1,000	-1,4085	1,6885	
	F1		,09000	,46100	1,000	-1,4585	1,6385	
	F2		,05000	,46100	1,000	-1,4985	1,5985	
	F3		-,03000	,46100	1,000	-1,5785	1,5185	
F5	-,04000		,46100	1,000	-1,5885	1,5085		



	F6	,11000	,46100	1,000	-1,4385	1,6585
F5	F1	,13000	,46100	1,000	-1,4185	1,6785
	F2	,09000	,46100	1,000	-1,4585	1,6385
	F3	,01000	,46100	1,000	-1,5385	1,5585
	F4	,04000	,46100	1,000	-1,5085	1,5885
	F6	,15000	,46100	,999	-1,3985	1,6985
F6	F1	-,02000	,46100	1,000	-1,5685	1,5285
	F2	-,06000	,46100	1,000	-1,6085	1,4885
	F3	-,14000	,46100	1,000	-1,6885	1,4085
	F4	-,11000	,46100	1,000	-1,6585	1,4385
	F5	-,15000	,46100	,999	-1,6985	1,3985



Lampiran 6.2 Analisis Statistik Uji Viskositas

Lampiran 6.2.1 Uji Viskositas (Suhu 4°C)

Tests of Normality

	Formula	Kolmogorov-Smirnov ^a			Shapiro-Wilk		
		Statistic	df	Sig.	Statistic	df	Sig.
Uji Viskositas	F1	,175	3	.	1,000	3	1,000
Suhu 4°C	F2	,175	3	.	1,000	3	1,000
	F3	,175	3	.	1,000	3	1,000
	F4	,175	3	.	1,000	3	1,000
	F5	,175	3	.	1,000	3	1,000
	F6	,175	3	.	1,000	3	1,000

a. Lilliefors Significance Correction

ANOVA

Uji Viskositas Suhu 4°C

	Sum of Squares	df	Mean Square	F	Sig.
Between Groups	9285,343	5	1857,069	4,367	,017
Within Groups	5103,507	12	425,292		
Total	14388,850	17			



Multiple Comparisons

Dependent Variable: Uji Viskositas Suhu 4°C

	(I) Formula	(J) Formula	Mean Difference		Sig.	95% Confidence Interval		
			(I-J)	Std. Error		Lower Bound	Upper Bound	
Tukey HSD	F1	F2	3,31660	16,83829	1,000	-53,2419	59,8751	
		F3	33,68100	16,83829	,395	-22,8775	90,2395	
		F4	-1,92610	16,83829	1,000	-58,4846	54,6324	
		F5	-10,27940	16,83829	,988	-66,8379	46,2791	
		F6	-43,57930	16,83829	,174	-100,1378	12,9792	
		F2	F1	-3,31660	16,83829	1,000	-59,8751	53,2419
	F2	F3	30,36440	16,83829	,498	-26,1941	86,9229	
		F4	-5,24270	16,83829	,999	-61,8012	51,3158	
		F5	-13,59600	16,83829	,961	-70,1545	42,9625	
		F6	-46,89590	16,83829	,128	-103,4544	9,6626	
		F3	F1	-33,68100	16,83829	,395	-90,2395	22,8775
			F2	-30,36440	16,83829	,498	-86,9229	26,1941
	F4		-35,60710	16,83829	,342	-92,1656	20,9514	
	F5		-43,96040	16,83829	,168	-100,5189	12,5981	
	F6		-77,26030*	16,83829	,006	-133,8188	-20,7018	
	F1		1,92610	16,83829	1,000	-54,6324	58,4846	
	F5	F2	5,24270	16,83829	,999	-51,3158	61,8012	
		F3	35,60710	16,83829	,342	-20,9514	92,1656	
F5		-8,35330	16,83829	,995	-64,9118	48,2052		



	F6	-41,65320	16,83829	,206	-98,2117	14,9053
F5	F1	10,27940	16,83829	,988	-46,2791	66,8379
	F2	13,59600	16,83829	,961	-42,9625	70,1545
	F3	43,96040	16,83829	,168	-12,5981	100,5189
	F4	8,35330	16,83829	,995	-48,2052	64,9118
	F6	-33,29990	16,83829	,407	-89,8584	23,2586
F6	F1	43,57930	16,83829	,174	-12,9792	100,1378
	F2	46,89590	16,83829	,128	-9,6626	103,4544
	F3	77,26030*	16,83829	,006	20,7018	133,8188
	F4	41,65320	16,83829	,206	-14,9053	98,2117
	F5	33,29990	16,83829	,407	-23,2586	89,8584

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



Lampiran 6.3 Analisis Statistik Uji Kekuatan Mukoadhesif

	Tests of Normality						
	Formula	Kolmogorov-Smirnov ^a			Shapiro-Wilk		
		Statistic	df	Sig.	Statistic	df	Sig.
Uji Kekuatan	F1	,175	3	.	1,000	3	1,000
Mukoadhesif	F2	,175	3	.	1,000	3	1,000
	F3	,175	3	.	1,000	3	,998
	F4	,175	3	.	1,000	3	,998
	F5	,175	3	.	1,000	3	,999
	F6	,385	3	.	,750	3	,000

a. Lilliefors Significance Correction

ANOVA

Uji Kekuatan Mukoadhesif

	Sum of Squares	df	Mean Square	F	Sig.
Between Groups	12860,527	5	2572,105	274,200	,000
Within Groups	112,565	12	9,380		
Total	12973,092	17			



Multiple Comparisons

Dependent Variable: Uji Kekuatan Mukoadhesif

	(I) Formula	(J) Formula	Mean Difference		Sig.	95% Confidence Interval		
			(I-J)	Std. Error		Lower Bound	Upper Bound	
Tukey HSD	F1	F2	-1,25000	2,50072	,995	-9,6497	7,1497	
		F3	1,00173	2,50072	,998	-7,3980	9,4015	
		F4	-7,99667	2,50072	,065	-16,3964	,4031	
		F5	-37,44667*	2,50072	,000	-45,8464	-29,0469	
		F6	-71,45000*	2,50072	,000	-79,8497	-63,0503	
		F2	F1	1,25000	2,50072	,995	-7,1497	9,6497
	F2	F3	2,25173	2,50072	,939	-6,1480	10,6515	
		F4	-6,74667	2,50072	,147	-15,1464	1,6531	
		F5	-36,19667*	2,50072	,000	-44,5964	-27,7969	
		F6	-70,20000*	2,50072	,000	-78,5997	-61,8003	
		F3	F1	-1,00173	2,50072	,998	-9,4015	7,3980
			F2	-2,25173	2,50072	,939	-10,6515	6,1480
	F4		-8,99840*	2,50072	,033	-17,3981	-,5987	
	F5		-38,44840*	2,50072	,000	-46,8481	-30,0487	
	F6		-72,45173*	2,50072	,000	-80,8515	-64,0520	
	F1		7,99667	2,50072	,065	-,4031	16,3964	
	F5	F2	6,74667	2,50072	,147	-1,6531	15,1464	
		F3	8,99840*	2,50072	,033	,5987	17,3981	
F4		-29,45000*	2,50072	,000	-37,8497	-21,0503		



	F6	-63,45333*	2,50072	,000	-71,8531	-55,0536
F5	F1	37,44667*	2,50072	,000	29,0469	45,8464
	F2	36,19667*	2,50072	,000	27,7969	44,5964
	F3	38,44840*	2,50072	,000	30,0487	46,8481
	F4	29,45000*	2,50072	,000	21,0503	37,8497
	F6	-34,00333*	2,50072	,000	-42,4031	-25,6036
F6	F1	71,45000*	2,50072	,000	63,0503	79,8497
	F2	70,20000*	2,50072	,000	61,8003	78,5997
	F3	72,45173*	2,50072	,000	64,0520	80,8515
	F4	63,45333*	2,50072	,000	55,0536	71,8531
	F5	34,00333*	2,50072	,000	25,6036	42,4031

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



Lampiran 6.4 Analisis Statistik Uji Mukoadhesif dalam Simulasi Pernapasan

	Tests of Normality						
	Formula	Kolmogorov-Smirnov ^a			Shapiro-Wilk		
		Statistic	df	Sig.	Statistic	df	Sig.
Uji Kekuatan	F1	,175	3	.	1,000	3	1,000
Mukoadhesif	F2	,175	3	.	1,000	3	1,000
Stimulasi	F3	,175	3	.	1,000	3	1,000
Pernafasan	F4	,175	3	.	1,000	3	1,000
	F5	,175	3	.	1,000	3	1,000
	F6	,175	3	.	1,000	3	1,000

a. Lilliefors Significance Correction

ANOVA

Uji Kekuatan Mukoadhesif Stimulasi Pernapasan

	Sum of Squares	df	Mean Square	F	Sig.
Between Groups	4185,922	5	837,184	20,568	,000
Within Groups	488,431	12	40,703		
Total	4674,353	17			



Multiple Comparisons

Dependent Variable: Uji Kekuatan Mukoadhesif Stimulasi Pernafasan

	(I) Formula	(J) Formula	Mean Difference		Sig.	95% Confidence Interval		
			(I-J)	Std. Error		Lower Bound	Upper Bound	
Tukey HSD	F1	F2	-,37000	5,20913	1,000	-17,8671	17,1271	
		F3	10,53000	5,20913	,385	-6,9671	28,0271	
		F4	1,44000	5,20913	1,000	-16,0571	18,9371	
		F5	-28,06000*	5,20913	,002	-45,5571	-10,5629	
		F6	-28,96000*	5,20913	,001	-46,4571	-11,4629	
		F2	F1	,37000	5,20913	1,000	-17,1271	17,8671
	F2	F3	10,90000	5,20913	,352	-6,5971	28,3971	
		F4	1,81000	5,20913	,999	-15,6871	19,3071	
		F5	-27,69000*	5,20913	,002	-45,1871	-10,1929	
		F6	-28,59000*	5,20913	,001	-46,0871	-11,0929	
		F3	F1	-10,53000	5,20913	,385	-28,0271	6,9671
			F2	-10,90000	5,20913	,352	-28,3971	6,5971
	F4		-9,09000	5,20913	,531	-26,5871	8,4071	
	F5		-38,59000*	5,20913	,000	-56,0871	-21,0929	
	F6		-39,49000*	5,20913	,000	-56,9871	-21,9929	
	F5		F1	-1,44000	5,20913	1,000	-18,9371	16,0571
		F2	-1,81000	5,20913	,999	-19,3071	15,6871	
		F3	9,09000	5,20913	,531	-8,4071	26,5871	
F4		-29,50000*	5,20913	,001	-46,9971	-12,0029		
F6		-29,50000*	5,20913	,001	-46,9971	-12,0029		



	F6	-30,40000*	5,20913	,001	-47,8971	-12,9029
F5	F1	28,06000*	5,20913	,002	10,5629	45,5571
	F2	27,69000*	5,20913	,002	10,1929	45,1871
	F3	38,59000*	5,20913	,000	21,0929	56,0871
	F4	29,50000*	5,20913	,001	12,0029	46,9971
	F6	-,90000	5,20913	1,000	-18,3971	16,5971
F6	F1	28,96000*	5,20913	,001	11,4629	46,4571
	F2	28,59000*	5,20913	,001	11,0929	46,0871
	F3	39,49000*	5,20913	,000	21,9929	56,9871
	F4	30,40000*	5,20913	,001	12,9029	47,8971
	F5	,90000	5,20913	1,000	-16,5971	18,3971

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



Lampiran 6.5 Analisis Statistik Uji Suhu Gelasi

Tests of Normality							
		Kolmogorov-Smirnov ^a			Shapiro-Wilk		
	Formula	Statistic	df	Sig.	Statistic	df	Sig.
Uji Suhu	F1	,175	3	.	1,000	3	1,000
Gelasi	F2	,175	3	.	1,000	3	1,000
	F3	,211	3	.	,991	3	,814
	F4	,175	3	.	1,000	3	1,000
	F5	,175	3	.	1,000	3	1,000
	F6	,178	3	.	,999	3	,953

a. Lilliefors Significance Correction

ANOVA

Uji Suhu Gelasi

	Sum of Squares	df	Mean Square	F	Sig.
Between Groups	955,199	5	191,040	26,524	,000
Within Groups	86,431	12	7,203		
Total	1041,630	17			



Multiple Comparisons

Dependent Variable: Uji Suhu Gelasi

	(I) Formula	(J) Formula	Mean Difference		Sig.	95% Confidence Interval		
			(I-J)	Std. Error		Lower Bound	Upper Bound	
Tukey HSD	F1	F2	-12,00000*	2,19129	,002	-19,3604	-4,6396	
		F3	-21,13667*	2,19129	,000	-28,4970	-13,7763	
		F4	-12,34000*	2,19129	,001	-19,7004	-4,9796	
		F5	-11,47000*	2,19129	,002	-18,8304	-4,1096	
		F6	-,86000	2,19129	,998	-8,2204	6,5004	
		F2	F1	12,00000*	2,19129	,002	4,6396	19,3604
	F2	F3	-9,13667*	2,19129	,013	-16,4970	-1,7763	
		F4	-,34000	2,19129	1,000	-7,7004	7,0204	
		F5	,53000	2,19129	1,000	-6,8304	7,8904	
		F6	11,14000*	2,19129	,003	3,7796	18,5004	
		F3	F1	21,13667*	2,19129	,000	13,7763	28,4970
			F2	9,13667*	2,19129	,013	1,7763	16,4970
	F4		8,79667*	2,19129	,017	1,4363	16,1570	
	F5		9,66667*	2,19129	,009	2,3063	17,0270	
	F6		20,27667*	2,19129	,000	12,9163	27,6370	
	F5		F1	12,34000*	2,19129	,001	4,9796	19,7004
		F2	,34000	2,19129	1,000	-7,0204	7,7004	
		F3	-8,79667*	2,19129	,017	-16,1570	-1,4363	
F4								
F6		,87000	2,19129	,998	-6,4904	8,2304		



	F6	11,48000*	2,19129	,002	4,1196	18,8404
F5	F1	11,47000*	2,19129	,002	4,1096	18,8304
	F2	-,53000	2,19129	1,000	-7,8904	6,8304
	F3	-9,66667*	2,19129	,009	-17,0270	-2,3063
	F4	-,87000	2,19129	,998	-8,2304	6,4904
	F6	10,61000*	2,19129	,004	3,2496	17,9704
F6	F1	,86000	2,19129	,998	-6,5004	8,2204
	F2	-11,14000*	2,19129	,003	-18,5004	-3,7796
	F3	-20,27667*	2,19129	,000	-27,6370	-12,9163
	F4	-11,48000*	2,19129	,002	-18,8404	-4,1196
	F5	-10,61000*	2,19129	,004	-17,9704	-3,2496

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



Lampiran 6.6 Analisis Statistik Uji Retensi secara *Ex Vivo*

Jam ke-24

Tests of Normality

	Formula	Kolmogorov-Smirnov ^a			Shapiro-Wilk		
		Statistic	df	Sig.	Statistic	df	Sig.
Uji Retensi	NE-TG-AmB	.211	3	.	.991	3	.817
	TG-AmB	.255	3	.	.963	3	.630

a. Lilliefors Significance Correction

Group Statistics

	Formula	N	Mean	Std. Deviation	Std. Error Mean
Uji Retensi	NE-TG-AmB	3	479.22967	84.960626	49.052040
	TG-AmB	3	3.58767	1.686507	.973705

Independent Samples Test

		Levene's Test for Equality of Variances		t-test for Equality of Means						
		F	Sig.	t	df	Sig. (2-tailed)	Mean Difference	Std. Error Difference	95% Confidence Interval of the Difference	
									Lower	Upper
Uji Retensi	Equal variances assumed	5.346	.082	9.695	4	.001	475.64200	49.061703	339.424874	611.859126
	Unequal variances			9.695	2.002	.010	475.64200	49.061703	264.705744	686.578256





Lampiran 7. Hasil Uji Kinetika Model Pelepasan Obat secara *Ex Vivo* menggunakan *add-ins* Microsoft Excel (DDsolver)

NE-TG-AmB (Zero Order)

Time (h)	No.1 F(%)	Mean	SD	RSD(%)
0,5	001	001		
1	001	001		
2	007	007		
3	011	011		
4	021	021		
5	029	029		
6	039	039		
7	057	057		
8	069	069		
12	098	098		
24	147	147		

Best-fit Values

Parameter	No.1	Mean	SD	RSD(%)
k0	0.007	0.007		

Secondary Parameter

Parameter	No.1	Mean	SD	RSD(%)
T25	0.004	0.004		
T50	0.007	0.007		
T75	0.011	0.011		
T80	0.012	0.012		
T90	0.013	0.013		

Goodness of Fit

Parameter	No.1
N_observed	11
DF	10
R_obs-pre	00.001
Rsqr	00.001
Rsqr_adj	00.001
MSE	00.105
MSE_root	00.010
Weighting	1
SS	01.054
WSS	01.054
AIC	00.079
MSC	00.003



TG-AmB (Zero Order)

Time (h)	No.1 F(%)	Mean	SD	RSD(%)
0,5	000	000		
1	001	001		
2	003	003		
3	005	005		
4	008	008		
5	013	013		
6	019	019		
7	027	027		
8	031	031		
12	038	038		
24	061	061		

Best-fit Values

Parameter	No.1	Mean	SD	RSD(%)
k0	0.003	0.003		

Secondary Parameter

Parameter	No.1	Mean	SD	RSD(%)
T25	0.009	0.009		
T50	0.018	0.018		
T75	0.027	0.027		
T80	0.029	0.029		
T90	0.032	0.032		

Goodness of Fit

Parameter	No.1
N_observed	11
DF	10
R_obs-pre	00.001
Rsqr	00.001
Rsqr_adj	00.001
MSE	00.023
MSE_root	00.005
Weighting	1
SS	00.228
WSS	00.228
AIC	00.062
MSC	00.003



Lampiran 8. Dokumentasi



Gambar 12. Penimbangan Pluronic®



Gambar 13. Orientasi Pluronic®



Gambar 14. Uji Kekuatan Mukoadhesif



Gambar 15. Pengukuran Spektrofotometer UV-Vis Hasil Uji Mukoadhesif dalam Simulasi Pernapasan



Optimization Software:
www.balesio.com



Gambar 17. Bubur Jaringan Mukosa Hidung Babi