Individual Noncompartmental Analysis Result

Date and Time: 2017-12-05 09:43:00 NA

Calculation Setting \_\_\_\_\_

Drug Administration: Extravascular

Observation count excluding trailing zero: 11

Dose at time 0: 320 mg

AUC Calculation Method: Linear-up Linear-down

Weighting for lambda z: Uniform (Ordinary Least Square, OLS)

Lambda z selection criterion: Heighest adjusted R-squared value with precision=1e-4

6.8900 6.8912 -1.228e-03 72.7565 322.2988 5.9400 5.9387 +1.324e-03 92.4505 528.5219

3.2800 3.2801 -1.465e-04 148.9231 1459.0711

### Fitting, AUC, AUMC Result \_\_\_\_\_\_

Time	Conc.	Pred.	Residual	AUC	AUMC
0.0000	0.7400			0.0000	0.0000
0.2500	2.8400			0.4475	0.0888
0.5700	6.5700			1.9531	0.8015
1.1200	10.5000			6.6474	5.0654
2.0200	9.6600			15.7194	19.1383
3.8200	8.5800			32.1354	66.1982
5.1000	8.3600			42.9769	114.4617
7.0300	7.4700			58.2529	206.2815

#### Calculated Values

9.0500 \*

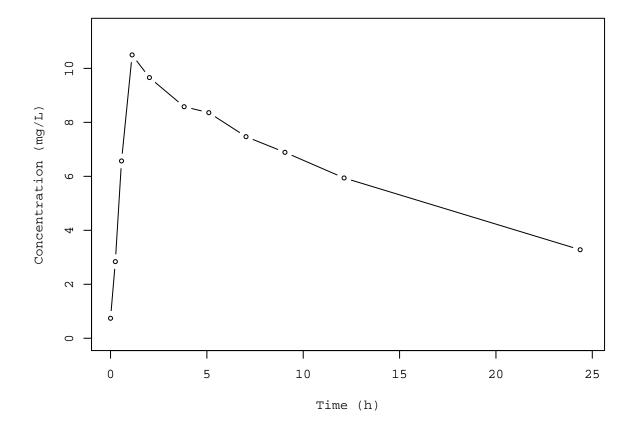
12.1200 \* 24.3700 \*

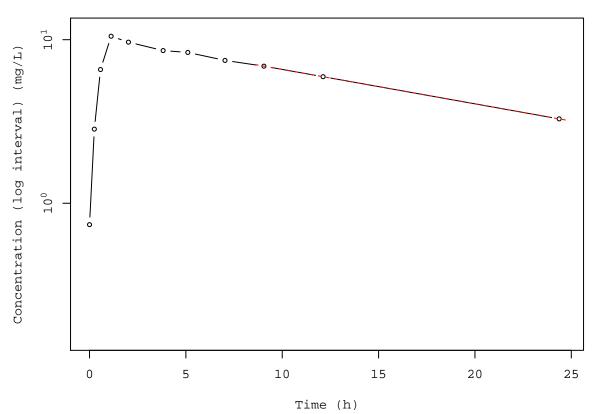
_	_	_	_	_	_	_	_	_	_	_	_	_	_	_	_	_

CMAX	Max Conc	10.5000	mg/L
CMAXD	Max Conc Norm by Dose	0.0328	mg/L/mg
TMAX	Time of CMAX	1.1200	h
TLAG	Time Until First Nonzero Conc	0.0000	h
CLST	Last Nonzero Conc	3.2800	mg/L
CLSTP	Last Nonzero Conc Pred	3.2801	mg/L
TLST	Time of Last Nonzero Conc	24.3700	h
LAMZHL	Half-Life Lambda z	14.3044	h
LAMZ	Lambda z	0.0485	/h
LAMZLL	Lambda z Lower Limit	9.0500	h
LAMZUL	Lambda z Upper Limit	24.3700	h
LAMZNPT	Number of Points for Lambda z	3	
CORRXY	Correlation Between TimeX and Log ConcY	-1.0000	
R2	R Squared	1.0000	
R2ADJ	R Squared Adjusted	1.0000	
AUCLST	AUC to Last Nonzero Conc	148.9231	h*mg/L
AUCALL	AUC All	148.9231	h*mg/L

<sup>\*:</sup> Used for the calculation of Lambda z.

AUCIFO	AUC Infinity Obs	216.6119	h*mg/L
AUCIFOD	AUC Infinity Obs Norm by Dose	0.6769	h*mg/L/mg
AUCIFP	AUC Infinity Pred	216.6150	h*mg/L
AUCIFPD	AUC Infinity Pred Norm by Dose	0.6769	h*mg/L/mg
AUCPEO	AUC %Extrapolation Obs	31.2489	%
AUCPEP	AUC %Extrapolation Pred	31.2499	%
AUMCLST	AUMC to Last Nonzero Conc	1459.0711	h2*mg/L
AUMCIFO	AUMC Infinity Obs	4505.5348	h2*mg/L
AUMCIFP	AUMC Infinity Pred	4505.6709	h2*mg/L
AUMCPEO	AUMC %Extrapolation Obs	67.6160	%
AUMCPEP	AUMC % Extrapolation Pred	67.6170	%
VZFO	Vz Obs by F	30.4867	L
VZFP	Vz Pred by F	30.4863	L
CLFO	Total CL Obs by F	1.4773	L/h
CLFP	Total CL Pred by F	1.4773	L/h
MRTEVLST	MRT Extravasc to Last Nonzero Conc	9.7975	h
MRTEVIFO	MRT Extravasc Infinity Obs	20.8000	h
MRTEVIFP	MRT Extravasc Infinity Pred	20.8004	h





Date and Time: 2017-12-05 09:43:00 NA

Calculation Setting

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Drug Administration: Extravascular

Observation count excluding trailing zero: 11

Dose at time 0: 320 mg

AUC Calculation Method: Linear-up Linear-down

Weighting for lambda z: Uniform (Ordinary Least Square, OLS)

Lambda z selection criterion: Heighest adjusted R-squared value with precision=le-4

0.9000 0.8886 +1.136e-02 91.5268 706.5866

# Fitting, AUC, AUMC Result

Time	Cor	nc. Pred	l. Residua	al AUC	AUMC
0.0000	0.0	 000		0.0000	0.0000
0.2700	1.7	200		0.2322	0.0627
0.5200	7.9	100		1.4360	0.6349
1.0000	8.3	100		5.3287	3.6165
1.9200	8.3	300		12.9832	14.7961
3.5000	6.8	500		24.9754	46.3713
5.0200	6.0	800		34.8022	87.7887
7.0300	* 5.4	000 5.362	29 +3.707e-0	2 46.3396	156.6147
9.0000	* 4.5	500 4.368	37 +1.813e-0	1 56.1403	234.3431
12.0000	* 3.0	100 3.197	70 -1.870e-0	1 67.4803	349.9481

<sup>\*:</sup> Used for the calculation of Lambda z.

# Calculated Values

AUCALL

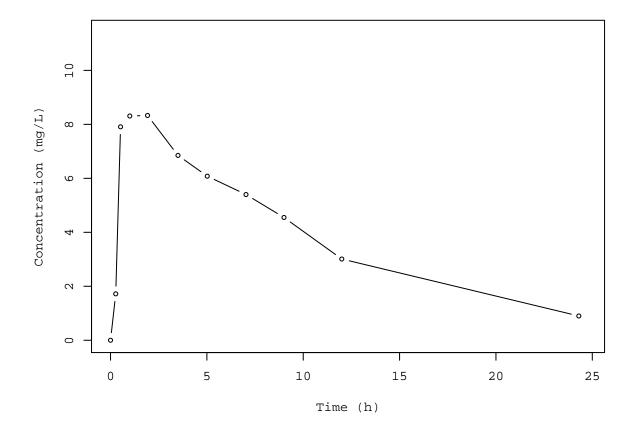
AUC All

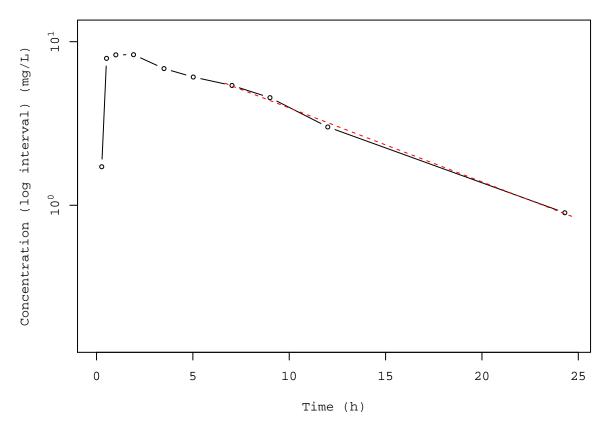
24.3000 \*

CMAX	Max Conc	8.3300	mg/L
CMAXD	Max Conc Norm by Dose	0.0260	mg/L/mg
TMAX	Time of CMAX	1.9200	h
TLAG	Time Until First Nonzero Conc	0.0000	h
CLST	Last Nonzero Conc	0.9000	mg/L
CLSTP	Last Nonzero Conc Pred	0.8886	mg/L
TLST	Time of Last Nonzero Conc	24.3000	h
LAMZHL	Half-Life Lambda z	6.6593	h
LAMZ	Lambda z	0.1041	/h
LAMZLL	Lambda z Lower Limit	7.0300	h
LAMZUL	Lambda z Upper Limit	24.3000	h
LAMZNPT	Number of Points for Lambda z	4	
CORRXY	Correlation Between TimeX and Log ConcY	-0.9986	
R2	R Squared	0.9972	
R2ADJ	R Squared Adjusted	0.9958	
AUCLST	AUC to Last Nonzero Conc	91.5268	h*mg/L

91.5268 h\*mg/L

AUCIFO	AUC Infinity Obs	100.1735	h*mg/L
AUCIFOD	AUC Infinity Obs Norm by Dose	0.3130	h*mg/L/mg
AUCIFP	AUC Infinity Pred	100.0643	h*mg/L
AUCIFPD	AUC Infinity Pred Norm by Dose	0.3127	h*mg/L/mg
AUCPEO	AUC %Extrapolation Obs	8.6317	%
AUCPEP	AUC %Extrapolation Pred	8.5320	%
AUMCLST	AUMC to Last Nonzero Conc	706.5866	h2*mg/L
AUMCIFO	AUMC Infinity Obs	999.7723	h2*mg/L
AUMCIFP	AUMC Infinity Pred	996.0716	h2*mg/L
AUMCPEO	AUMC %Extrapolation Obs	29.3252	%
AUMCPEP	AUMC % Extrapolation Pred	29.0627	%
VZFO	Vz Obs by F	30.6904	L
VZFP	Vz Pred by F	30.7239	L
CLFO	Total CL Obs by F	3.1945	L/h
CLFP	Total CL Pred by F	3.1979	L/h
MRTEVLST	MRT Extravasc to Last Nonzero Conc	7.7200	h
MRTEVIFO	MRT Extravasc Infinity Obs	9.9804	h
MRTEVIFP	MRT Extravasc Infinity Pred	9.9543	h





Date and Time: 2017-12-05 09:43:00 NA

Calculation Setting \_\_\_\_\_

Drug Administration: Extravascular

Observation count excluding trailing zero: 11

Dose at time 0: 320 mg

AUC Calculation Method: Linear-up Linear-down

Weighting for lambda z: Uniform (Ordinary Least Square, OLS)

Lambda z selection criterion: Heighest adjusted R-squared value with precision=1e-4

1.0551 -5.097e-03 99.2865 803.1859

### Fitting, AUC, AUMC Result \_\_\_\_\_

Time	Conc.	Pred.	Residual	AUC	AUMC
0.0000	0.0000			0.0000	0.0000
0.2700	4.4000			0.5940	0.1604
0.5800	6.9000			2.3455	0.9648
1.0200	8.2000			5.6675	3.6854
2.0200	7.8000			13.6675	15.7453
3.6200	7.5000			25.9075	50.0702
5.0800	6.2000			35.9085	92.8817
7.0700	5.3000			47.3510	161.5039
9.0000 *	4.9000	4.9914 -	-9.138e-02	57.1940	240.2199
12.1500 *	3.7000	3.6147 -	+8.528e-02	70.7390	380.4815

<sup>\*:</sup> Used for the calculation of Lambda z.

1.0500

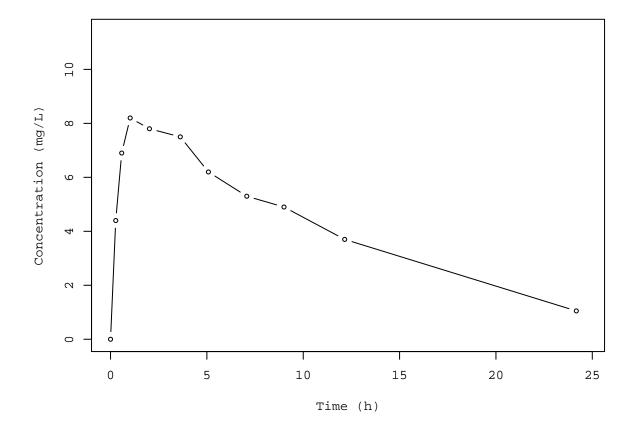
#### Calculated Values

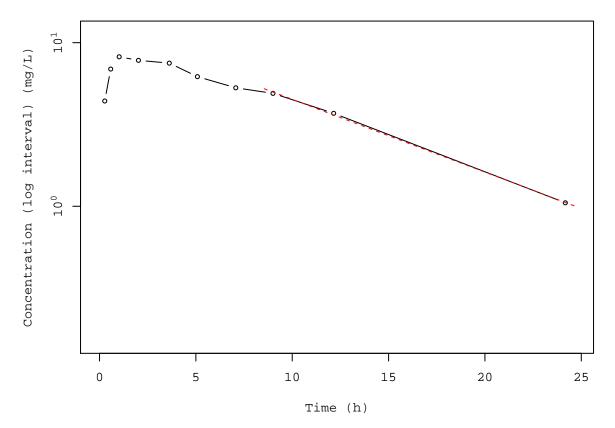
24.1700 \*

	 _	-	-	-	-	-	-	-	-	-	-	-	-	-	
~-	 												_		

CMAX	Max Conc	8.2000	mg/L
CMAXD	Max Conc Norm by Dose	0.0256	mg/L/mg
TMAX	Time of CMAX	1.0200	h
TLAG	Time Until First Nonzero Conc	0.0000	h
CLST	Last Nonzero Conc	1.0500	mg/L
CLSTP	Last Nonzero Conc Pred	1.0551	mg/L
TLST	Time of Last Nonzero Conc	24.1700	h
LAMZHL	Half-Life Lambda z	6.7661	h
LAMZ	Lambda z	0.1024	/h
LAMZLL	Lambda z Lower Limit	9.0000	h
LAMZUL	Lambda z Upper Limit	24.1700	h
LAMZNPT	Number of Points for Lambda z	3	
CORRXY	Correlation Between TimeX and Log ConcY	-0.9997	
R2	R Squared	0.9993	
R2ADJ	R Squared Adjusted	0.9986	
AUCLST	AUC to Last Nonzero Conc	99.2865	h*mg/L
AUCALL	AUC All	99.2865	h*mg/L

AUCIFO	AUC Infinity Obs	109.5360	h*mg/L
AUCIFOD	AUC Infinity Obs Norm by Dose	0.3423	h*mg/L/mg
AUCIFP	AUC Infinity Pred	109.5857	h*mg/L
AUCIFPD	AUC Infinity Pred Norm by Dose	0.3425	h*mg/L/mg
AUCPEO	AUC %Extrapolation Obs	9.3572	%
AUCPEP	AUC %Extrapolation Pred	9.3983	%
AUMCLST	AUMC to Last Nonzero Conc	803.1859	h2*mg/L
AUMCIFO	AUMC Infinity Obs	1150.9648	h2*mg/L
AUMCIFP	AUMC Infinity Pred	1152.6529	h2*mg/L
AUMCPEO	AUMC %Extrapolation Obs	30.2163	%
AUMCPEP	AUMC % Extrapolation Pred	30.3185	%
VZFO	Vz Obs by F	28.5171	L
VZFP	Vz Pred by F	28.5042	L
CLFO	Total CL Obs by F	2.9214	L/h
CLFP	Total CL Pred by F	2.9201	L/h
MRTEVLST	MRT Extravasc to Last Nonzero Conc	8.0896	h
MRTEVIFO	MRT Extravasc Infinity Obs	10.5076	h
MRTEVIFP	MRT Extravasc Infinity Pred	10.5183	h





Date and Time: 2017-12-05 09:43:00 NA

Calculation Setting \_\_\_\_\_

Drug Administration: Extravascular

Observation count excluding trailing zero: 11

Dose at time 0: 320 mg

AUC Calculation Method: Linear-up Linear-down

Weighting for lambda z: Uniform (Ordinary Least Square, OLS)

Lambda z selection criterion: Heighest adjusted R-squared value with precision=1e-4

1.1564 -6.422e-03 106.7963 901.0842

### Fitting, AUC, AUMC Result \_\_\_\_\_

	Time	Conc.	Pred.	Residual	AUC	AUMC
_	0.0000	0.0000			0.0000	0.0000
	0.3500	1.8900			0.3308	0.1158
	0.6000	4.6000			1.1420	0.5435
	1.0700	8.6000			4.2440	3.3545
	2.1300	8.3800			13.2434	17.6918
	3.5000	7.5400			24.1486	47.9958
	5.0200	6.8800			35.1078	94.3007
	7.0200	5.7800			47.7678	169.4139
	9.0200 *	5.3300	5.4586	-1.286e-01	58.8778	258.0661
	11.9800 *	4.1900	4.0686	+1.214e-01	72.9674	403.5099

<sup>\*:</sup> Used for the calculation of Lambda z.

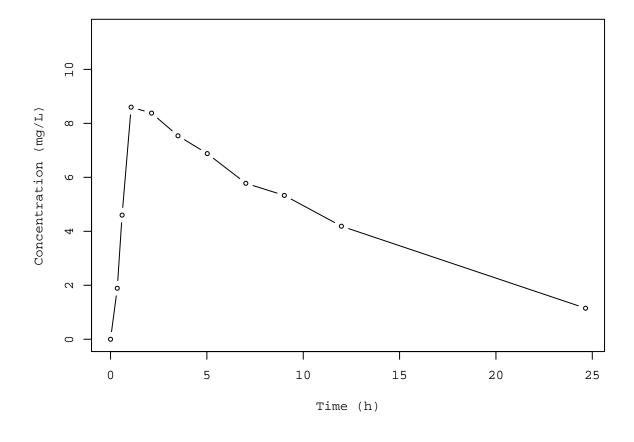
1.1500

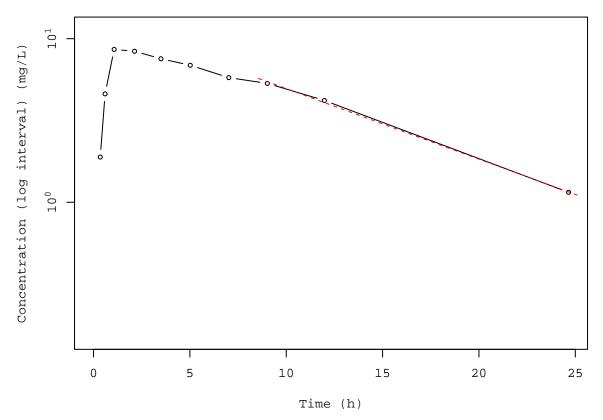
#### Calculated Values \_\_\_\_\_

24.6500 \*

CMAX	Max Conc	8.6000	mg/L
CMAXD	Max Conc Norm by Dose	0.0269	mg/L/mg
TMAX	Time of CMAX	1.0700	h
TLAG	Time Until First Nonzero Conc	0.0000	h
CLST	Last Nonzero Conc	1.1500	mg/L
CLSTP	Last Nonzero Conc Pred	1.1564	mg/L
TLST	Time of Last Nonzero Conc	24.6500	h
LAMZHL	Half-Life Lambda z	6.9812	h
LAMZ	Lambda z	0.0993	/h
LAMZLL	Lambda z Lower Limit	9.0200	h
LAMZUL	Lambda z Upper Limit	24.6500	h
LAMZNPT	Number of Points for Lambda z	3	
CORRXY	Correlation Between TimeX and Log ConcY	-0.9995	
R2	R Squared	0.9989	
R2ADJ	R Squared Adjusted	0.9978	
AUCLST	AUC to Last Nonzero Conc	106.7963	h*mg/L
AUCALL	AUC All	106.7963	h*mg/L

AUCIFO	AUC Infinity Obs	118.3789	h*mg/L
AUCIFOD	AUC Infinity Obs Norm by Dose	0.3699	h*mg/L/mg
AUCIFP	AUC Infinity Pred	118.4436	h*mg/L
AUCIFPD	AUC Infinity Pred Norm by Dose	0.3701	h*mg/L/mg
AUCPEO	AUC %Extrapolation Obs	9.7843	%
AUCPEP	AUC %Extrapolation Pred	9.8336	%
AUMCLST	AUMC to Last Nonzero Conc	901.0842	h2*mg/L
AUMCIFO	AUMC Infinity Obs	1303.2524	h2*mg/L
AUMCIFP	AUMC Infinity Pred	1305.4981	h2*mg/L
AUMCPEO	AUMC %Extrapolation Obs	30.8588	%
AUMCPEP	AUMC % Extrapolation Pred	30.9777	%
VZFO	Vz Obs by F	27.2260	L
VZFP	Vz Pred by F	27.2111	L
CLFO	Total CL Obs by F	2.7032	L/h
CLFP	Total CL Pred by F	2.7017	L/h
MRTEVLST	MRT Extravasc to Last Nonzero Conc	8.4374	h
MRTEVIFO	MRT Extravasc Infinity Obs	11.0092	h
MRTEVIFP	MRT Extravasc Infinity Pred	11.0221	h





Date and Time: 2017-12-05 09:43:00 NA

Calculation Setting

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Drug Administration: Extravascular

Observation count excluding trailing zero: 11

Dose at time 0: 320 mg

AUC Calculation Method: Linear-up Linear-down

Weighting for lambda z: Uniform (Ordinary Least Square, OLS)

Lambda z selection criterion: Heighest adjusted R-squared value with precision=le-4

#### Fitting, AUC, AUMC Result

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Time		Conc.	Pred.	Residual	AUC	AUMC
0.0000		0.0000			0.0000	0.0000
0.3000		2.0200			0.3030	0.0909
0.5200		5.6300			1.1445	0.4796
1.0000		11.4000			5.2317	3.9182
2.0200		9.3300			15.8040	19.3440
3.5000		8.7400			29.1758	55.9271
5.0200		7.5600			41.5638	108.0184
7.0200	*	7.0900	6.9799	+1.101e-01	56.2138	195.7414
9.1000	*	5.9000	5.8291	+7.091e-02	69.7234	303.3417
12.0000	*	4.3700	4.5343	-1.643e-01	84.6149	457.2302
24.3500	*	1.5700	1.5557	+1.430e-02	121.2944	1017.1143

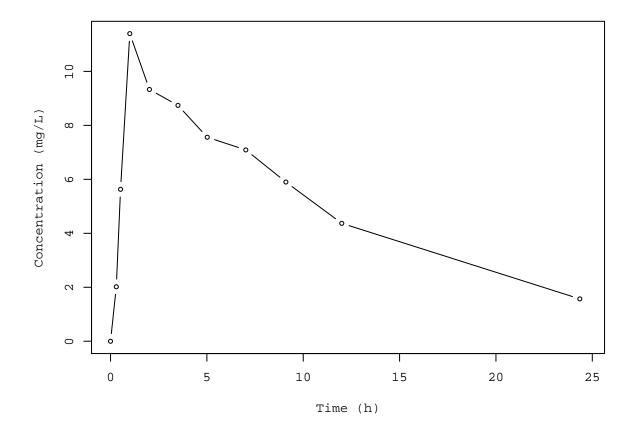
<sup>\*:</sup> Used for the calculation of Lambda z.

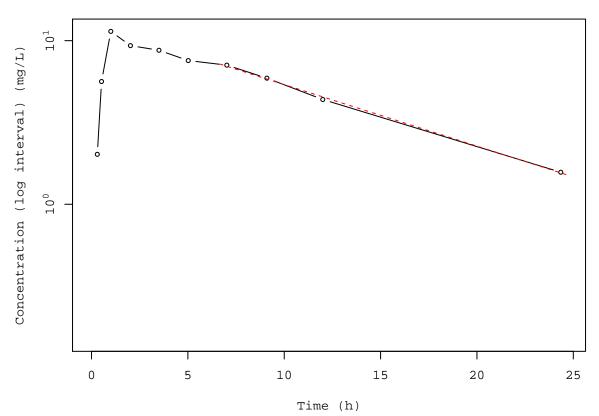
#### Calculated Values

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Max Conc	11.4000	mg/L
Max Conc Norm by Dose	0.0356	mg/L/mg
Time of CMAX	1.0000	h
Time Until First Nonzero Conc	0.0000	h
Last Nonzero Conc	1.5700	mg/L
Last Nonzero Conc Pred	1.5557	mg/L
Time of Last Nonzero Conc	24.3500	h
Half-Life Lambda z	8.0023	h
Lambda z	0.0866	/h
Lambda z Lower Limit	7.0200	h
Lambda z Upper Limit	24.3500	h
Number of Points for Lambda z	4	
Correlation Between TimeX and Log ConcY	-0.9993	
R Squared	0.9986	
R Squared Adjusted	0.9980	
AUC to Last Nonzero Conc	121.2944	h*mg/L
AUC All	121.2944	h*mg/L
	Max Conc Norm by Dose Time of CMAX Time Until First Nonzero Conc Last Nonzero Conc Last Nonzero Conc Pred Time of Last Nonzero Conc Half-Life Lambda z Lambda z Lambda z Lambda z Lower Limit Lambda z Upper Limit Number of Points for Lambda z Correlation Between TimeX and Log ConcY R Squared R Squared Adjusted AUC to Last Nonzero Conc	Max Conc Norm by Dose 0.0356 Time of CMAX 1.0000 Time Until First Nonzero Conc 0.0000 Last Nonzero Conc 1.5700 Last Nonzero Conc Pred 1.5557 Time of Last Nonzero Conc 24.3500 Half-Life Lambda z 8.0023 Lambda z 0.0866 Lambda z Lower Limit 7.0200 Lambda z Upper Limit 24.3500 Number of Points for Lambda z 4 Correlation Between TimeX and Log ConcY -0.9993 R Squared 0.9986 R Squared Adjusted 0.9980 AUC to Last Nonzero Conc 121.2944

AUCIFO	AUC Infinity Obs	139.4198	h*mg/L
AUCIFOD	AUC Infinity Obs Norm by Dose	0.4357	h*mg/L/mg
AUCIFP	AUC Infinity Pred	139.2546	h*mg/L
AUCIFPD	AUC Infinity Pred Norm by Dose	0.4352	h*mg/L/mg
AUCPEO	AUC %Extrapolation Obs	13.0006	8
AUCPEP	AUC %Extrapolation Pred	12.8974	8
AUMCLST	AUMC to Last Nonzero Conc	1017.1143	h2*mg/L
AUMCIFO	AUMC Infinity Obs	1667.7216	h2*mg/L
AUMCIFP	AUMC Infinity Pred	1661.7937	h2*mg/L
AUMCPEO	AUMC %Extrapolation Obs	39.0117	%
AUMCPEP	AUMC % Extrapolation Pred	38.7942	%
VZFO	Vz Obs by F	26.4980	L
VZFP	Vz Pred by F	26.5294	L
CLFO	Total CL Obs by F	2.2952	L/h
CLFP	Total CL Pred by F	2.2979	L/h
MRTEVLST	MRT Extravasc to Last Nonzero Conc	8.3855	h
MRTEVIFO	MRT Extravasc Infinity Obs	11.9619	h
MRTEVIFP	MRT Extravasc Infinity Pred	11.9335	h





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Calculation Setting \_\_\_\_\_

Drug Administration: Extravascular

Observation count excluding trailing zero: 11

Dose at time 0: 320 mg

AUC Calculation Method: Linear-up Linear-down

Weighting for lambda z: Uniform (Ordinary Least Square, OLS)

Lambda z selection criterion: Heighest adjusted R-squared value with precision=le-4

### Fitting, AUC, AUMC Result

Time	Conc.

Time		Conc.	Pred.	Residual	AUC	AUMC
0.0000		0.0000			0.0000	0.0000
0.0000		0.0000			0.0000	0.0000
0.2700		1.2900			0.1742	0.0470
0.5800		3.0800			0.8515	0.3779
1.1500		6.4400			3.5647	2.9977
2.0300	*	6.3200	6.3928	-7.284e-02	9.1791	11.9014
3.5700	*	5.5300	5.5844	-5.438e-02	18.3036	36.9816
5.0000	*	4.9400	4.9255	+1.450e-02	25.7897	68.7577
7.0000	*	4.0200	4.1323	-1.123e-01	34.7497	121.5977
9.2200	*	3.4600	3.4005	+5.948e-02	43.0525	188.2434
12.1000	*	2.7800	2.6408	+1.392e-01	52.0381	282.6199
23.8500	*	0.9200	0.9413	-2.127e-02	73.7756	609.1524

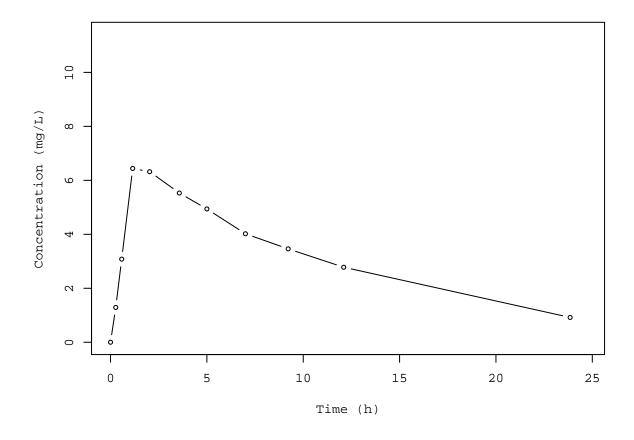
<sup>\*:</sup> Used for the calculation of Lambda z.

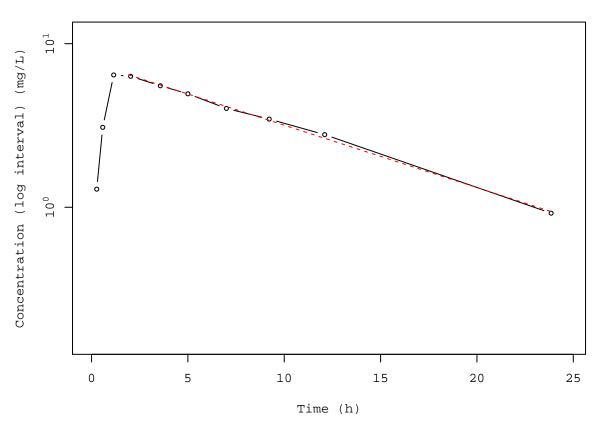
#### Calculated Values

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CMAX	Max Conc	6.4400	mg/L
CMAXD	Max Conc Norm by Dose	0.0201	mg/L/mg
TMAX	Time of CMAX	1.1500	h
TLAG	Time Until First Nonzero Conc	0.0000	h
CLST	Last Nonzero Conc	0.9200	mg/L
CLSTP	Last Nonzero Conc Pred	0.9413	mg/L
TLST	Time of Last Nonzero Conc	23.8500	h
LAMZHL	Half-Life Lambda z	7.8950	h
LAMZ	Lambda z	0.0878	/h
LAMZLL	Lambda z Lower Limit	2.0300	h
LAMZUL	Lambda z Upper Limit	23.8500	h
LAMZNPT	Number of Points for Lambda z	7	
CORRXY	Correlation Between TimeX and Log ConcY	-0.9991	
R2	R Squared	0.9982	
R2ADJ	R Squared Adjusted	0.9979	
AUCLST	AUC to Last Nonzero Conc	73.7756	h*mg/L
AUCALL	AUC All	73.7756	h*mg/L

AUCIFO	AUC Infinity Obs	84.2544	h*mg/L
AUCIFOD	AUC Infinity Obs Norm by Dose	0.2633	h*mg/L/mg
AUCIFP	AUC Infinity Pred	84.4967	h*mg/L
AUCIFPD	AUC Infinity Pred Norm by Dose	0.2641	h*mg/L/mg
AUCPEO	AUC %Extrapolation Obs	12.4372	%
AUCPEP	AUC %Extrapolation Pred	12.6882	%
AUMCLST	AUMC to Last Nonzero Conc	609.1524	h2*mg/L
AUMCIFO	AUMC Infinity Obs	978.4285	h2*mg/L
AUMCIFP	AUMC Infinity Pred	986.9665	h2*mg/L
AUMCPEO	AUMC %Extrapolation Obs	37.7418	%
AUMCPEP	AUMC % Extrapolation Pred	38.2803	%
VZFO	Vz Obs by F	43.2597	L
VZFP	Vz Pred by F	43.1357	L
CLFO	Total CL Obs by F	3.7980	L/h
CLFP	Total CL Pred by F	3.7871	L/h
MRTEVLST	MRT Extravasc to Last Nonzero Conc	8.2568	h
MRTEVIFO	MRT Extravasc Infinity Obs	11.6128	h
MRTEVIFP	MRT Extravasc Infinity Pred	11.6805	h





Date and Time: 2017-12-05 09:43:00 NA

Calculation Setting

Drug Administration: Extravascular

Observation count excluding trailing zero: 11

Dose at time 0: 320 mg

AUC Calculation Method: Linear-up Linear-down

Weighting for lambda z: Uniform (Ordinary Least Square, OLS)

Lambda z selection criterion: Heighest adjusted R-squared value with precision=1e-4

1.1500 1.1607 -1.072e-02 90.7534 782.4199

# Fitting, AUC, AUMC Result

Time		Conc.	Pred.	Residual	AUC	AUMC
0.0000		0.1500			0.0000	0.0000
0.2500		0.8500			0.1250	0.0266
0.5000		2.3500			0.5250	0.2000
1.0200		5.0200			2.4412	1.8368
2.0200		6.5800			8.2412	11.0428
3.4800		7.0900			18.2203	38.7571
5.0000		6.6600			28.6703	82.8167
6.9800	*	5.2500	5.3226	-7.260e-02	40.4612	152.0623
9.0000	*	4.3900	4.4527	-6.275e-02	50.1976	228.9788
12.0500	*	3.5300	3.4011	+1.289e-01	62.2756	354.0998

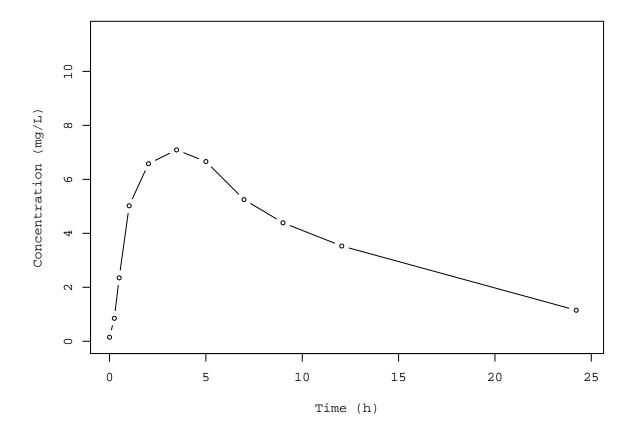
<sup>\*:</sup> Used for the calculation of Lambda z.

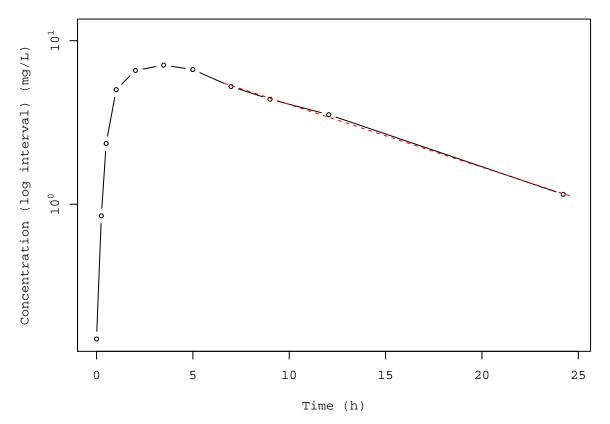
# Calculated Values

24.2200 \*

CMAX	Max Conc	7.0900	mg/L
CMAXD	Max Conc Norm by Dose	0.0222	mg/L/mg
TMAX	Time of CMAX	3.4800	h
TLAG	Time Until First Nonzero Conc	0.0000	h
CLST	Last Nonzero Conc	1.1500	mg/L
CLSTP	Last Nonzero Conc Pred	1.1607	mg/L
TLST	Time of Last Nonzero Conc	24.2200	h
LAMZHL	Half-Life Lambda z	7.8467	h
LAMZ	Lambda z	0.0883	/h
LAMZLL	Lambda z Lower Limit	6.9800	h
LAMZUL	Lambda z Upper Limit	24.2200	h
LAMZNPT	Number of Points for Lambda z	4	
CORRXY	Correlation Between TimeX and Log ConcY	-0.9993	
R2	R Squared	0.9987	
R2ADJ	R Squared Adjusted	0.9980	
AUCLST	AUC to Last Nonzero Conc	90.7534	h*mg/L
AUCALL	AUC All	90.7534	h*mg/L

AUCIFO	AUC Infinity Obs	103.7718	h*mg/L
AUCIFOD	AUC Infinity Obs Norm by Dose	0.3243	h*mg/L/mg
AUCIFP	AUC Infinity Pred	103.8931	h*mg/L
AUCIFPD	AUC Infinity Pred Norm by Dose	0.3247	h*mg/L/mg
AUCPEO	AUC %Extrapolation Obs	12.5452	8
AUCPEP	AUC %Extrapolation Pred	12.6474	8
AUMCLST	AUMC to Last Nonzero Conc	782.4199	h2*mg/L
AUMCIFO	AUMC Infinity Obs	1245.0984	h2*mg/L
AUMCIFP	AUMC Infinity Pred	1249.4111	h2*mg/L
AUMCPEO	AUMC %Extrapolation Obs	37.1600	%
AUMCPEP	AUMC % Extrapolation Pred	37.3769	%
VZFO	Vz Obs by F	34.9084	L
VZFP	Vz Pred by F	34.8677	L
CLFO	Total CL Obs by F	3.0837	L/h
CLFP	Total CL Pred by F	3.0801	L/h
MRTEVLST	MRT Extravasc to Last Nonzero Conc	8.6214	h
MRTEVIFO	MRT Extravasc Infinity Obs	11.9984	h
MRTEVIFP	MRT Extravasc Infinity Pred	12.0259	h





Date and Time: 2017-12-05 09:43:00 NA

Calculation Setting

Drug Administration: Extravascular

Observation count excluding trailing zero: 11

Dose at time 0: 320 mg

AUC Calculation Method: Linear-up Linear-down

Weighting for lambda z: Uniform (Ordinary Least Square, OLS)

Lambda z selection criterion: Heighest adjusted R-squared value with precision=le-4

1.2500 1.2285 +2.147e-02 88.5600 739.5346

# Fitting, AUC, AUMC Result

Time		Conc.	Pred.	Residual	AUC	AUMC
2 2222		0 0000			0.0000	0.0000
0.0000		0.0000			0.0000	0.0000
0.2500		3.0500			0.3813	0.0953
0.5200		3.0500			1.2048	0.4124
0.9800		7.3100			3.5875	2.4248
2.0200		7.5600			11.3200	14.0910
3.5300	*	6.5900	6.5724	+1.758e-02	22.0032	43.1841
5.0500	*	5.8800	5.8071	+7.292e-02	31.4804	83.4312
7.1500	*	4.7300	4.8941	-1.641e-01	42.6209	150.1204
9.0700	*	4.5700	4.1856	+3.844e-01	51.5489	222.3790
12.1000	*	3.0000	3.2702	-2.702e-01	63.0175	340.1701

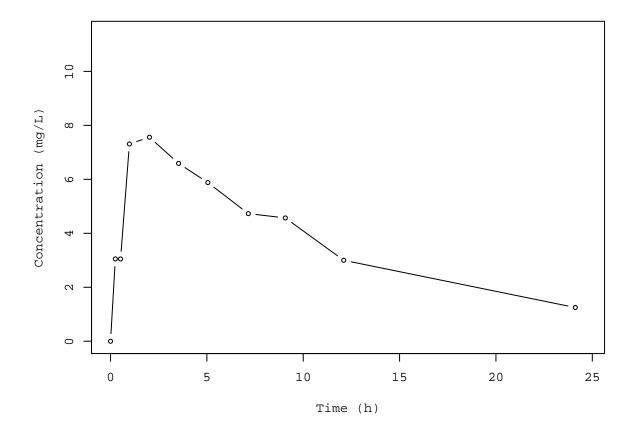
<sup>\*:</sup> Used for the calculation of Lambda z.

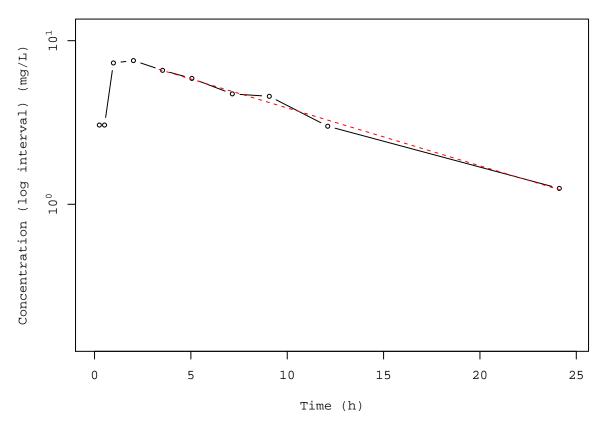
# Calculated Values

24.1200 \*

CMAX	Max Conc	7.5600	mg/L
CMAXD	Max Conc Norm by Dose	0.0236	mg/L/mg
TMAX	Time of CMAX	2.0200	h
TLAG	Time Until First Nonzero Conc	0.0000	h
CLST	Last Nonzero Conc	1.2500	mg/L
CLSTP	Last Nonzero Conc Pred	1.2285	mg/L
TLST	Time of Last Nonzero Conc	24.1200	h
LAMZHL	Half-Life Lambda z	8.5100	h
LAMZ	Lambda z	0.0815	/h
LAMZLL	Lambda z Lower Limit	3.5300	h
LAMZUL	Lambda z Upper Limit	24.1200	h
LAMZNPT	Number of Points for Lambda z	6	
CORRXY	Correlation Between TimeX and Log ConcY	-0.9955	
R2	R Squared	0.9910	
R2ADJ	R Squared Adjusted	0.9888	
AUCLST	AUC to Last Nonzero Conc	88.5600	h*mg/L
AUCALL	AUC All	88.5600	h*mg/L

AUCIFO	AUC Infinity Obs	103.9067	h*mg/L
AUCIFOD	AUC Infinity Obs Norm by Dose	0.3247	h*mg/L/mg
AUCIFP	AUC Infinity Pred	103.6431	h*mg/L
AUCIFPD	AUC Infinity Pred Norm by Dose	0.3239	h*mg/L/mg
AUCPEO	AUC %Extrapolation Obs	14.7697	8
AUCPEP	AUC %Extrapolation Pred	14.5529	8
AUMCLST	AUMC to Last Nonzero Conc	739.5346	h2*mg/L
AUMCIFO	AUMC Infinity Obs	1298.1158	h2*mg/L
AUMCIFP	AUMC Infinity Pred	1288.5201	h2*mg/L
AUMCPEO	AUMC %Extrapolation Obs	43.0302	%
AUMCPEP	AUMC % Extrapolation Pred	42.6059	%
VZFO	Vz Obs by F	37.8105	L
VZFP	Vz Pred by F	37.9067	L
CLFO	Total CL Obs by F	3.0797	L/h
CLFP	Total CL Pred by F	3.0875	L/h
MRTEVLST	MRT Extravasc to Last Nonzero Conc	8.3507	h
MRTEVIFO	MRT Extravasc Infinity Obs	12.4931	h
MRTEVIFP	MRT Extravasc Infinity Pred	12.4323	h





Date and Time: 2017-12-05 09:43:00 NA

Calculation Setting \_\_\_\_\_

Drug Administration: Extravascular

Observation count excluding trailing zero: 11

Dose at time 0: 320 mg

AUC Calculation Method: Linear-up Linear-down

Weighting for lambda z: Uniform (Ordinary Least Square, OLS)

Lambda z selection criterion: Heighest adjusted R-squared value with precision=1e-4

1.1165 +3.517e-03 86.3262 705.2296

### Fitting, AUC, AUMC Result \_\_\_\_\_\_

Time	Conc.	Pred.	Residual	AUC	AUMC
 0.0000	0.0000			0.0000	0.0000
0.3000	7.3700			1.1055	0.3316
0.6300	9.0300			3.8115	1.6351
1.0500	7.1400			7.2072	4.4042
2.0200	6.3300			13.7402	14.2417
3.5300	5.6600			22.7926	38.9804
5.0200	5.6700			31.2335	75.0705
7.1700	4.2400			41.8867	138.3495
8.8000 *	4.1100	4.0512	+5.880e-02	48.6920	192.6031
11.6000 *	3.1600	3.2160	-5.597e-02	58.8700	294.5567

<sup>\*:</sup> Used for the calculation of Lambda z.

1.1200

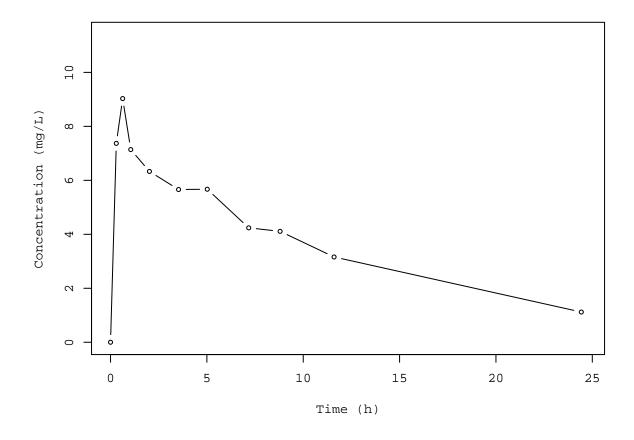
#### Calculated Values \_\_\_\_\_

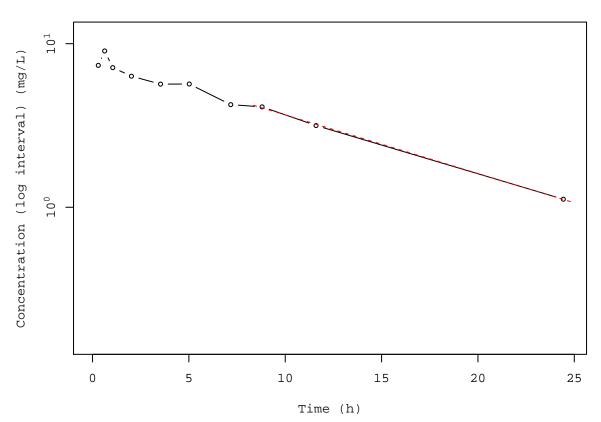
24.4300 \*

CMAX	Max	Con

CMAX	Max Conc	9.0300	mg/L
CMAXD	Max Conc Norm by Dose	0.0282	${\rm mg/L/mg}$
TMAX	Time of CMAX	0.6300	h
TLAG	Time Until First Nonzero Conc	0.0000	h
CLST	Last Nonzero Conc	1.1200	mg/L
CLSTP	Last Nonzero Conc Pred	1.1165	mg/L
TLST	Time of Last Nonzero Conc	24.4300	h
LAMZHL	Half-Life Lambda z	8.4060	h
LAMZ	Lambda z	0.0825	/h
LAMZLL	Lambda z Lower Limit	8.8000	h
LAMZUL	Lambda z Upper Limit	24.4300	h
LAMZNPT	Number of Points for Lambda z	3	
CORRXY	Correlation Between TimeX and Log ConcY	-0.9997	
R2	R Squared	0.9994	
R2ADJ	R Squared Adjusted	0.9989	
AUCLST	AUC to Last Nonzero Conc	86.3262	h*mg/L
AUCALL	AUC All	86.3262	h*mg/L

AUCIFO	AUC Infinity Obs	99.9087	h*mg/L
AUCIFOD	AUC Infinity Obs Norm by Dose	0.3122	h*mg/L/mg
AUCIFP	AUC Infinity Pred	99.8661	h*mg/L
AUCIFPD	AUC Infinity Pred Norm by Dose	0.3121	h*mg/L/mg
AUCPEO	AUC %Extrapolation Obs	13.5950	8
AUCPEP	AUC %Extrapolation Pred	13.5581	8
AUMCLST	AUMC to Last Nonzero Conc	705.2296	h2*mg/L
AUMCIFO	AUMC Infinity Obs	1201.7715	h2*mg/L
AUMCIFP	AUMC Infinity Pred	1200.2124	h2*mg/L
AUMCPEO	AUMC %Extrapolation Obs	41.3175	%
AUMCPEP	AUMC % Extrapolation Pred	41.2413	8
VZFO	Vz Obs by F	38.8428	L
VZFP	Vz Pred by F	38.8594	L
CLFO	Total CL Obs by F	3.2029	L/h
CLFP	Total CL Pred by F	3.2043	L/h
MRTEVLST	MRT Extravasc to Last Nonzero Conc	8.1694	h
MRTEVIFO	MRT Extravasc Infinity Obs	12.0287	h
MRTEVIFP	MRT Extravasc Infinity Pred	12.0182	h





Date and Time: 2017-12-05 09:43:00 NA

Calculation Setting \_\_\_\_\_

Drug Administration: Extravascular

Observation count excluding trailing zero: 11

Dose at time 0: 320 mg

AUC Calculation Method: Linear-up Linear-down

Weighting for lambda z: Uniform (Ordinary Least Square, OLS)

Lambda z selection criterion: Heighest adjusted R-squared value with precision=1e-4

2.4200 2.4137 +6.308e-03 138.3681 1278.1800

91.3881 546.9044

### Fitting, AUC, AUMC Result \_\_\_\_\_\_

Time	Conc.	Pred.	Residual	AUC	AUMC
0.0000	0.2400			0.0000	0.0000
0.3700	2.8900			0.5790	0.1978
0.7700	5.2200			2.2011	1.2156
1.0200	6.4100			3.6548	2.5353
2.0500	7.8300			10.9884	14.1690
3.5500	10.2100			24.5184	53.3917
5.0500	9.1800			39.0609	115.3451
7.0800	8.0200			56.5189	220.0328
9.3800	* 7.1400	7.0610	+7.903e-02	73.9529	362.3508

5.6800 5.7586 -7.858e-02

#### Calculated Values

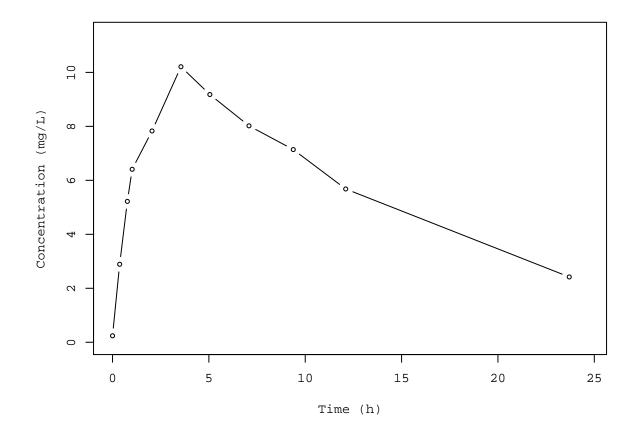
12.1000 \* 23.7000 \*

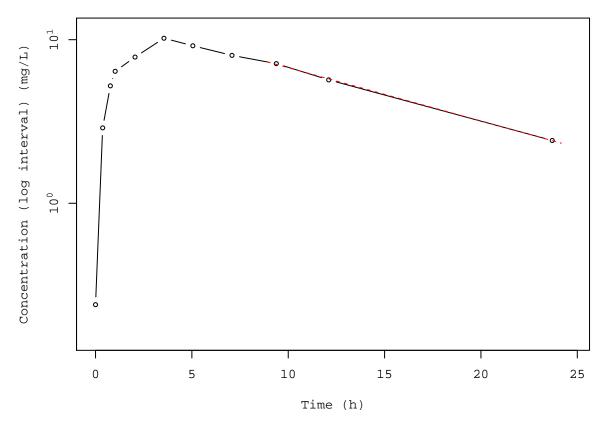
_	_	_	_	_	_	_	_	_	_	_	_	_	_	_	_	_	

CMAX	Max Conc	10.2100	mg/L
CMAXD	Max Conc Norm by Dose	0.0319	mg/L/mg
TMAX	Time of CMAX	3.5500	h
TLAG	Time Until First Nonzero Conc	0.0000	h
CLST	Last Nonzero Conc	2.4200	mg/L
CLSTP	Last Nonzero Conc Pred	2.4137	mg/L
TLST	Time of Last Nonzero Conc	23.7000	h
LAMZHL	Half-Life Lambda z	9.2469	h
LAMZ	Lambda z	0.0750	/h
LAMZLL	Lambda z Lower Limit	9.3800	h
LAMZUL	Lambda z Upper Limit	23.7000	h
LAMZNPT	Number of Points for Lambda z	3	
CORRXY	Correlation Between TimeX and Log ConcY	-0.9998	
R2	R Squared	0.9995	
R2ADJ	R Squared Adjusted	0.9990	
AUCLST	AUC to Last Nonzero Conc	138.3681	h*mg/L
AUCALL	AUC All	138.3681	h*mg/L

<sup>\*:</sup> Used for the calculation of Lambda z.

AUCIFO	AUC Infinity Obs	170.6521	h*mg/L
AUCIFOD	AUC Infinity Obs Norm by Dose	0.5333	h*mg/L/mg
AUCIFP	AUC Infinity Pred	170.5679	h*mg/L
AUCIFPD	AUC Infinity Pred Norm by Dose	0.5330	h*mg/L/mg
AUCPEO	AUC %Extrapolation Obs	18.9180	8
AUCPEP	AUC %Extrapolation Pred	18.8780	8
AUMCLST	AUMC to Last Nonzero Conc	1278.1800	h2*mg/L
AUMCIFO	AUMC Infinity Obs	2473.9934	h2*mg/L
AUMCIFP	AUMC Infinity Pred	2470.8765	h2*mg/L
AUMCPEO	AUMC %Extrapolation Obs	48.3354	%
AUMCPEP	AUMC % Extrapolation Pred	48.2702	%
VZFO	Vz Obs by F	25.0155	L
VZFP	Vz Pred by F	25.0279	L
CLFO	Total CL Obs by F	1.8752	L/h
CLFP	Total CL Pred by F	1.8761	L/h
MRTEVLST	MRT Extravasc to Last Nonzero Conc	9.2375	h
MRTEVIFO	MRT Extravasc Infinity Obs	14.4973	h
MRTEVIFP	MRT Extravasc Infinity Pred	14.4862	h





Date and Time: 2017-12-05 09:43:00 NA

Calculation Setting

Drug Administration: Extravascular

Observation count excluding trailing zero: 11

Dose at time 0: 320 mg

AUC Calculation Method: Linear-up Linear-down

Weighting for lambda z: Uniform (Ordinary Least Square, OLS)

Lambda z selection criterion: Heighest adjusted R-squared value with precision=le-4

# Fitting, AUC, AUMC Result

Time	Conc.	Pred.	Residual	AUC	AUMC
0.0000	0.0000			0.0000	0.0000
0.2500	4.8600			0.6075	0.1519
0.5000	7.2400			2.1200	0.7563
0.9800	8.0000			5.7776	3.5067
1.9800	6.8100			13.1826	14.1686
3.6000	5.8700			23.4534	42.2073
5.0200	5.2200			31.3273	75.8162
7.0300	4.4500			41.0457	133.5915
9.0300 *	3.6200	3.6169	+3.150e-03	49.1156	197.5636

2.6929 -2.948e-03

0.8600 0.8598 +1.934e-04 80.0936 617.2422

58.8646 298.4388

2.6900

# Calculated Values

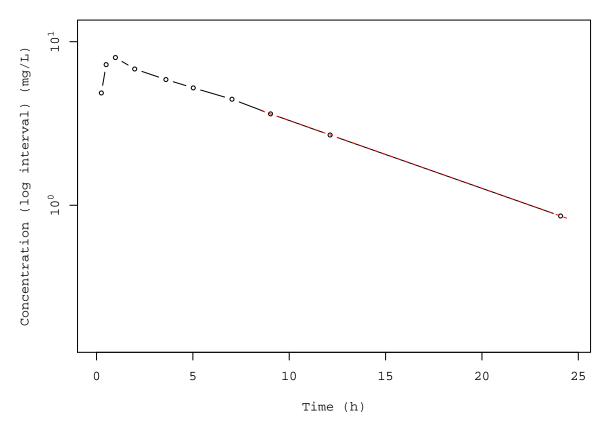
12.1200 \* 24.0800 \*

CMAX	Max Conc	8.0000	mg/L
CMAXD	Max Conc Norm by Dose	0.0250	${\rm mg/L/mg}$
TMAX	Time of CMAX	0.9800	h
TLAG	Time Until First Nonzero Conc	0.0000	h
CLST	Last Nonzero Conc	0.8600	mg/L
CLSTP	Last Nonzero Conc Pred	0.8598	mg/L
TLST	Time of Last Nonzero Conc	24.0800	h
LAMZHL	Half-Life Lambda z	7.2612	h
LAMZ	Lambda z	0.0955	/h
LAMZLL	Lambda z Lower Limit	9.0300	h
LAMZUL	Lambda z Upper Limit	24.0800	h
LAMZNPT	Number of Points for Lambda z	3	
CORRXY	Correlation Between TimeX and Log ConcY	-1.0000	
R2	R Squared	1.0000	
R2ADJ	R Squared Adjusted	1.0000	
AUCLST	AUC to Last Nonzero Conc	80.0936	h*mg/L
AUCALL	AUC All	80.0936	h*mg/L

 $<sup>\</sup>ast\colon$  Used for the calculation of Lambda z.

AUCIFO	AUC Infinity Obs	89.1027	h*mg/L
AUCIFOD	AUC Infinity Obs Norm by Dose	0.2784	h*mg/L/mg
AUCIFP	AUC Infinity Pred	89.1007	h*mg/L
AUCIFPD	AUC Infinity Pred Norm by Dose	0.2784	h*mg/L/mg
AUCPEO	AUC %Extrapolation Obs	10.1110	%
AUCPEP	AUC %Extrapolation Pred	10.1089	%
AUMCLST	AUMC to Last Nonzero Conc	617.2422	h2*mg/L
AUMCIFO	AUMC Infinity Obs	928.5600	h2*mg/L
AUMCIFP	AUMC Infinity Pred	928.4900	h2*mg/L
AUMCPEO	AUMC %Extrapolation Obs	33.5269	%
AUMCPEP	AUMC % Extrapolation Pred	33.5219	%
VZFO	Vz Obs by F	37.6222	L
VZFP	Vz Pred by F	37.6230	L
CLFO	Total CL Obs by F	3.5914	L/h
CLFP	Total CL Pred by F	3.5914	L/h
MRTEVLST	MRT Extravasc to Last Nonzero Conc	7.7065	h
MRTEVIFO	MRT Extravasc Infinity Obs	10.4212	h
MRTEVIFP	MRT Extravasc Infinity Pred	10.4207	h





Date and Time: 2017-12-05 09:43:00 NA

Calculation Setting

Drug Administration: Extravascular

Observation count excluding trailing zero: 11

Dose at time 0: 320 mg

AUC Calculation Method: Linear-up Linear-down

Weighting for lambda z: Uniform (Ordinary Least Square, OLS)

Lambda z selection criterion: Heighest adjusted R-squared value with precision=le-4

4.4632 +1.068e-01 85.2505 473.7705

1.1700 1.1755 -5.539e-03 119.9775 977.8807

# Fitting, AUC, AUMC Result

Time	Conc.	Pred.	Residual	AUC	AUMC
0.0000	0.0000			0.0000	0.0000
0.2500	1.2500			0.1563	0.0391
0.5000	3.9600			0.8075	0.3256
1.0000	7.8200			3.7525	2.7756
2.0000	9.7200			12.5225	16.4056
3.5200	9.7500			27.3197	57.2632
5.0700	8.5700			41.5177	117.5349
7.0700	6.5900			56.6777	207.5761
9.0300 *	6.1100	6.2267 -	-1.167e-01	69.1237	307.3054

4.5700

#### Calculated Values

12.0500 \*

24.1500 \*

\_\_\_\_\_

CMAX	Max Conc	9.7500	mg/L
CMAXD	Max Conc Norm by Dose	0.0305	mg/L/mg
TMAX	Time of CMAX	3.5200	h
TLAG	Time Until First Nonzero Conc	0.0000	h
CLST	Last Nonzero Conc	1.1700	mg/L
CLSTP	Last Nonzero Conc Pred	1.1755	mg/L
TLST	Time of Last Nonzero Conc	24.1500	h
LAMZHL	Half-Life Lambda z	6.2865	h
LAMZ	Lambda z	0.1103	/h
LAMZLL	Lambda z Lower Limit	9.0300	h
LAMZUL	Lambda z Upper Limit	24.1500	h
LAMZNPT	Number of Points for Lambda z	3	
CORRXY	Correlation Between TimeX and Log ConcY	-0.9997	
R2	R Squared	0.9994	
R2ADJ	R Squared Adjusted	0.9988	
AUCLST	AUC to Last Nonzero Conc	119.9775	h*mg/L
AUCALL	AUC All	119.9775	h*mg/L

<sup>\*:</sup> Used for the calculation of Lambda z.

AUCIFO	AUC Infinity Obs	130.5888	h*mg/L
AUCIFOD	AUC Infinity Obs Norm by Dose	0.4081	h*mg/L/mg
AUCIFP	AUC Infinity Pred	130.6391	h*mg/L
AUCIFPD	AUC Infinity Pred Norm by Dose	0.4082	h*mg/L/mg
AUCPEO	AUC %Extrapolation Obs	8.1258	%
AUCPEP	AUC %Extrapolation Pred	8.1611	%
AUMCLST	AUMC to Last Nonzero Conc	977.8807	h2*mg/L
AUMCIFO	AUMC Infinity Obs	1330.3840	h2*mg/L
AUMCIFP	AUMC Infinity Pred	1332.0528	h2*mg/L
AUMCPEO	AUMC %Extrapolation Obs	26.4964	%
AUMCPEP	AUMC % Extrapolation Pred	26.5884	%
VZFO	Vz Obs by F	22.2243	L
VZFP	Vz Pred by F	22.2157	L
CLFO	Total CL Obs by F	2.4504	L/h
CLFP	Total CL Pred by F	2.4495	L/h
MRTEVLST	MRT Extravasc to Last Nonzero Conc	8.1505	h
MRTEVIFO	MRT Extravasc Infinity Obs	10.1876	h
MRTEVIFP	MRT Extravasc Infinity Pred	10.1964	h



