

FIGURE 1A

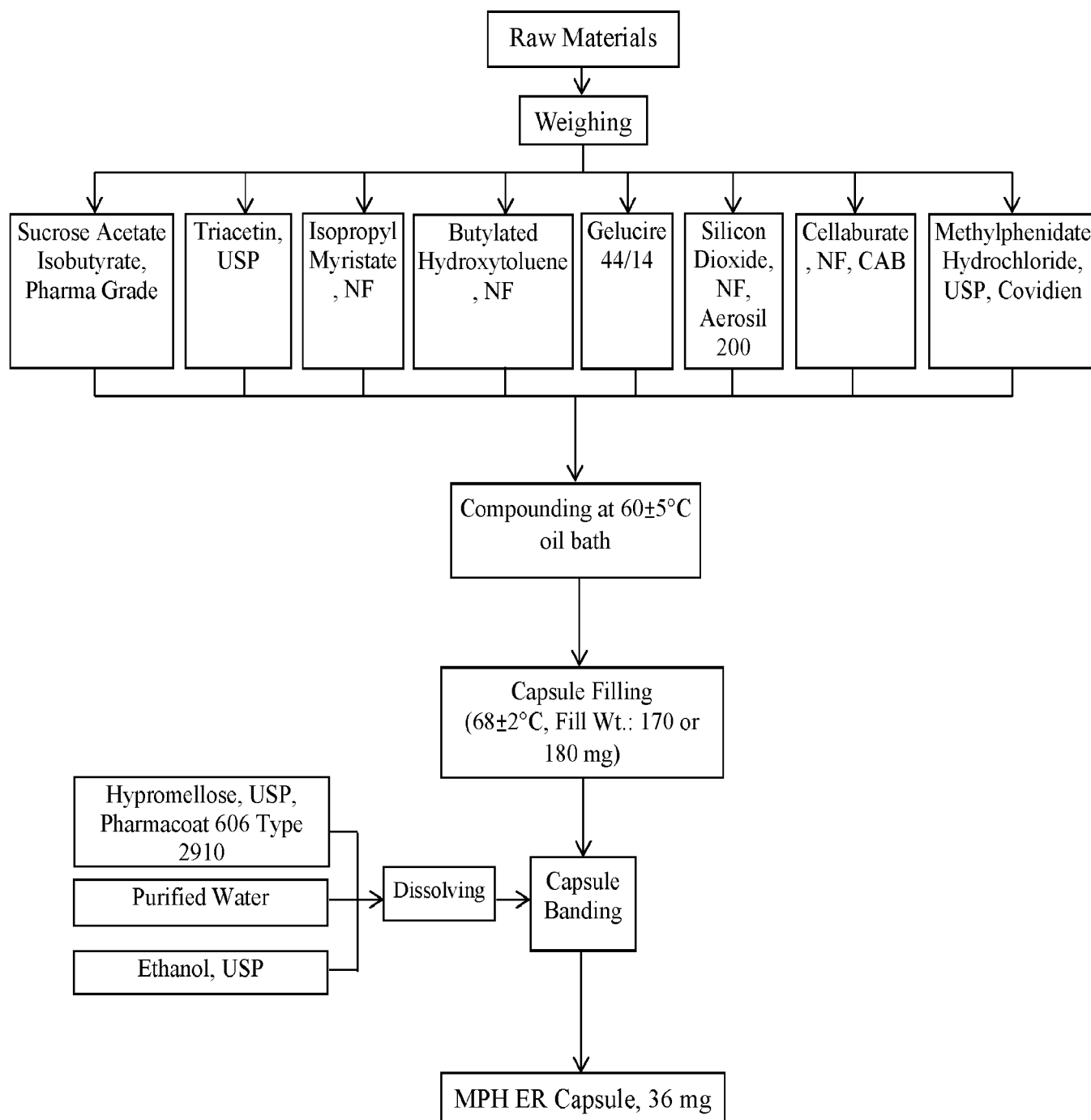
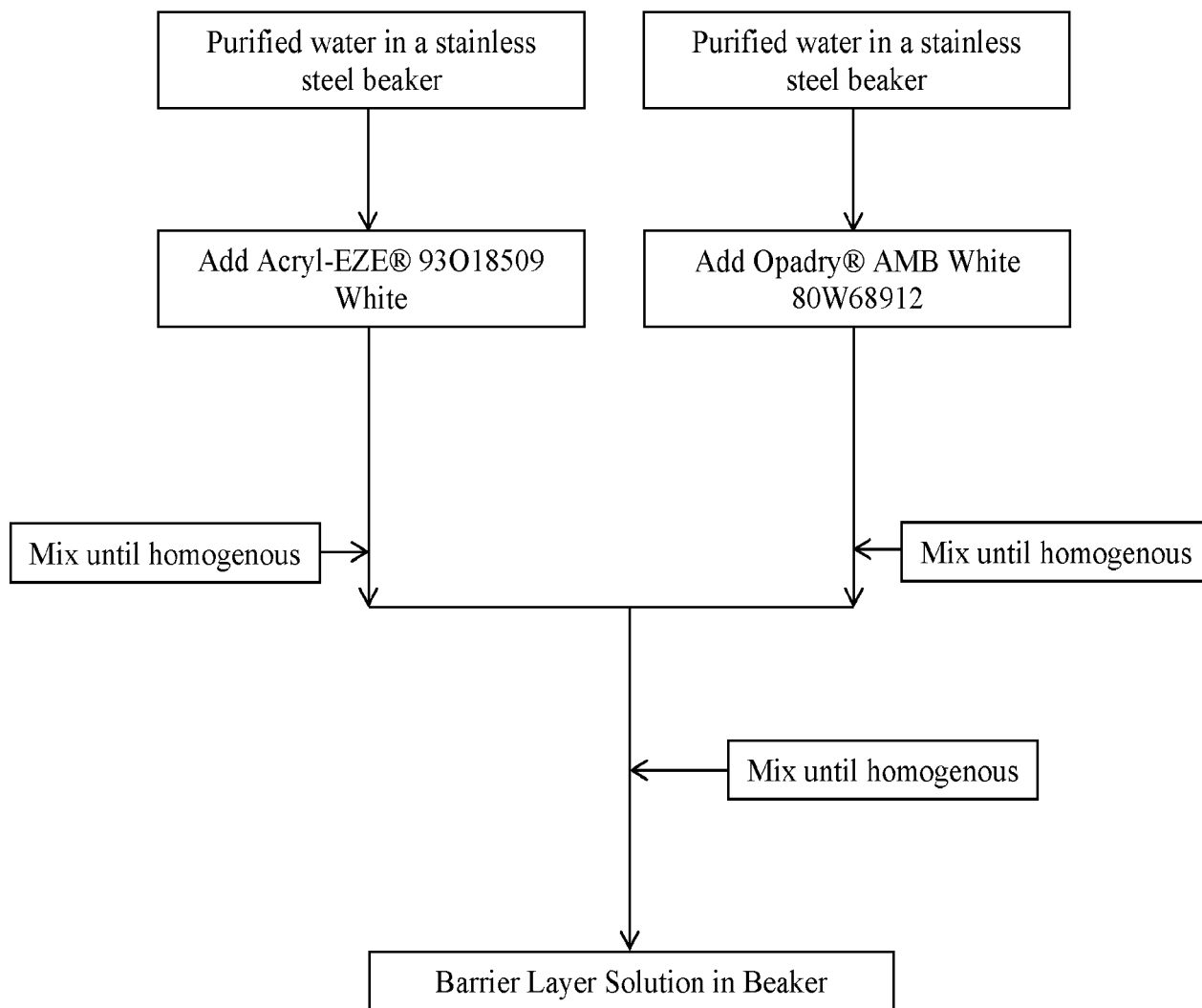
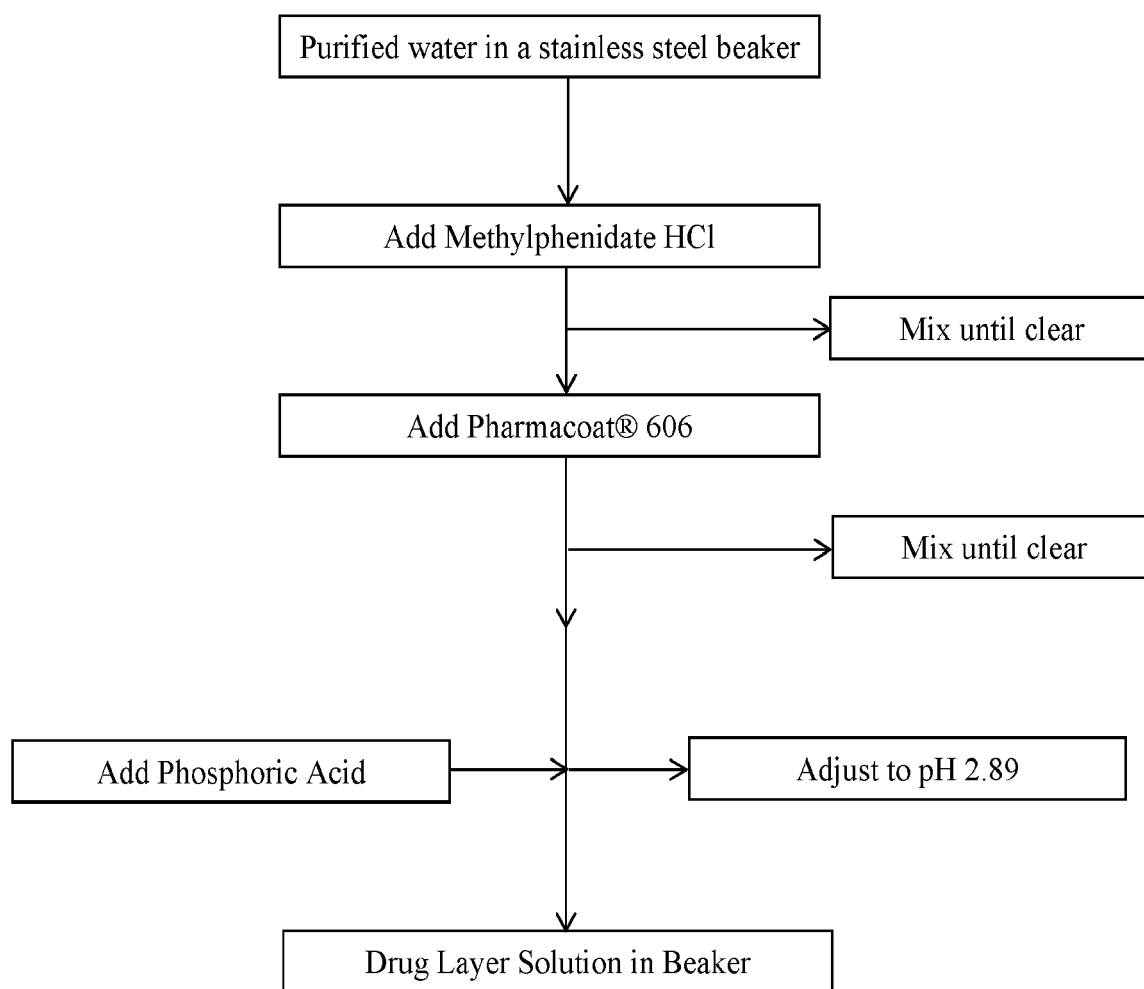


FIGURE 1B



3/24

FIGURE 2



4/24

FIGURE 3

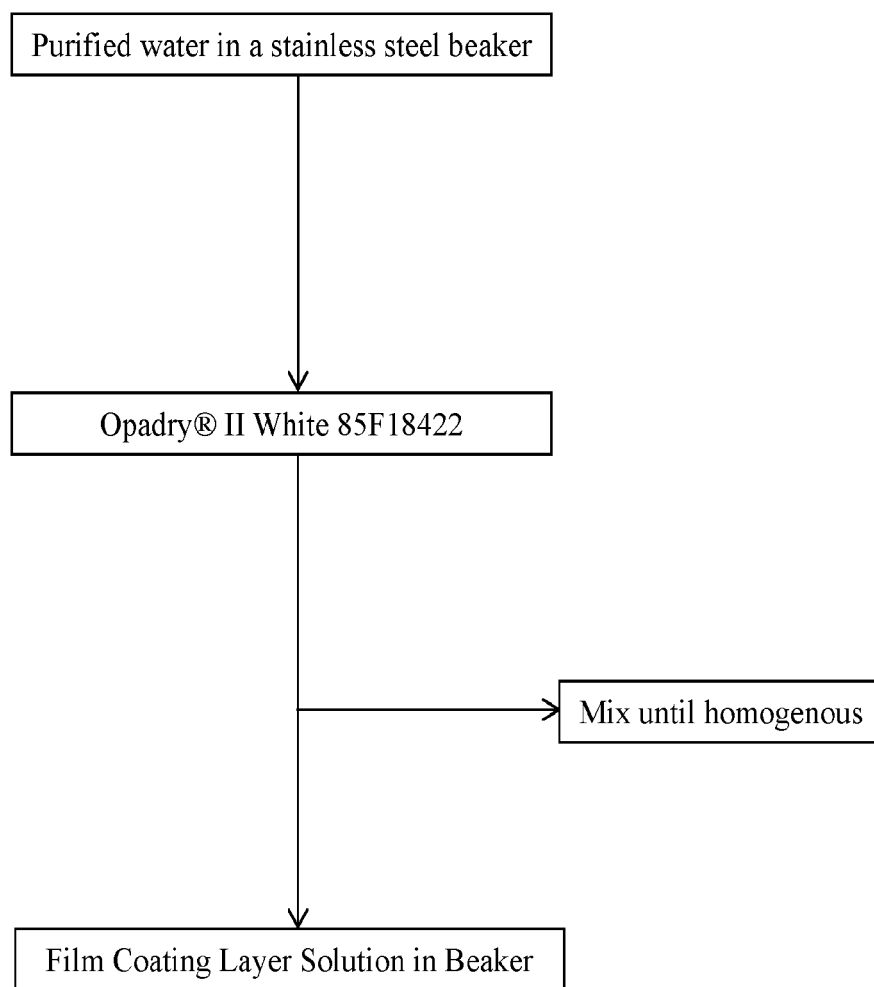
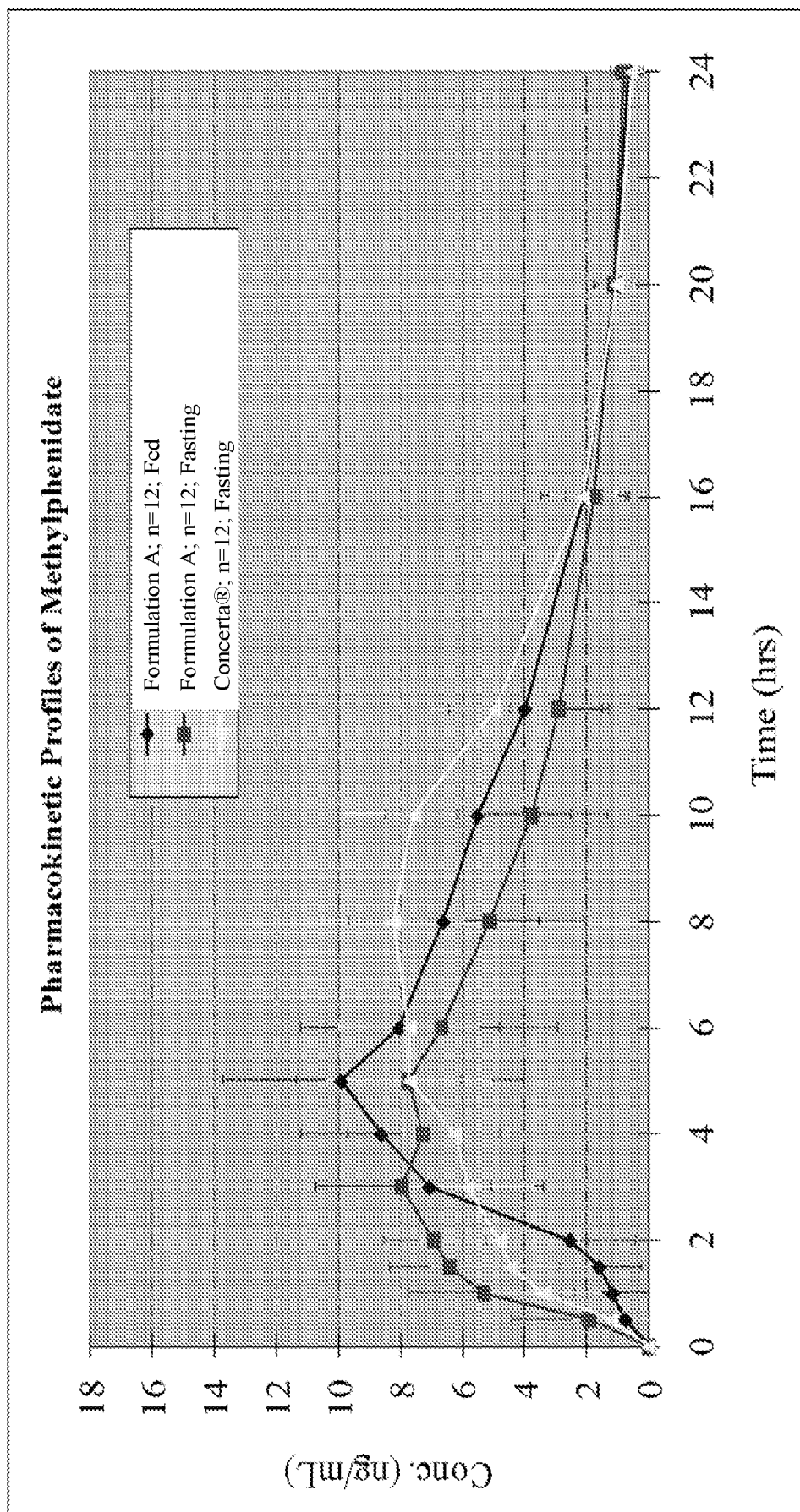


FIGURE 4



6/24

FIGURE 5(A)

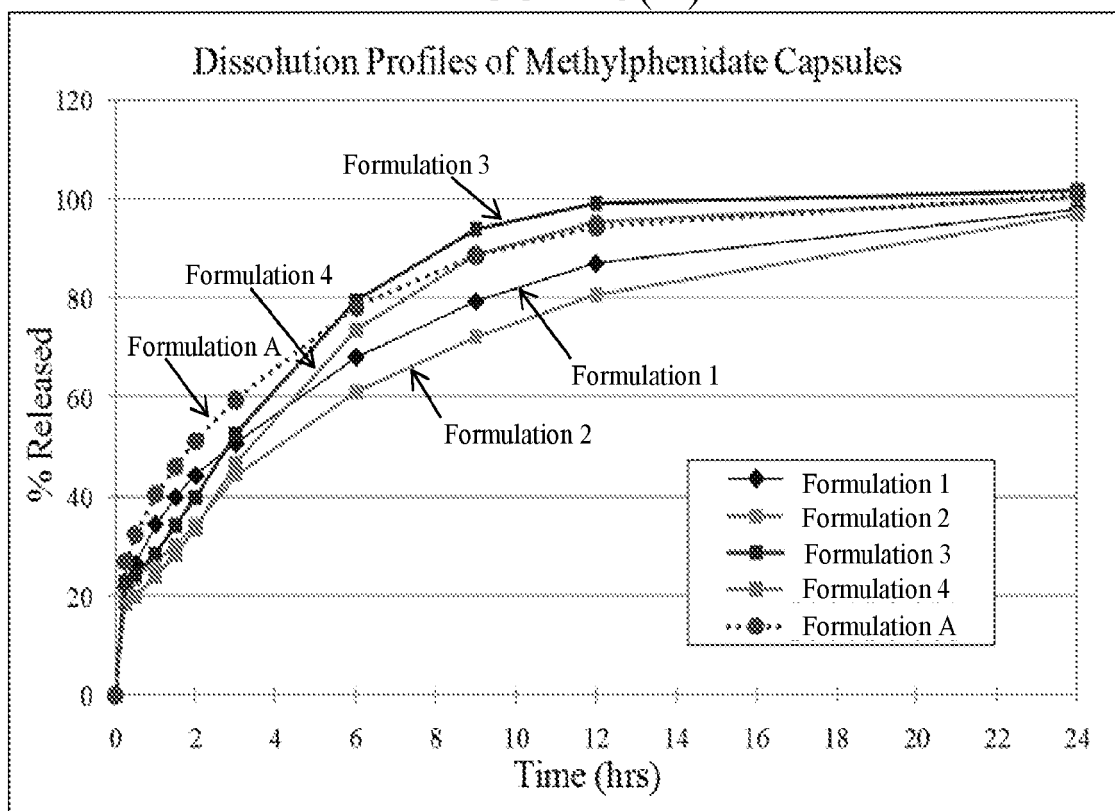
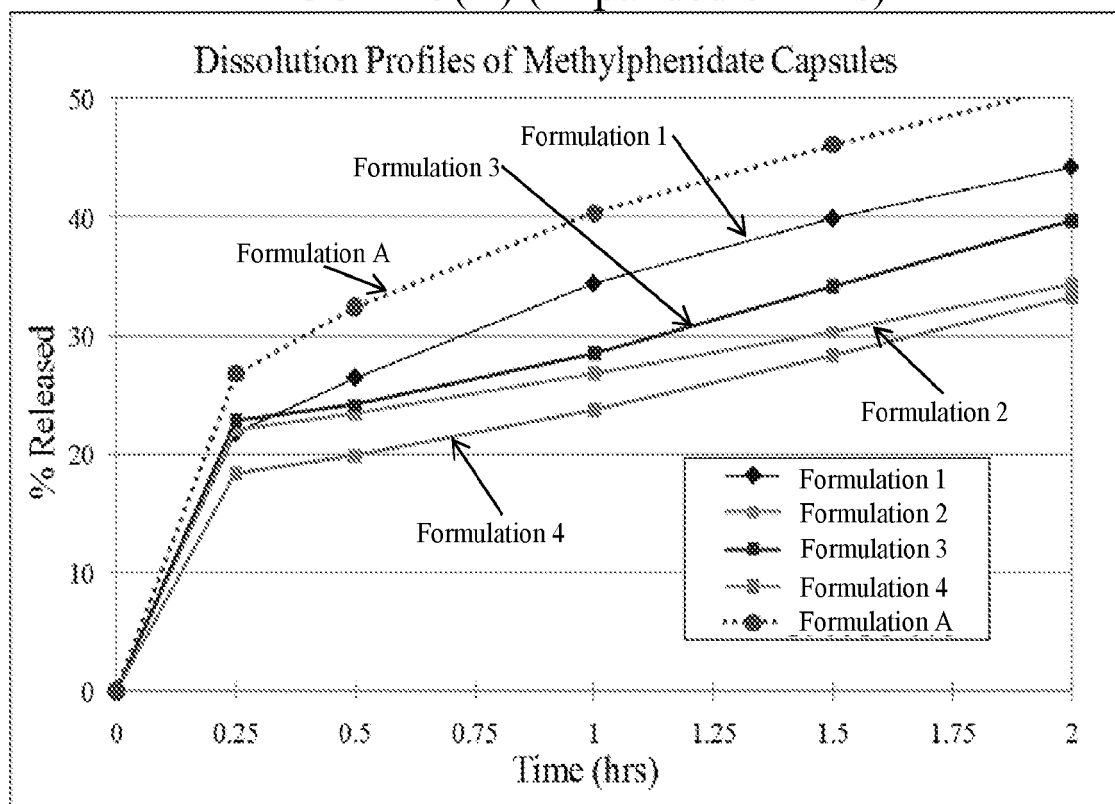


FIGURE 5(B) (Expanded 0-2 hrs)



7/24

FIGURE 5(C) (Higuchi-Plot)

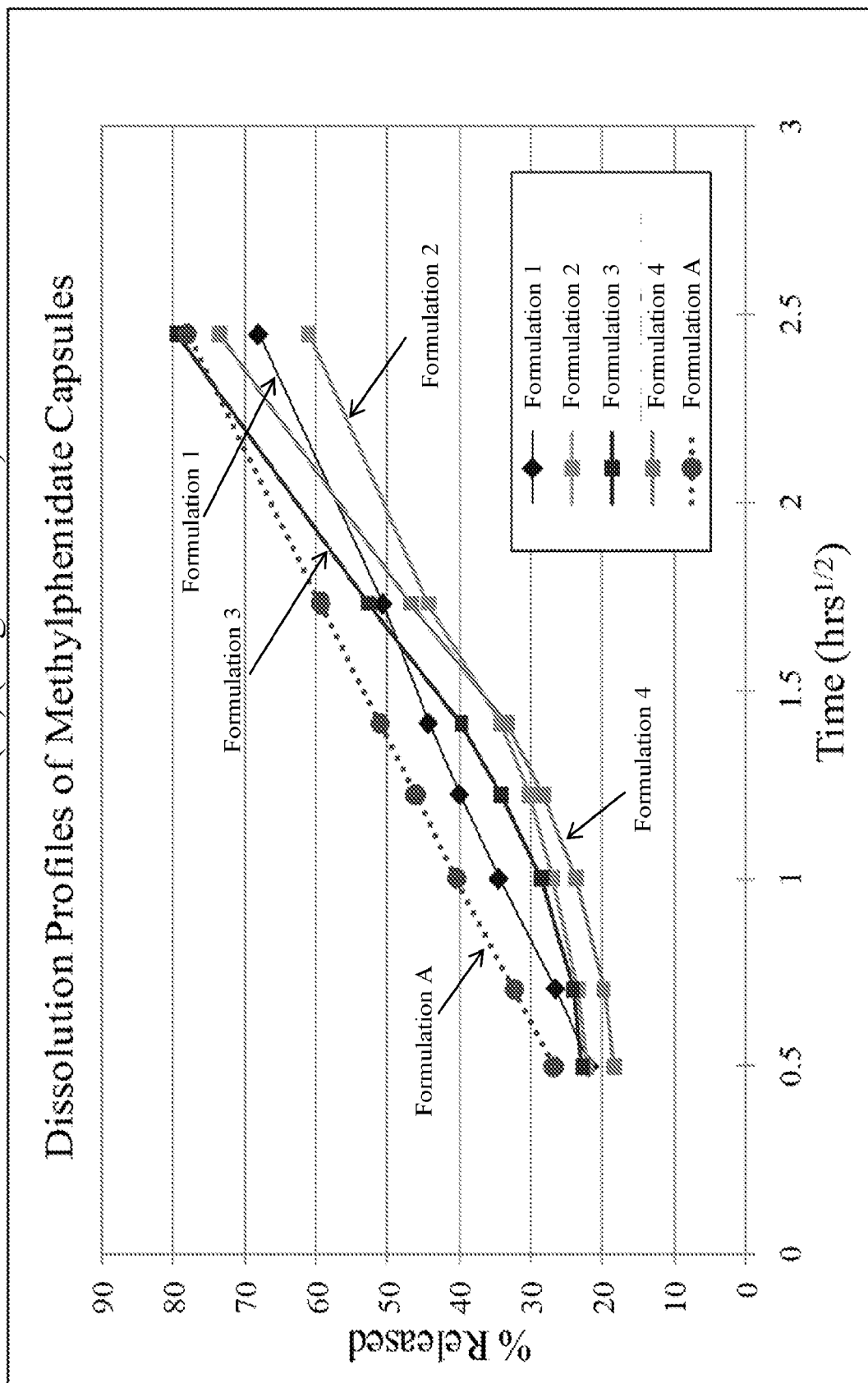


FIGURE 6
Dissolution Profiles of Methylphenidate Capsules

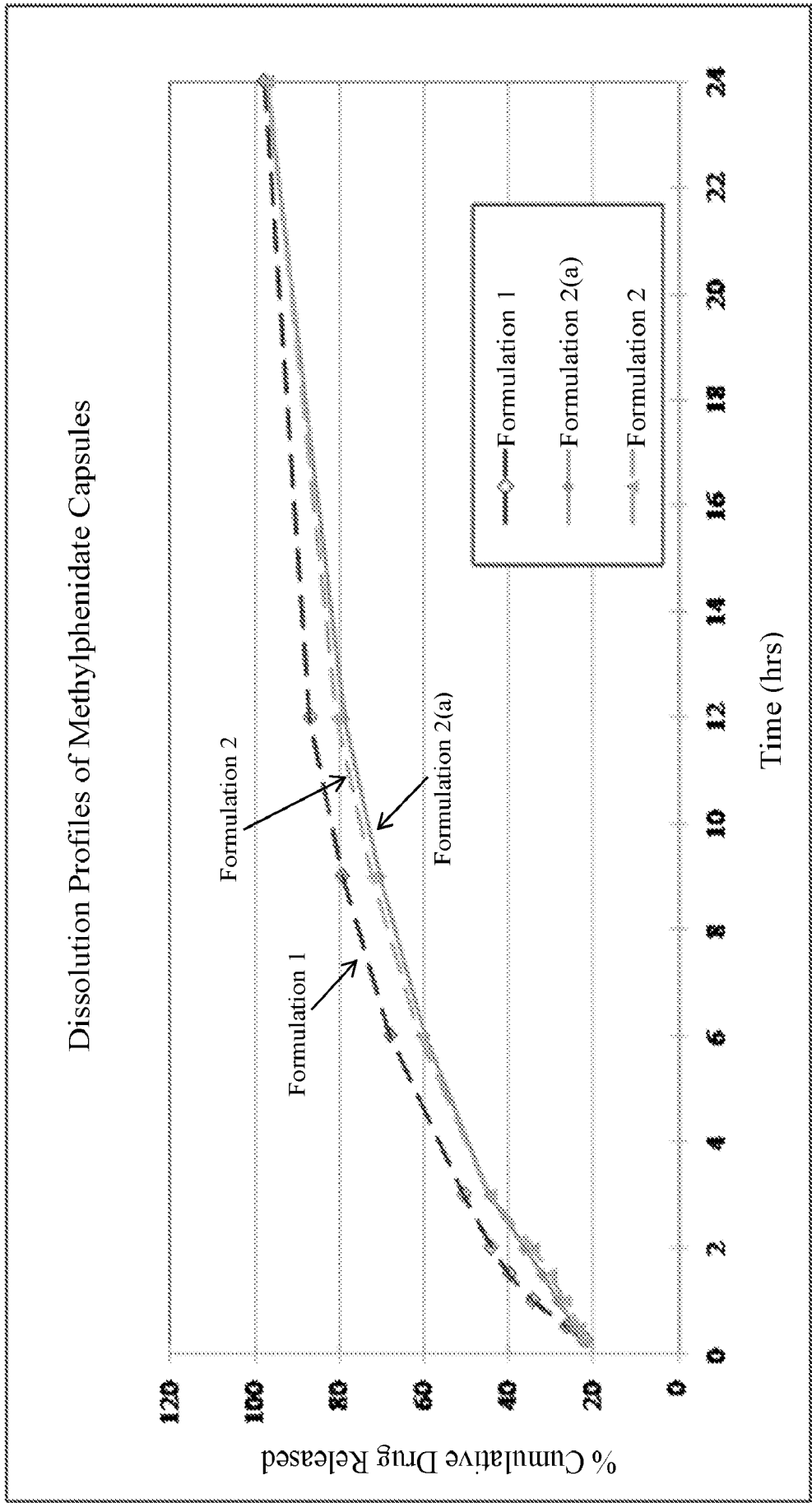
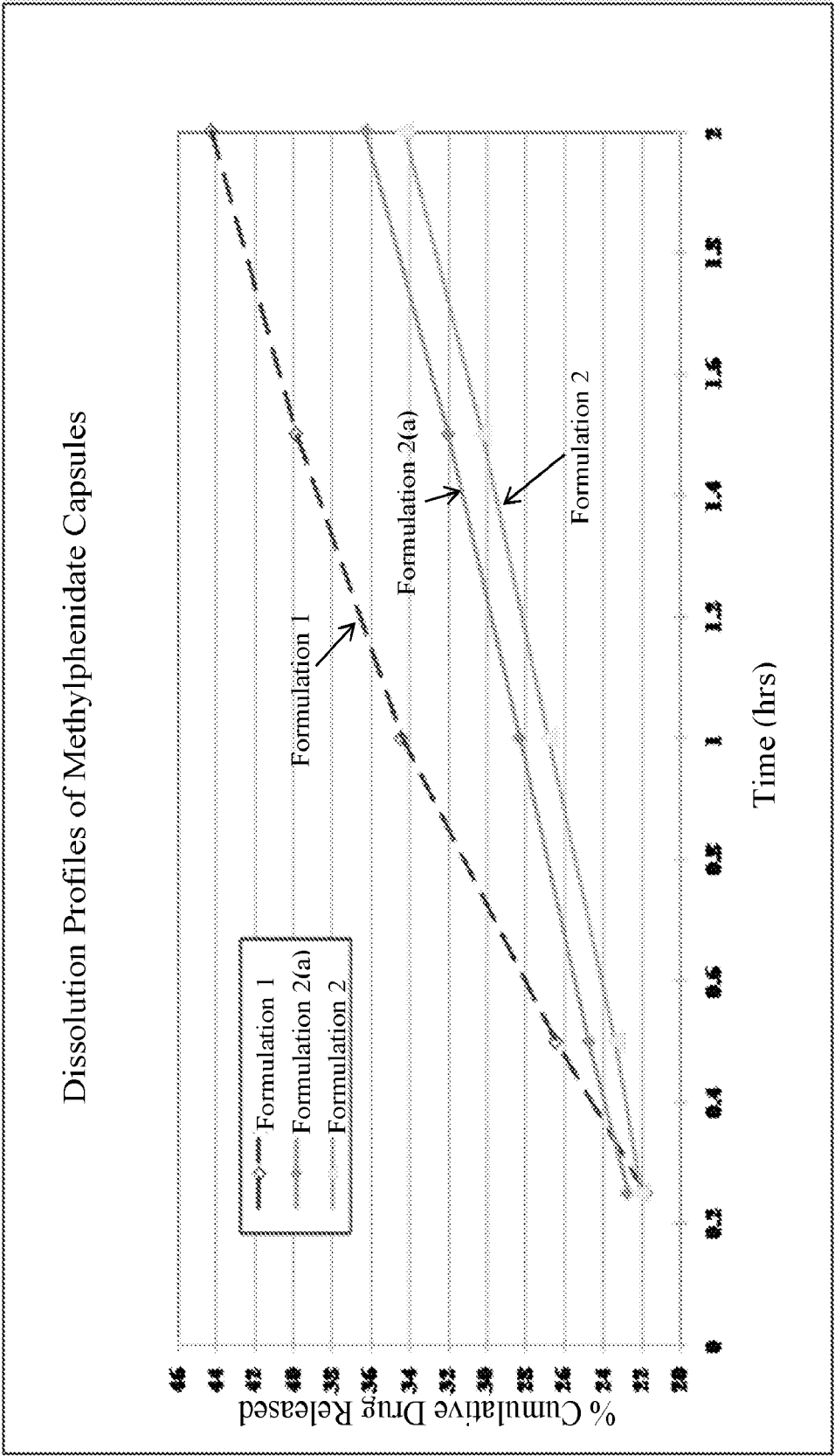


FIGURE 7



10/24

FIGURE 8

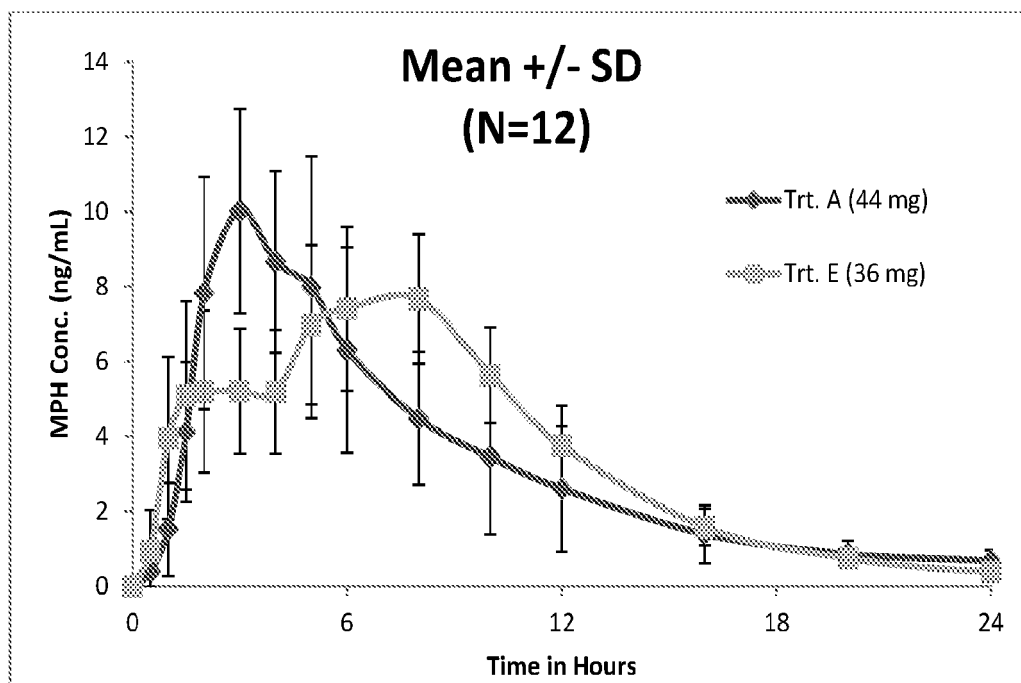
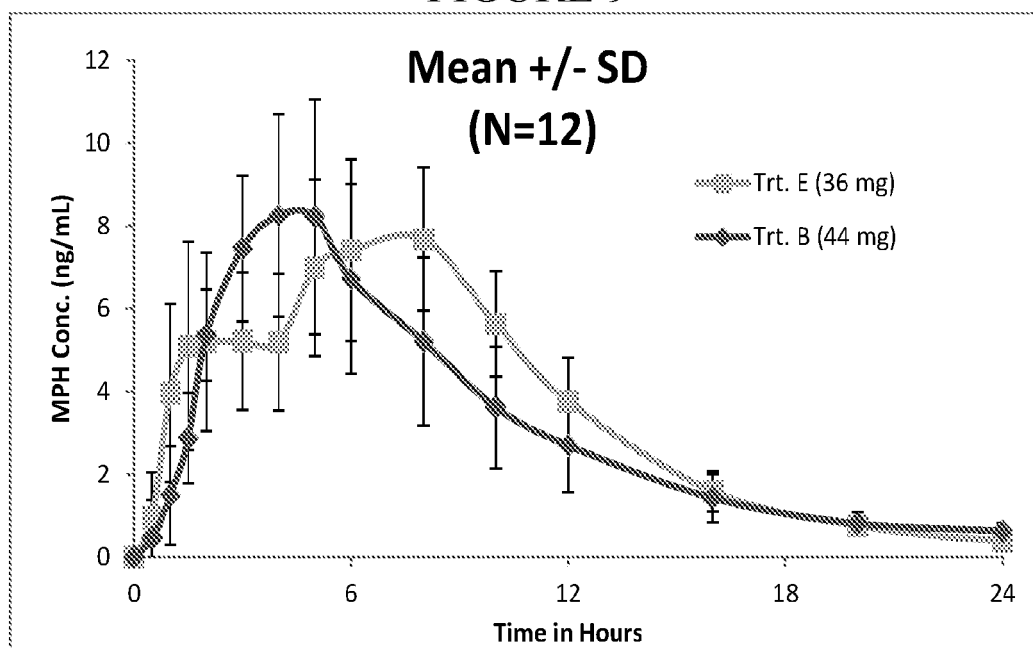


FIGURE 9



11/24

FIGURE 10

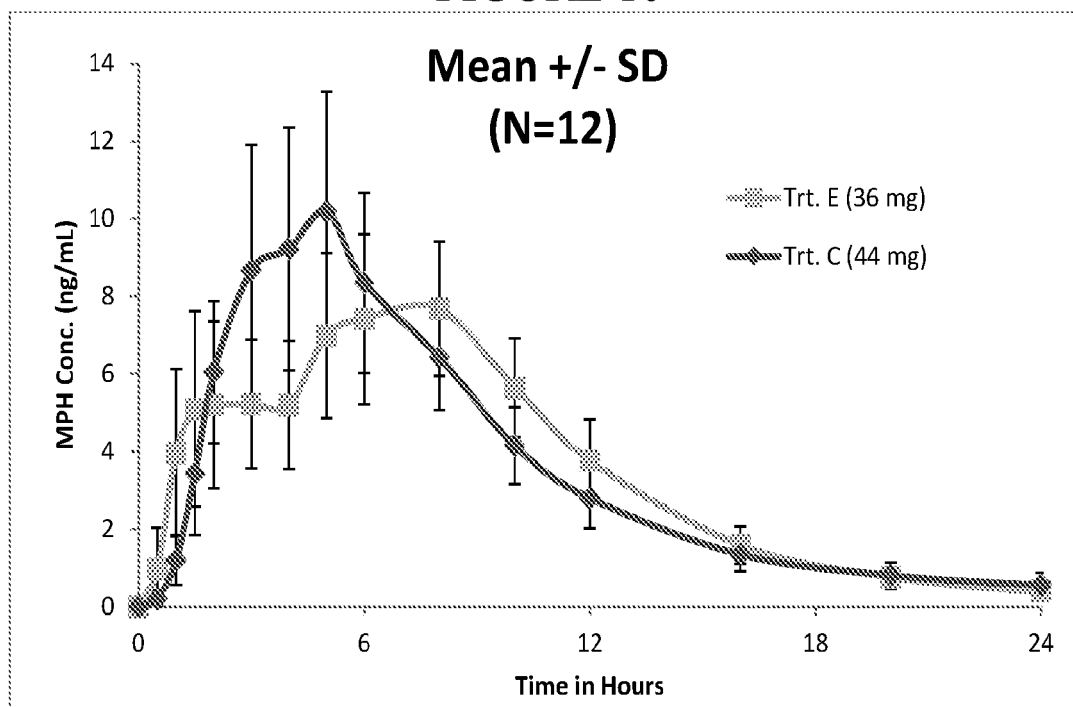
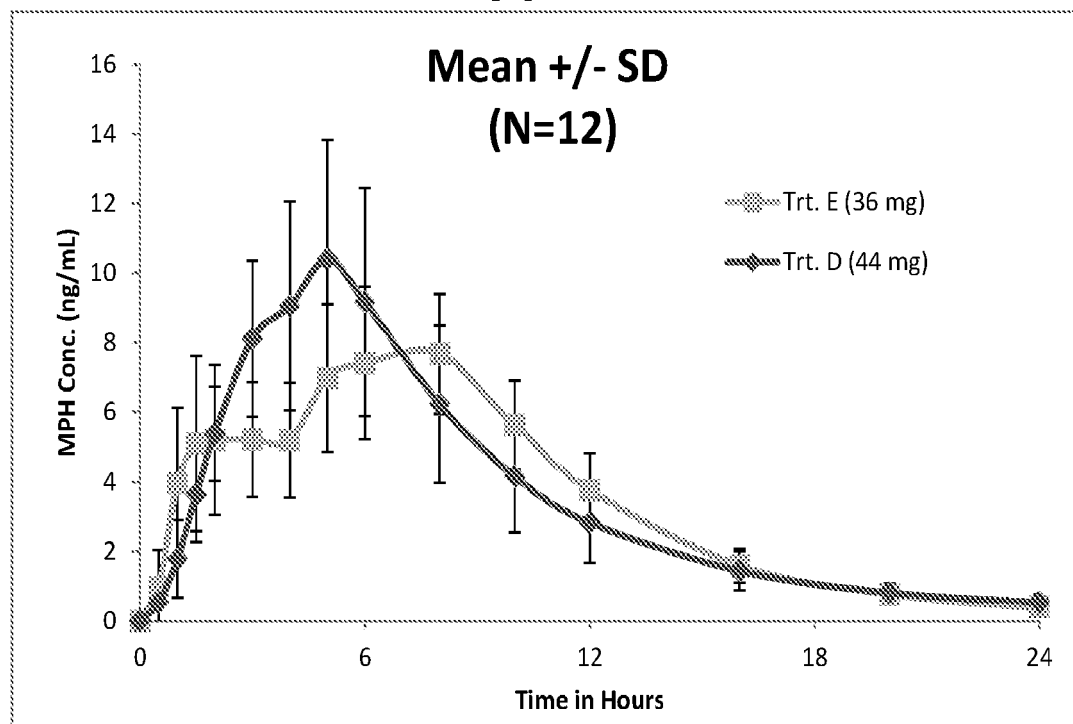


FIGURE 11



12/24

FIGURE 12

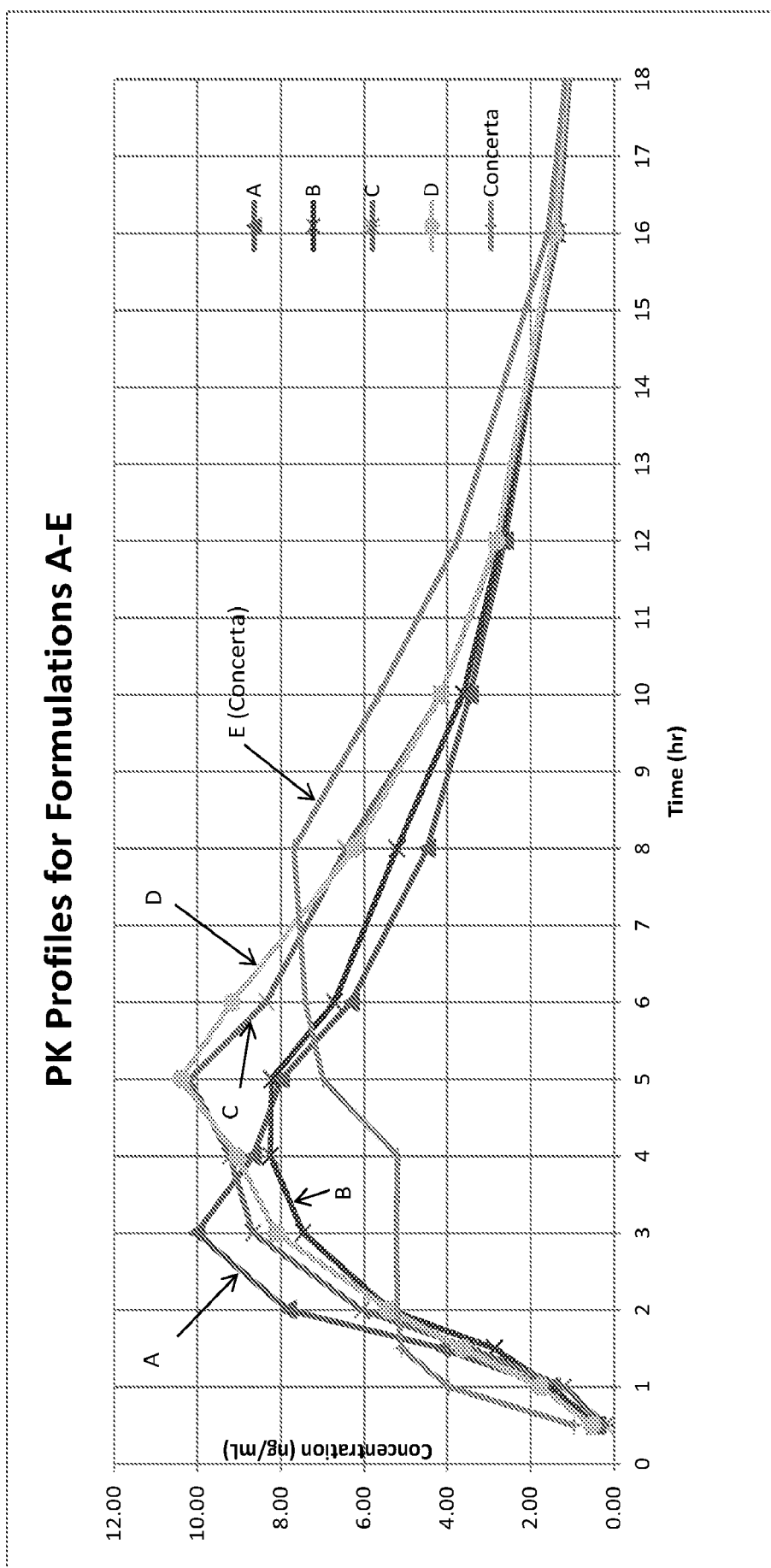


FIGURE 13
PK Profiles for Examples 2 and 6

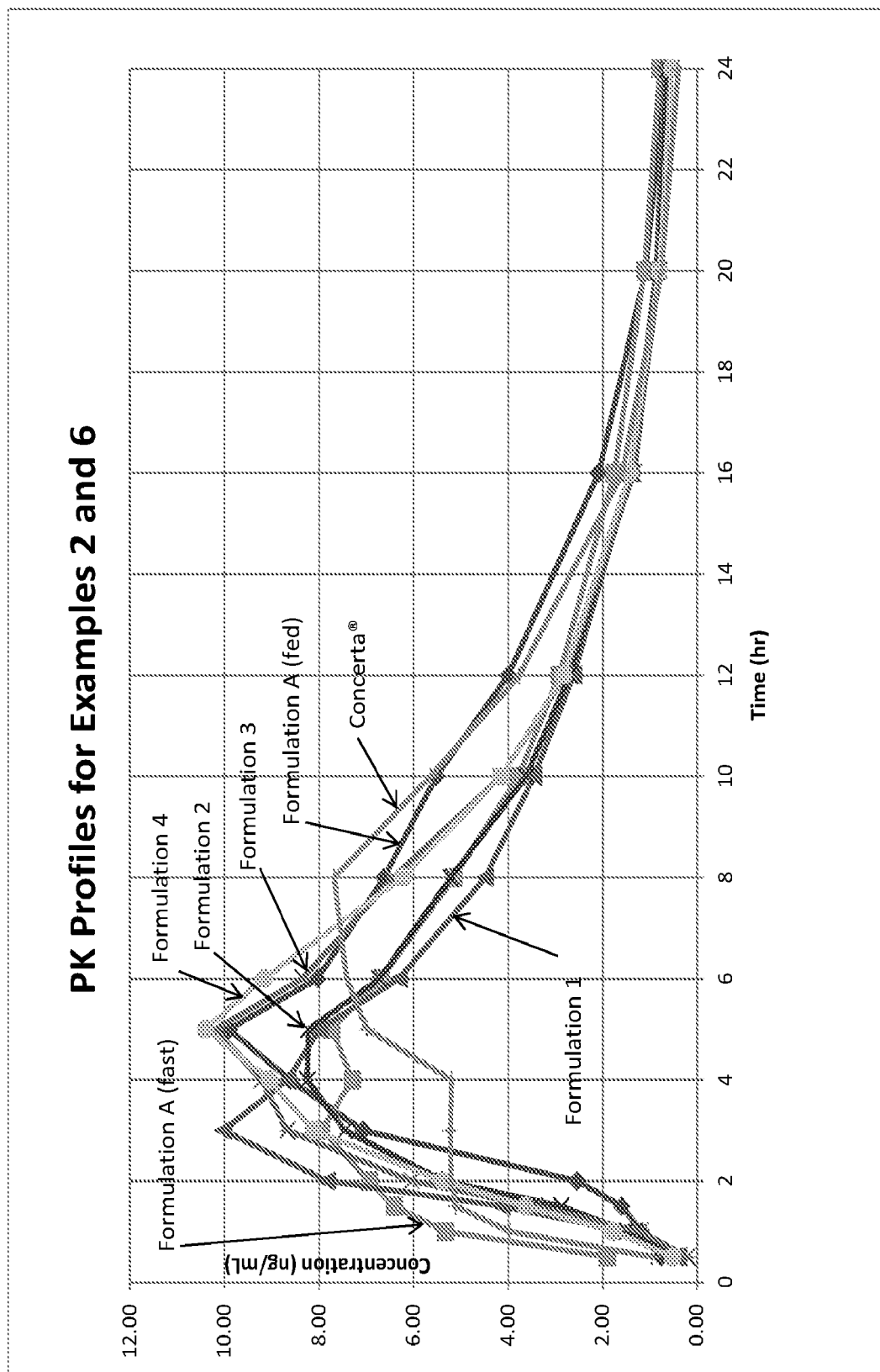


FIGURE 14

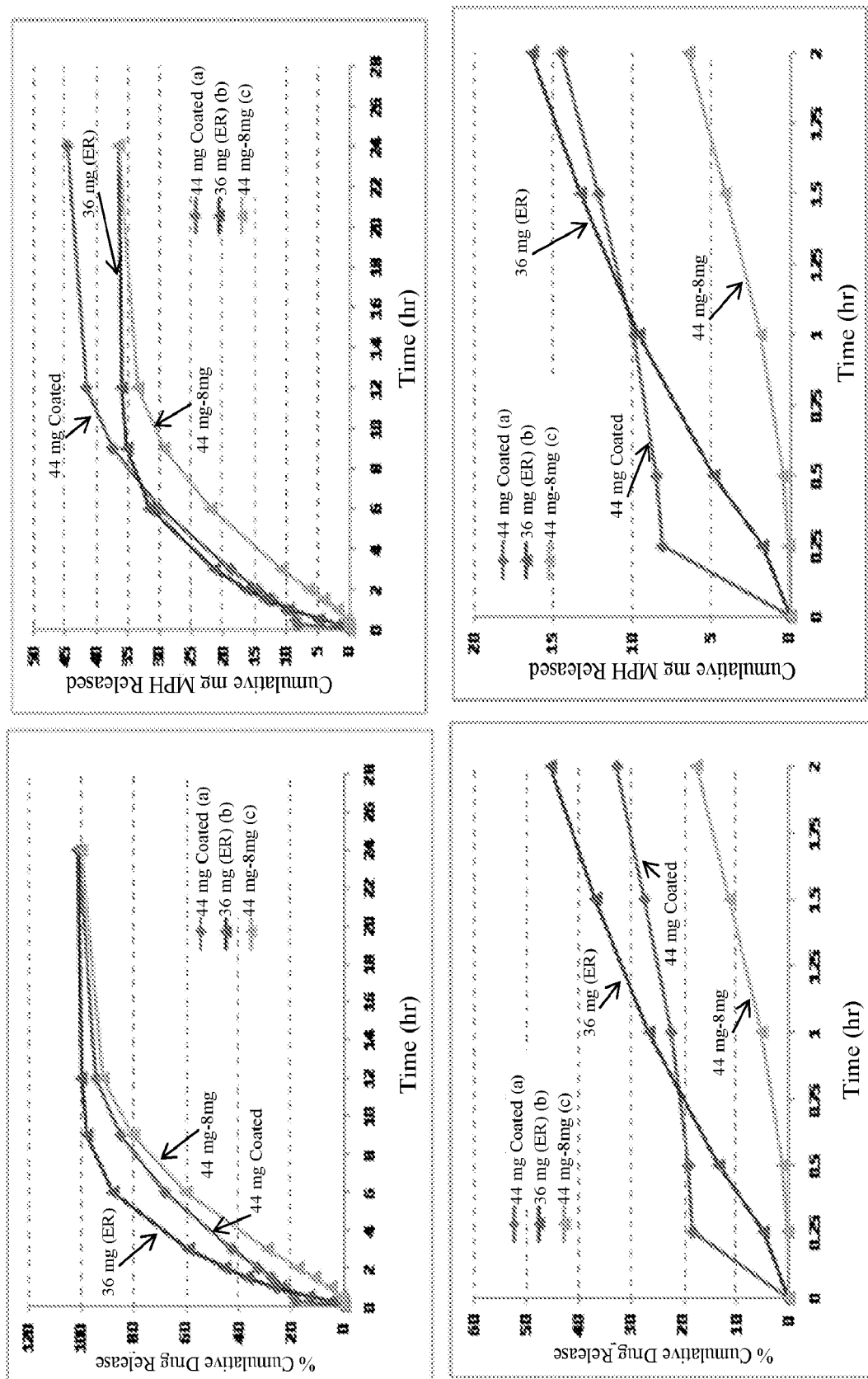


FIGURE 15

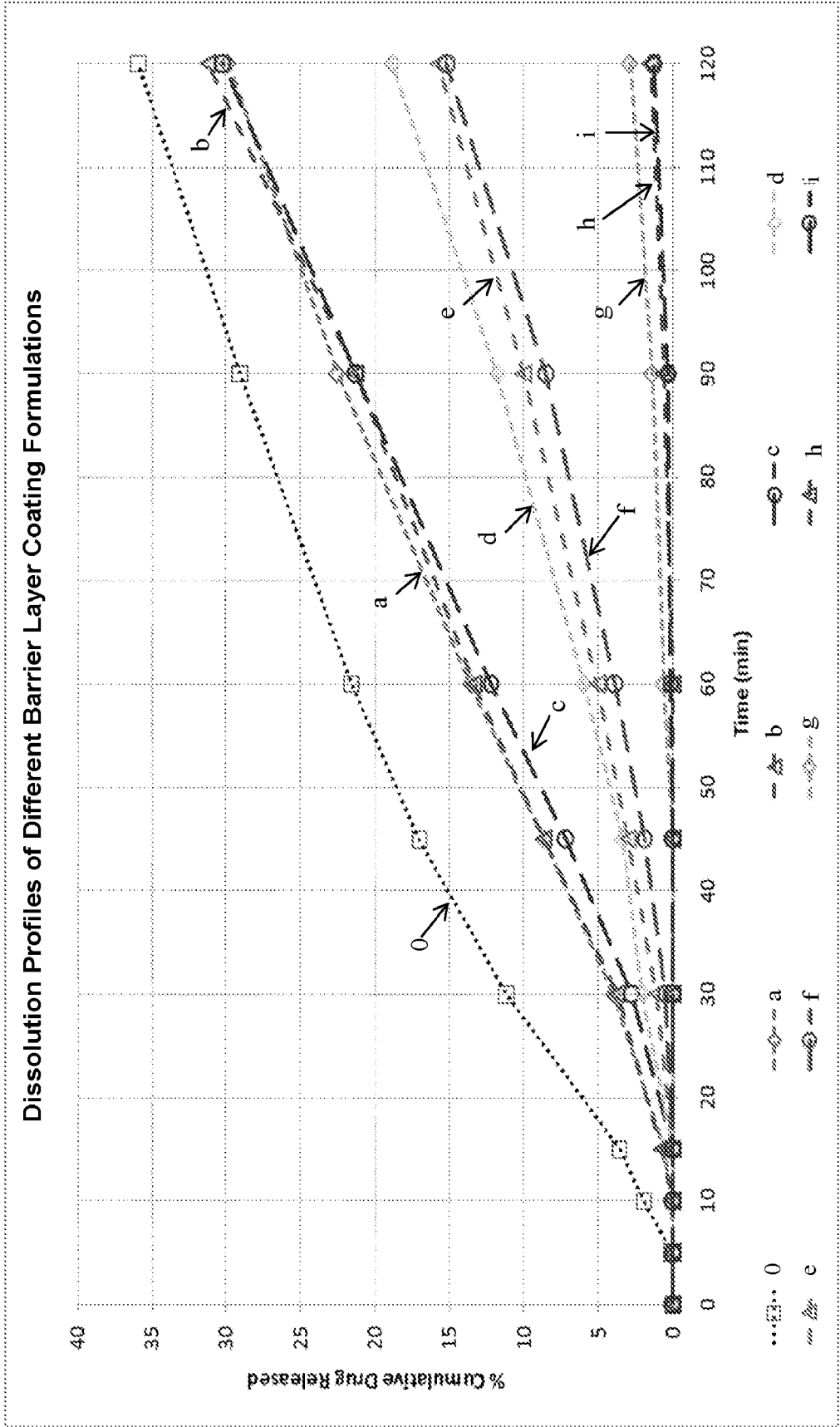


FIGURE 16

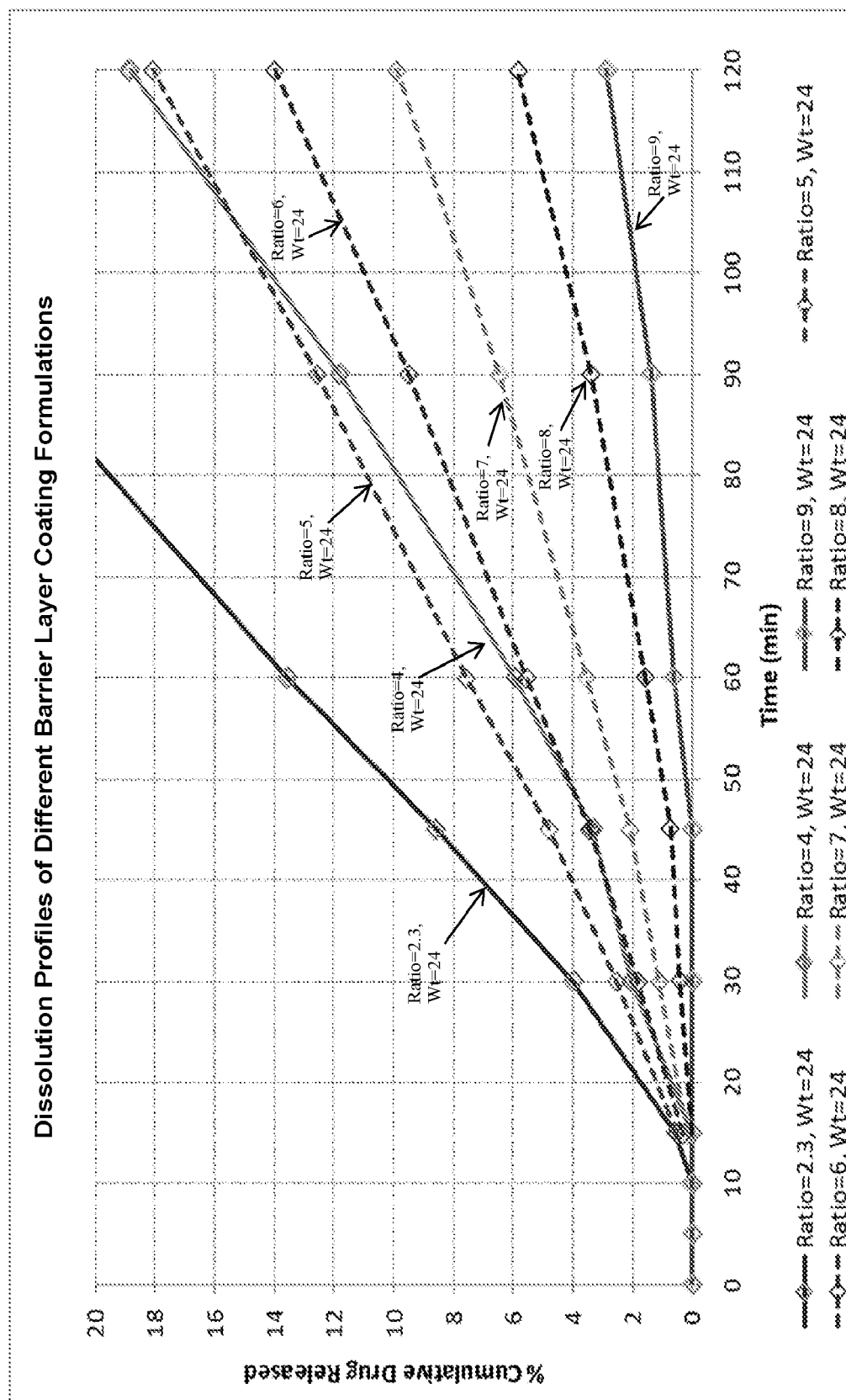


FIGURE 17

Dissolution Profiles of Different Delay Layer Coating Formulations

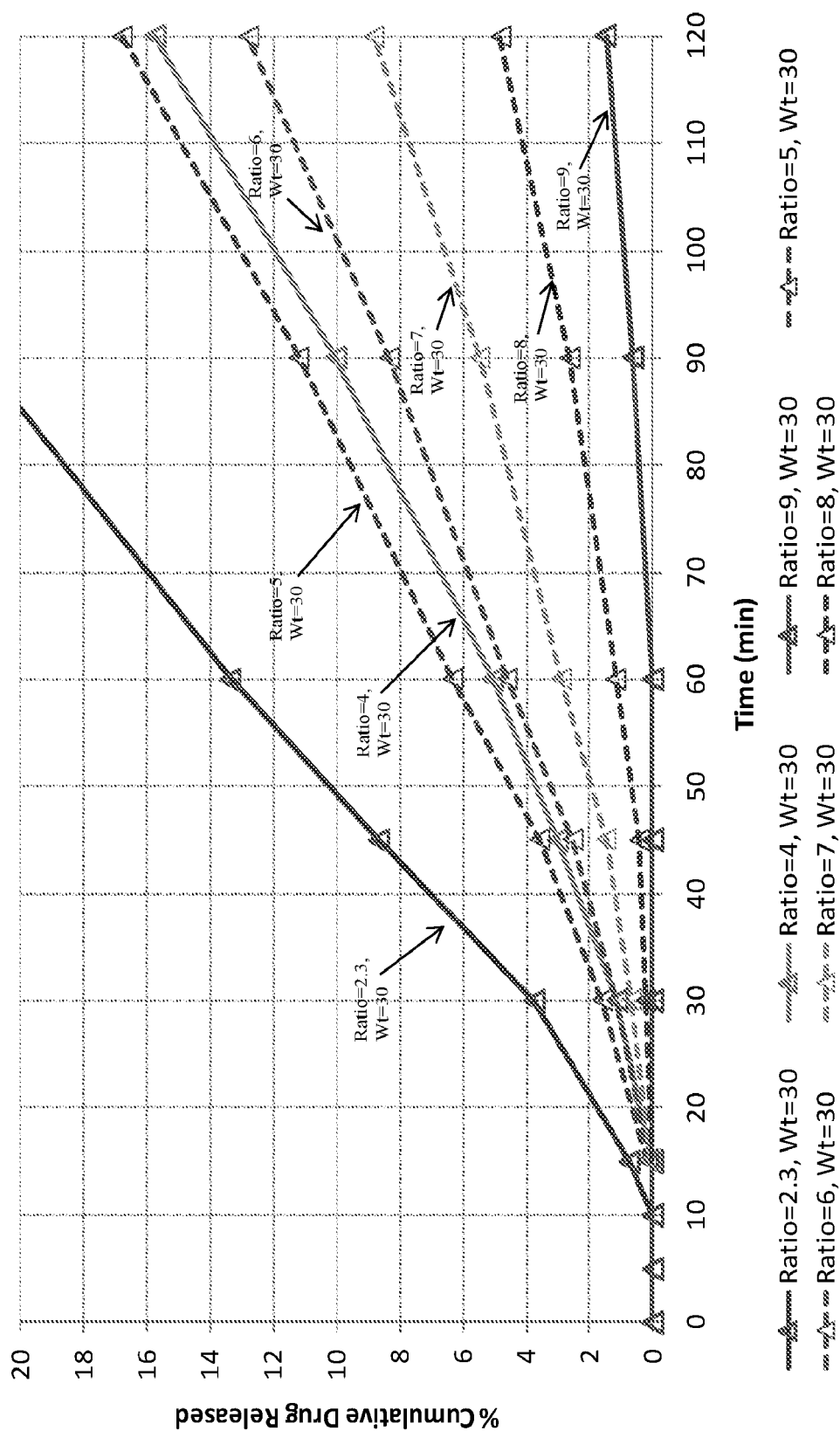


FIGURE 18

Dissolution Profiles of Different Delay Layer Coating Formulations

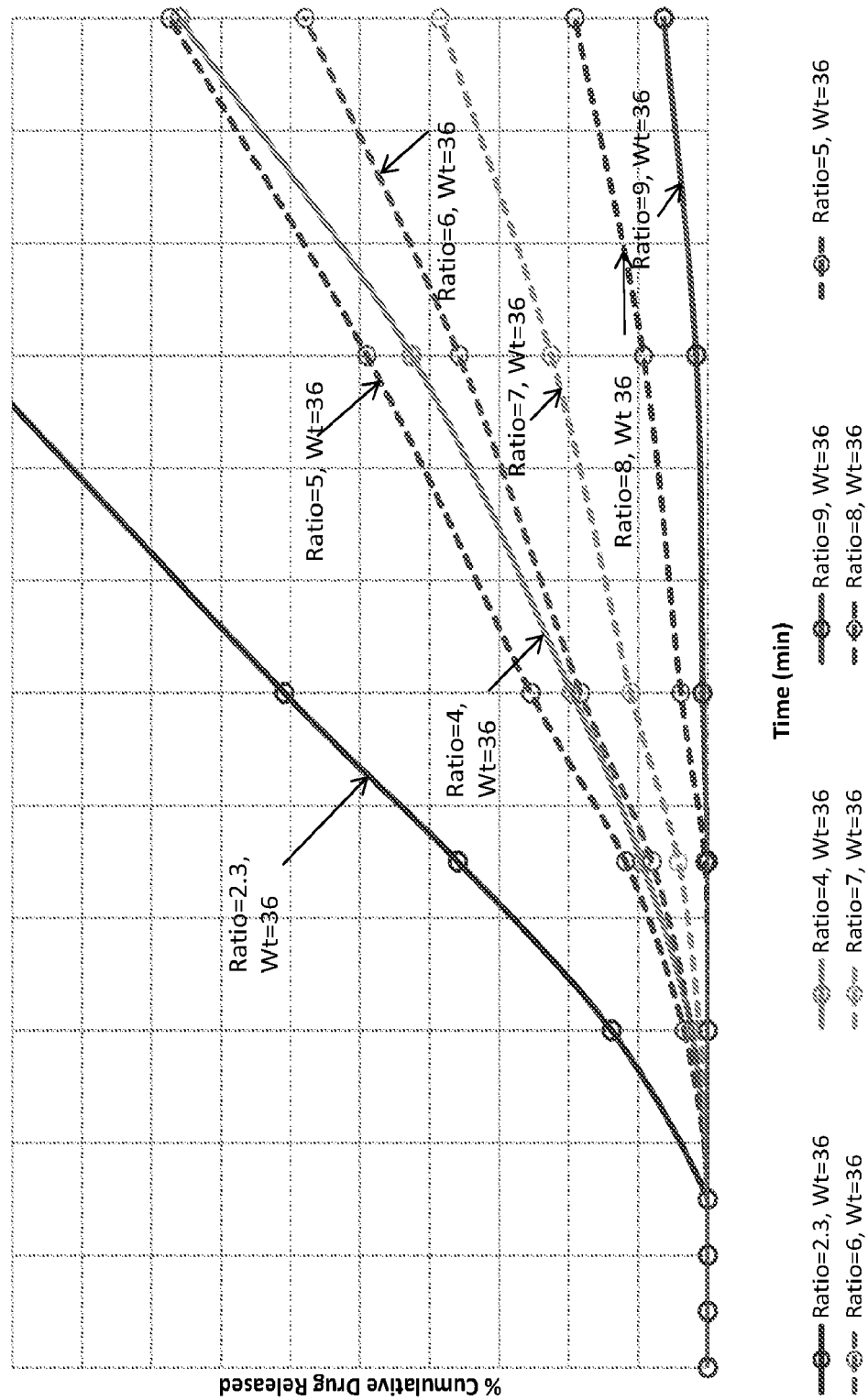


FIGURE 19A
22 mg MPH capsules

Lot No.	Packaging Material / Stability Conditions	Time Point (Month)	% Cumulative Drug Released				Organic Impurities				Microbial Limit Tests			
			1 HR	3 HR	9 HR	24 HR	MPH Related Compound A (LOQ=0.99%)	Erythor Isomer (LOQ=0.06%)	Other Degradation Product (LOQ=0.10%)	Total Impurity	Assay (% L.C)	TAMC (cfu/g)	TYMC (cfu/g)	E. Coli
A	120cc HDPE Bottle 25°C/60%RH	0	29	62	102	105	ND	ND	ND	<LOQ	103.7	<10	<10	Negative
		3	29	61	101	105	ND	ND	ND	<LOQ	103.2	-	-	-
		6	28	61	100	104	0.11	ND	ND	0.11	103.8	-	-	-
		9	28	58	99	103	0.16	ND	ND	0.16	104.5	-	-	-
		12	29	65	97	101	0.20	ND	ND	0.20	102.8	<10	<10	Negative
		18	30	56	96	103	0.32	ND	ND	0.32	102.5	-	-	-
	120cc HDPE Bottle 30°C/75%RH	24	29	56	93	103	0.43	ND	ND	0.43	104.3	<10	<10	Negative
		0	29	62	102	105	ND	ND	ND	<LOQ	103.7	<10	<10	Negative
		3	27	59	100	104	0.16	ND	ND	0.16	105.5	-	-	-
		6	29	58	98	103	0.24	ND	ND	0.24	103.2	-	-	-
		9	29	62	98	103	0.39	ND	ND	0.39	103.3	-	-	-
		12	27	55	96	102	0.52	ND	ND	0.52	102.6	<10	<10	Negative
	120cc HDPE Bottle 40°C/75%RH	18	30	57	92	103	0.81	ND	ND	0.81	101.3	-	-	-
		24	28	53	90	103	1.04	0.068	ND	1.11	103.0	<10	<10	Negative
		0	29	62	102	105	ND	ND	ND	<LOQ	103.7	<10	<10	Negative
		1	28	62	103	106	0.26	ND	ND	0.20	103.6	-	-	-
		3	28	62	100	102	0.56	ND	ND	0.60	104.4	-	-	-
		6	29	60	99	103	0.83	0.06	ND	0.90	102.4	-	-	-

ND = Not Detected, LOQ = Limit of Quantification, TAMC = Total Aerobic Microbial Count, TYMC = Total Combined Yeast/Molds Count

FIGURE 19B
22 mg MPH capsules

Lot No.	Packaging Material / Stability Conditions	Time Point (Month)	Water Content	% Cumulative Drug Released				Organic Impurities				Assay (%LC)	Microbial Limit Tests	
				1 HR	3 HR	9 HR	24 HR	MPH Related Compound A (LOQ=0.09%)	Erythor Isomer (LOQ=0.06%)	Degradation Product (LOQ=0.10%)	Total Impurity		TAMC (cfu/g)	TYMC (cfu/g)
B	Blister-PVC/PVDC 40g with aluminum bag 25°C/60%RH	0	1.1	31	48	74	90	0.10	ND	ND	0.10	102.9	690	10
		3	1.9	32	48	74	88	0.15	ND	ND	0.15	104.3	-	-
		6	1.5	33	50	76	88	0.18	ND	ND	0.18	104.6	-	-
		9	1.0	33	50	72	82	0.2	ND	ND	0.2	104.2	-	-
		13	1.6	31	49	78	92	0.2	ND	ND	0.2	105.3	700	<10

ND = Not Detected, TAMC = Total Aerobic Microbial Count, TYMC = Total Combined Yeast/Molds Count

FIGURE 20A
33 mg MPH capsules

Lot No.	Packaging Material / Stability Conditions	Time Point (Month)	% Cumulative Drug Released				Organic Impurities				Assay (% L.C.)	Microbial Limit Tests		
			1 HR	3 HR	9 HR	24 HR	MPH Related Compound A (LOQ=0.13%)	Erythor Isomer (LOQ=0.08%)	Other Degradation Product (LOQ=0.13%)	Total Impurity		TAMC (cfu/g)	TYMC (cfu/g)	E. Coli
C	120cc HDPE Bottle 25°C/60%RH	0	24	46	90	103	<LOD	ND	ND	<LOQ	103.5	<10	<10	Negative
		3	24	49	89	101	<LOD	ND	ND	<LOQ	102.1	-	-	-
		6	24	47	86	101	<LOD	ND	ND	<LOQ	100.8	-	-	-
		9	26	52	90	102	0.16	ND	ND	0.16	101.3	-	-	-
		12	23	44	83	100	0.21	ND	ND	0.21	101.1	<10	<10	Negative
		18	24	48	86	101	0.35	ND	ND	0.35	100.3	-	-	-
	120cc HDPE Bottle 30°C/75%RH	24	24	47	84	103	0.49	ND	ND	0.49	102.1	<10	<10	Negative
		0	24	46	90	103	<LOD	ND	ND	<LOQ	103.5	<10	<10	Negative
		3	25	50	86	100	0.16	ND	ND	0.16	102.2	-	-	-
		6	24	49	91	101	0.25	ND	ND	0.25	101.4	-	-	-
		9	25	51	89	102	0.42	ND	ND	0.42	100.1	-	-	-
		12	24	48	85	102	0.55	ND	ND	0.55	100.4	<10	<10	Negative
	120cc HDPE Bottle 40°C/75%RH	18	24	46	86	102	0.85	ND	ND	0.85	99.2	-	-	-
		24	25	46	81	105	1.11	ND	ND	1.11	99.3	<10	<10	Negative
		0	24	46	90	103	<LOD	<LOD	<LOD	<LOQ	103.5	<10	<10	Negative
		1	25	49	92	104	0.20	<LOD	<LOD	0.20	101.5	-	-	-
		3	25	50	90	102	0.58	<LOD	<LOD	0.60	102.0	-	-	-
		6	27	55	96	103	0.91	<LOD	<LOD	0.91	100.5	-	-	-

ND = Not Detected, LOD = Limit of Detection, LOQ = Limit of Quantification, TAMC = Total Aerobic Microbial Count, TYMC = Total Combined Yeast/Molds Count

FIGURE 20B
33 mg MPH capsules

Lot No.	Packaging Material / Stability Conditions	Time Point (Month)	% Cumulative Drug Released				Organic Impurities				Microbial Limit Tests		
			1 HR	3 HR	9 HR	24 HR	MPH Related Compound A (LOQ=0.13%)	Erythor Isomer (LOQ=0.08%)	Other Degradation Product (LOQ=0.13%)	Total Impurity	Assay (% L.C.)	TAMC (cfu/g)	TYMC (cfu/g)
D	Blister-PVC/PVDC 40g with aluminum bag 25°C/60%RH	0	24	51	86	99	< LOQ	ND	ND	< LOQ	100.5	140	<10
		3	28	53	86	102	0.14	ND	ND	0.14	102.8	-	-
		6	27	52	83	100	0.16	ND	ND	0.16	101.8	-	-
		9	26	54	86	98	0.20	ND	ND	0.20	102.3	-	-
		13	26	51	84	100	0.40	ND	ND	0.40	101.1	170	<10

ND = Not Detected, LOQ = Limit of Quantification, TAMC = Total Aerobic Microbial Count, TYMC = Total Combined Yeast/Molds Count

FIGURE 21A
44 mg MPH capsules

Lot No.	Packaging Material / Stability Conditions	Time Point (Month)	Water Content	% Cumulative Drug Released				Organic Impurities				Assay (%LC)	Microbial Limit Tests	
				1 HR	3 HR	9 HR	24 HR	MPH Related Compound A (LOQ=0.05%)	Erythor Isomer (LOQ=0.06%)	Degradation Product (LOQ=0.10%)	Total Impurity		TAMC (cfu/g)	TYMC (cfu/g)
E	120cc HDPE Bottle 25°C/60%RH	0	3.9	22	43	85	101	<LOD	ND	ND	<LOQ	101.0	<10	<10
		3	1.5	22	42	84	102	<LOD	ND	ND	<LOQ	100.4	-	-
		6	1.6	23	42	81	101	0.10	ND	ND	0.10	99.8	-	-
		9	2.0	22	43	84	97	0.15	ND	ND	0.15	100.2	-	-
		12	1.7	22	45	86	100	0.20	ND	ND	0.20	100.0	<10	<10
		18	2.9	22	43	83	99	0.35	ND	ND	0.35	97.9	-	-
		24	2.0	22	44	82	100	0.48	ND	ND	0.48	101.4	<10	<10
		0	3.9	22	43	85	101	<LOD	ND	ND	<LOQ	101.0	<10	<10
	120cc HDPE Bottle 30°C/75%RH	3	2.0	22	47	87	102	0.14	ND	ND	0.14	99.5	-	-
		6	2.2	23	46	86	99	0.24	ND	ND	0.24	99.8	-	-
		9	2.3	23	45	84	101	0.40	ND	ND	0.40	99.4	-	-
		12	2.1	23	46	88	102	0.56	ND	ND	0.56	99.7	<10	<10
	120cc HDPE Bottle 40°C/75%RH	18	2.7	23	44	82	101	0.90	ND	ND	0.90	98.0	-	-
		24	2.8	24	45	82	101	1.07	<LOD	ND	1.07	98.7	<10	<10
		0	3.85	22	43	85	101	<LOD	<LOD	<LOD	<LOQ	101.0	<10	<10
		1	1.40	25	50	90	102	0.19	<LOD	<LOD	0.20	100.1	-	-
		3	1.80	24	49	88	101	0.56	<LOD	<LOD	0.60	98.7	-	-
		6	2.30	24	47	86	101	0.89	<LOD	<LOD	0.89	98.7	-	-

ND = Not Detected, LOD = Limit of Detection, LOQ = Limit of Quantification, TAMC = Total Aerobic Microbial Count, TYMC = Total Combined Yeast/Molds Count

24/24

FIGURE 21B
44 mg MPH capsules

Lot No.	Packaging Material / Stability Conditions	Time Point (Month)	% Cumulative Drug Released				Organic Impurities				Microbial Limit Tests		
			1 HR	3 HR	9 HR	24 HR	MPH Related Compound A (LOQ=0.09%)	Erythor Isomer (LOQ=0.06%)	Other Degradation Product (LOQ=0.10%)	Total Impurity	Assay (% I.C)	TAMC	E. Coli
F	Blister-PVC/PVDC 40g with aluminum bag 25°C/60%RH	0	22	47	89	102	<LOQ	ND	ND	<LOQ	100.1	230	<10
		3	24	47	83	101	0.13	ND	ND	0.13	103.1	-	-
		6	23	48	90	104	0.16	ND	ND	0.16	102.9	-	-
		9	23	48	88	102	0.2	ND	ND	0.20	100.6	-	-
		13	23	46	83	103	0.2	ND	ND	0.20	102.2	210	<10

ND = Not Detected, LOQ = Limit of Quantification, TAMC = Total Aerobic Microbial Count, TYMC = Total Combined Yeast/Molds Count