

**Argatroban** **PROTOCOL FOR ISOLATED HIT, HIT & HITS IN PATIENTS WITH HEPATIC DYSFUNCTION (HEPATIC DISEASE SCORE GREATER THAN 6) NON CATH LAB**

Indication Anticoagulant for prophylaxis or treatment of thrombosis in patients with Isolated HIT, HIT, HITS (Heparin induced Thrombocytopenia/thrombosis syndrome)

Contraindications Overt major bleeding; argatroban should be avoided if LFT are greater than or equal to 3 times the upper limit of normal

Baseline Labs aPTT, PT/INR, Serum Creatinine, CMP, HCT, and platelets

Monitoring Platelets daily while on argatroban

Monitoring Guaiaac all stools, gastrocull all emesis, visually check for hematuria or other bleeding. CALL PHYSICIAN FOR BLEEDING

Notify Physician immediately for any of the following: unexplained drop in blood pressure, development of hematoma, drop in hematocrit, significant bleed, flank pain, bright red urine, or bruising

Discontinue All Heparin, Lovenox (enoxaparin), dalteparin (Fragmin)

Discontinue Discontinue warfarin until platelets are back to baseline or > 100,000 (preferably when count is 150,000)

Discontinue IM injections while on Argatroban, obtain physician order for alternative route

Antidote No antagonist is available to reverse the action of argatroban

Transition to Warfarin Start when platelet count > 100,000 (preferably when count is 150,000). No starting doses > 5 mg. Overlap with argatroban for at least 5 days.

INR Monitoring INR daily. If INR > 4 on combination therapy stop argatroban, repeat INR in 4-6 hours, if INR is less than therapeutic restart argatroban at previous rate. Repeat the procedure daily. Discontinue argatroban when two consecutive daily INRs are therapeutic on warfarin alone after therapy has overlapped for at least 5 days.

aPTT Monitoring Draw aPTT 2 hours and 6 hours after initiation of therapy and every 4 hours after any change in infusion rate until two consecutive aPTT are therapeutic then draw aPTT every 24 Hours  
If aPTT is greater than 100 seconds, stop argatroban, drawn aPTT every 2 hours until aPTT within therapeutic range, restart argatroban at new rate  
All aPTTs should be run stat

aPTT Goal aPTT 40-83 seconds (round actual aPTT to closest whole number)

Mixing instructions **Dosage adjustments based on aPTT see Argatroban Rates Change Based on aPTT Chart**  
Dilute each 2.5ml vial with 250ml of 0.9% Sodium Chloride Injection, 5% Dextrose Injection or Lactated Ringer's Injection to obtain a final concentration of 1mg/ml. The constituted solution must be mixed by repeated inversion of the diluent bag for 1 minute. Upon preparation, the solution may show slight, but brief haziness due to the formation of microprecipitates that rapidly dissolve upon mixing.

Drug Amount **250 mg**  
Volume **500 ml**  
Final Concentration **500 mcg/ml**

Starting Dose 0.5 mcg/kg/min (Hepatic Disease score greater than 6)  
Hepatic Disease Score and creatinine clearance calculators are available on the pharmacy web site under calculators  
If Liver Function Test are greater than 3 times upper limit of normal do not use argatroban

Dosage Range 0.5-10 mcg/kg/min  
Maximum Dose 10 mcg/kg/min

Patient Weight (kg)	Patient Weight (lbs)	INITIAL DOSE (mcg/kg/min)														
		0.1	0.2	0.3	0.4	0.5	1	2	3	4	5	6	7	8	9	10
40	88	0.5	1	1.4	1.9	2.4	4.8	10	14	19	24	29	34	38	43	48
45	99	0.5	1.1	1.6	2.2	2.7	5	11	16	22	27	32	38	43	49	54
50	110	0.6	1.2	1.8	2.4	3	6	12	18	24	30	36	42	48	54	60
55	121	0.7	1.3	2	2.6	3.3	7	13	20	26	33	40	46	53	59	66
60	132	0.7	1.4	2.2	2.9	3.6	7	14	22	29	36	43	50	58	65	72
65	143	0.8	1.6	2.3	3.1	3.9	8	16	23	31	39	47	55	62	70	78
70	154	0.8	1.7	2.5	3.4	4.2	8	17	25	34	42	50	59	67	76	84
75	165	0.9	1.8	2.7	3.6	4.5	9	18	27	36	45	54	63	72	81	90
80	176	1	1.9	2.9	3.8	4.8	10	19	29	38	48	58	67	77	86	96
85	187	1	2	3.1	4.1	5	10	20	31	41	51	61	71	82	92	102
90	198	1.1	2.2	3.2	4.3	5	11	22	32	43	54	65	76	86	97	108
95	209	1.1	2.3	3.4	4.6	6	11	23	34	46	57	68	80	91	103	114
100	220	1.2	2.4	3.6	4.8	6	12	24	36	48	60	72	84	96	108	120
105	231	1.3	2.5	3.8	5	6	13	25	38	50	63	76	88	101	113	126
110	242	1.3	2.6	4	5	7	13	26	40	53	66	79	92	106	119	132
115	253	1.4	2.8	4.1	6	7	14	28	41	55	69	83	97	110	124	138
120	264	1.4	2.9	4.3	6	7	14	29	43	58	72	86	101	115	130	144
125	275	1.5	3	4.5	6	8	15	30	45	60	75	90	105	120	135	150
130	286	1.6	3.1	4.7	6	8	16	31	47	62	78	94	109	125	140	156
135	297	1.6	3.2	4.9	6	8	16	32	49	65	81	97	113	130	146	162
140	308	1.7	3.4	5	7	8	17	34	50	67	84	101	118	134	151	168
145	319	1.7	3.5	5	7	9	17	35	52	70	87	104	122	139	157	174
150	330	1.8	3.6	5	7	9	18	36	54	72	90	108	126	144	162	180
155	341	1.9	3.7	6	7	9	19	37	56	74	93	112	130	149	167	186
160	352	1.9	3.8	6	8	10	19	38	58	77	96	115	134	154	173	192
165	363	2	4	6	8	10	20	40	59	79	99	119	139	158	178	198
170	374	2	4.1	6	8	10	20	41	61	82	102	122	143	163	184	204

RATE (ml/hr)

Marshall Pierce PharmD.

Infusion Rate Change Based on aPTT				
Please see Rate Change Chart for rate in ml/hr				
aPTT (Seconds)				
33 or less	34-39	40-83	84-96	97 or higher
50% Increase	25% Increase	No Change	25% Decrease	50% Decrease
Rate Change				
If aPTT is greater than 100 seconds, stop argatroban, drawn aPTT every 2 hours until aPTT is within therapeutic range, then restart argatroban at new rate				
Round the aPTT to the closest whole number				