

### 1.1.1 Biomarkers results of studies

Paper	Population	Biomarker	Reported timing of test (presentation or onset of symptoms)	Natural History of test (when does it peak)	N (Population)	Prevalance	Sensitivity	Specificity	Positive Likelihood ratio	Postiv e predic tive Value	Negative Likeliho od ratio	Negati ve predic tive Value
Balk	Acute Patients Emergency department	Troponin I (single test)	1 study at 2 hours after presentation, 1 at 7 hours after presentation, 5 studies not reported	Initial peak: 3-12 hours Mean time to peak elevation : 24 hours	1149	17%	39%	93%	5.57143	0.527625	0.655914	0.883784
Ebell	Acute Patients Emergency department	Troponin I (single test)	1 hour after onset of pain		1510		13%	95%	2.6	0	0.915789	1
Ebell	Acute Patients Emergency department	Troponin I (single test)	2 hours after onset of pain		1510		34%	95%	6.8	0	0.694737	1
Ebell	Acute Patients Emergency department	Troponin I (single test)	3 hours after onset of pain		1510		52%	95%	10.4	0	0.505263	1
Ebell	Acute Patients Emergency department	Troponin I (single test)	4 hours after onset of pain		1510		67%	95%	13.4	0	0.347368	1
Ebell	Acute Patients Emergency department	Troponin I (single test)	5 hours after onset of pain		1510		80%	95%	16	0	0.210526	1
Ebell	Acute Patients Emergency department	Troponin I (single test)	6 hours after onset of pain		1510		90%	95%	18	0	0.105263	1
Kost	Acute Patients Emergency department	Troponin I (single test)	Presentation and 3, 6 and 12 hours after presentation		97	28%	90%	100%	#DIV/O!	1	0.1	0.962567
Alp	Acute Patients CCU	Troponin I (single test)	6 hours after onset of pain		397	28%	76%	92%	9.5	0.786982	0.26087	0.907895

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Paper	Population	Biomarker	Reported timing of test (presentation or onset of symptoms)	Natural History of test (when does it peak)	N (Population)	Prevalence	Sensitivity	Specificity	Positive Likelihood ratio	Positive predictive Value	Negative Likelihood ratio	Negative predictive Value
Chui	Acute Patients Emergency department or CCU	Troponin I (single test)	0-4 hours after onset of pain		87	100%	35%		0.353	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	Troponin I (single test)	4-8 hours after onset of pain		87	100%	80%		0.8	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	Troponin I (single test)	8-12 hours after onset of pain		87	100%	93%		0.93	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	Troponin I (single test)	12-24 hours after onset of pain		87	100%	97%		0.97	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	Troponin I (single test)	24-48 hours after onset of pain		87	100%	96%		0.96	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	Troponin I (single test)	48-72 hours after onset of pain		87	100%	97%		0.97	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	Troponin I (single test)	>72 hours after onset of pain		87	100%	93%		0.93	1	#DIV/0!	0
Eggers	Acute Patients CCU	Troponin I (single test)	Presentation		176	22%	79%	95%	15.8	0.816729	0.221053	0.941311
Eggers	Acute Patients CCU	Troponin I (single test)	6 hours after presentation		180	22%	89%	95%	17.8	0.833901	0.115789	0.968374
Eggers	Acute Patients CCU	Troponin I (single test)	12 hours after presentation Every 6-8 hours after admission for 24-48 hours		172	22%	100%	95%	20	0.849421	0	1
Falahati	Acute Patients Hospital	Troponin I (single test)	24-48 hours		327	19%	100%	96%	25	0.854317	0	1
al Harbi	Acute Patients CCU	Troponin I (single test)	Presentation		124	59%	60%	82%	3.333333	0.827489	0.487805	0.587557
al Harbi	Acute Patients CCU	Troponin I (single test)	8 hours after presentation		124	59%	88%	72%	3.14286	0.818927	0.166667	0.806557

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al Harbi	Acute Patients CCU	Troponin I (single test)	16 hours after presentation		124	59%	93%	79%	4.42857	0.864367	0.088608	0.886911
Zimmerman	Acute Patients Hospital	Troponin I (single test)	2 hours after onset of pain		955	100%	16%	97%	5.33333	1	0.865979	0
Zimmerman	Acute Patients Hospital	Troponin I (single test)	4 hours after onset of pain		955	100%	36%	94%	6	1	0.680851	0
Zimmerman	Acute Patients Hospital	Troponin I (single test)	6 hours after onset of pain		955	100%	58%	94%	9.66667	1	0.446809	0
Zimmerman	Acute Patients Hospital	Troponin I (single test)	10 hours after onset of pain		955	100%	92%	95%	18.4	1	0.084211	0
Zimmerman	Acute Patients Hospital	Troponin I (single test)	14 hours after onset of pain		955	100%	91%	92%	11.375	1	0.097826	0
Zimmerman	Acute Patients Hospital	Troponin I (single test)	18 hours after onset of pain		955	100%	96%	93%	13.7143	1	0.043011	0
Zimmerman	Acute Patients Hospital	Troponin I (single test)	22 hours after onset of pain		955	100%	90%	94%	15	1	0.106383	0
Balk	Acute Patients Emergency department	Troponin T (single test)	Presentation	Initial peak: 3-12 hours	1348	34%	39%	93%	5.57143	0.741611	0.655914	0.747443
Ebell	Acute Patients Emergency department	Troponin T >0.1 ng/ml (single test)	1 hour after onset of pain	Mean time to peak elevation : 12-48 hours	4362		47%	87%	3.61538	0	0.609195	1

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Ebell	Acute Patients Emergency department	Troponin T >0.1 ng/ml (single test)	2 hours after onset of pain		4362		53%	87%	4.07692	0	0.54023	1
Ebell	Acute Patients Emergency department	Troponin T >0.1 ng/ml (single test)	3 hours after onset of pain		4362		58%	86%	4.14286	0	0.488372	1
Ebell	Acute Patients Emergency department	Troponin T >0.1 ng/ml (single test)	4 hours after onset of pain		4362		64%	85%	4.26667	0	0.423529	1
Ebell	Acute Patients Emergency department	Troponin T >0.1 ng/ml (single test)	6 hours after onset of pain		4362		74%	83%	4.35294	0	0.313253	1
Ebell	Acute Patients Emergency department	Troponin T >0.1 ng/ml (single test)	8 hours after onset of pain		4362		84%	81%	4.42105	0	0.197531	1
Ebell	Acute Patients Emergency department	Troponin T >0.1 ng/ml (single test)	10 hours after onset of pain		4362		93%	80%	4.65	0	0.0875	1
Ebell	Acute Patients Emergency department	Troponin T >0.2 ng/ml (single test)	1 hour after onset of pain		4362		14%	87%	1.07692	0	0.988506	1
Ebell	Acute Patients Emergency department	Troponin T >0.2 ng/ml (single test)	2 hours after onset of pain		4362		33%	87%	2.53846	0	0.770115	1
Ebell	Acute Patients Emergency department	Troponin T >0.2 ng/ml (single test)	3 hours after onset of pain		4362		50%	86%	3.57143	0	0.581395	1
Ebell	Acute Patients Emergency department	Troponin T >0.2 ng/ml (single test)	4 hours after onset of pain		4362		65%	85%	4.33333	0	0.411765	1
Ebell	Acute Patients Emergency department	Troponin T >0.2 ng/ml (single test)	6 hours after onset of pain		4362		86%	83%	5.05882	0	0.168675	1
Ebell	Acute Patients Emergency department	Troponin T >0.2 ng/ml (single test)	8 hours after onset of pain		4362		96%	81%	5.05263	0	0.049383	1

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Ebell	Acute Patients Emergency department	Troponin T >0.2 ng/ml (single test)	10 hours after onset of pain		4362		96%	80%	4.8	0	0.05	1
Guo	Acute Patients Cardiac department or CCU	Troponin T (single test)	Presentation, 6 and 12 hours after presentation		502	89%	95%	94%	15.8667	0.992271	0.051064	0.707638
Kost	Acute Patients Emergency department	Troponin T (single test)	Presentation and 3, 6 and 12 hours after presentation		97	28%	89%	90%	8.9	0.775841	0.122222	0.954626
Gust	Acute patients Out-of-hospital (EMS)	Troponin T (single test)	Not reported		68	24%	25%	98%	12.5	0.797872	0.765306	0.805363
Planer	Acute patients Out-of-hospital (community)	Troponin T (single test)	Not reported		349	2%	83%	100%	#DIV/0!	1	0.17	0.997069
Zarich	Acute Patients Emergency department	Troponin T (single test)	Presentation		267	32%	87%	94%	14.5	0.87218	0.138298	0.938895
Vatansever	Acute Patients Emergency department	Troponin T (single test)	Presentation		60	55%	76%	90%	7.6	0.902808	0.266667	0.75419
Vatansever	Acute Patients Emergency department	Troponin T (single test)	2 hours after presentation		60	55%	97%	90%	9.7	0.922213	0.033333	0.960854
Zimmerman	Acute Patients Hospital	Troponin T (single test)	2 hours after onset of pain		955	100%	11%	98%	5.5	1	0.908163	0
Zimmerman	Acute Patients Hospital	Troponin T (single test)	4 hours after onset of pain		955	100%	36%	98%	18	1	0.653061	0
Zimmerman	Acute Patients Hospital	Troponin T (single test)	6 hours after onset of pain		955	100%	62%	96%	15.5	1	0.395833	0
Zimmerman	Acute Patients Hospital	Troponin T (single test)	10 hours after onset of pain		955	100%	87%	96%	21.75	1	0.135417	0
Zimmerman	Acute Patients Hospital	Troponin T (single test)	14 hours after onset of pain		955	100%	85%	96%	21.25	1	0.15625	0

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Zimmerman	Acute Patients Hospital	Troponin T (single test)	18 hours after onset of pain		955	100%	79%	96%	19.75	1	0.21875	0
Zimmerman	Acute Patients Hospital	Troponin T (single test)	22 hours after onset of pain		955	100%	86%	95%	17.2	1	0.147368	0
Balk	Acute Patients Emergency department	CK (single test)	Presentation Every 6-8 hours after admission for 24-48 hours		3195	25%	37%	87%	2.84615	0.481485	0.724138	0.808893
Falahati	Acute Patients Hospital	CK (single test)			327	19%	74%	85%	4.9	0.53475	0.311765	0.931854
Vatansever	Acute Patients Emergency department	CK (single test)	Presentation		60	55%	64%	90%				
Vatansever	Acute Patients Emergency department	CK (single test)	2 hours after presentation		60	55%	79%	90%	7.9	0.906152	0.233333	0.778098
Balk	Acute Patients Emergency department	CK-MB (single test)	Presentation	Initial peak: 3-12 hours Mean time to peak elevation : 24 hours	6425	25%	42%	97%	14	0.823529	0.597938	0.833811
Kost	Acute Patients Emergency department	CK-MB (single test)	Presentation and 3, 6 and 12 hours after presentation		97	28%	89%	90%	8.9	0.775841	0.122222	0.954626
Chui	Acute Patients Emergency department or CCU	CK-MB (single test)	0-4 hours after onset of pain		87	100%	44%		0.441	1	#DIV/0!	0
Chui	Acute Patients Emergency	CK-MB (single test)	4-8 hours after onset of pain		87	100%	96%		0.96	1	#DIV/0!	0

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	department or CCU											
Chui	Acute Patients Emergency department or CCU	CK-MB (single test)	8-12 hours after onset of pain		87	100%	98%		0.98	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	CK-MB (single test)	12-24 hours after onset of pain		87	100%	97%		0.97	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	CK-MB (single test)	24-48 hours after onset of pain		87	100%	93%		0.93	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	CK-MB (single test)	48-72 hours after onset of pain		87	100%	71%		0.71	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	CK-MB (single test)	>72 hours after onset of pain		87	100%	23%		0.23	1	#DIV/0!	0
Eggers	Acute Patients CCU	CK-MB (single test)	Presentation		176	22%	66%	95%	13.2	0.788274	0.357895	0.908311
Eggers	Acute Patients CCU	CK-MB (single test)	6 hours after presentation		180	22%	81%	95%	16.2	0.820442	0.2	0.946602
Eggers	Acute Patients CCU	CK-MB (single test)	12 hours after presentation		172	22%	77%	95%	15.4	0.81286	0.242105	0.936079
Falahati	Acute Patients Hospital	CK-MB (single test)	Every 6-8 hours after admission for 24-48 hours		327	19%	88%	93%	12.5714	0.746762	0.129032	0.970622
Fesmire	Acute Patients Hospital	CK-MB (single test)	2 hours after presentation		975	5%	91%	78%	4.13636	0.163115	0.115385	0.994592
Zarich	Acute Patients Emergency department	CK-MB (single test)	Presentation		267	32%	47%	83%	2.76471	0.565414	0.638554	0.768937
Zimmerman	Acute Patients Hospital	CK-MB (single test)	2 hours after onset of pain		955	100%	21%	91%	2.33333	1	0.868132	0

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Paper	Population	Biomarker	Reported timing of test (presentation or onset of symptoms)	Natural History of test (when does it peak)	N (Population)	Prevalence	Sensitivity	Specificity	Positive Likelihood ratio	Positive predictive Value	Negative Likelihood ratio	Negative predictive Value
Zimmerman	Acute Patients Hospital	CK-MB (single test)	4 hours after onset of pain		955	100%	46%	89%	4.18182	1	0.606742	0
Zimmerman	Acute Patients Hospital	CK-MB (single test)	6 hours after onset of pain		955	100%	92%	89%	8.36364	1	0.089888	0
Zimmerman	Acute Patients Hospital	CK-MB (single test)	10 hours after onset of pain		955	100%	96%	90%	9.6	1	0.044444	0
Zimmerman	Acute Patients Hospital	CK-MB (single test)	14 hours after onset of pain		955	100%	91%	90%	9.1	1	0.1	0
Zimmerman	Acute Patients Hospital	CK-MB (single test)	18 hours after onset of pain		955	100%	81%	90%	8.1	1	0.211111	0
Zimmerman	Acute Patients Hospital	CK-MB (single test)	22 hours after onset of pain		955	100%	53%	92%	6.625	1	0.51087	0

Balk	Acute Patients Emergency department	Myoglobin (single test)	Presentation	Initial peak: 1-4 hours Mean time to peak elevation : 6-7 hours	4172	28%	49%	91%	5.44444	0.679208	0.56044	0.821053
Kost	Acute Patients Emergency department	Myoglobin (single test)	Presentation and 3, 6 and 12 hours after presentation		97	28%	80%	75%	3.2	0.554455	0.266667	0.90604
Chui	Acute Patients Emergency department or CCU	Myoglobin (single test)	0-4 hours after onset of pain		87	100%	56%		0.558	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	Myoglobin (single test)	4-8 hours after onset of pain		87	100%	92%		0.92	1	#DIV/0!	0



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Paper	Population	Biomarker	Reported timing of test (presentation or onset of symptoms)	Natural History of test (when does it peak)	N (Population)	Prevalence	Sensitivity	Specificity	Positive Likelihood ratio	Positive predictive Value	Negative Likelihood ratio	Negative predictive Value
Chui	Acute Patients Emergency department or CCU	Myoglobin (single test)	8-12 hours after onset of pain		87	100%	85%		0.85	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	Myoglobin (single test)	12-24 hours after onset of pain		87	100%	75%		0.75	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	Myoglobin (single test)	24-48 hours after onset of pain		87	100%	43%		0.43	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	Myoglobin (single test)	48-72 hours after onset of pain		87	100%	20%		0.2	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	Myoglobin (single test)	>72 hours after onset of pain		87	100%	14%		0.14	1	#DIV/0!	0
Eggers	Acute Patients CCU	Myoglobin (single test)	Presentation		176	22%	63%	95%	12.6	0.780405	0.389474	0.901021
Eggers	Acute Patients CCU	Myoglobin (single test)	6 hours after presentation		180	22%	43%	95%	8.6	0.708084	0.6	0.855263
Fesmire	Acute Patients Hospital	Myoglobin (single test)	2 hours after presentation		975	5%	75%	80%	3.75	0.150167	0.3125	0.985489
Vatansever	Acute Patients Emergency department	Myoglobin (single test)	Presentation		60	55%	85%	90%	8.5	0.912195	0.166667	0.830769
Vatansever	Acute Patients Emergency department	Myoglobin (single test)	2 hours after presentation		60	55%	97%	90%	9.7	0.922213	0.033333	0.960854
Zimmerman	Acute Patients Hospital	Myoglobin (single test)	2 hours after onset of pain		955	100%	26%	87%	2	1	0.850575	0
Zimmerman	Acute Patients Hospital	Myoglobin (single test)	4 hours after onset of pain		955	100%	42%	89%	3.81818	1	0.651685	0
Zimmerman	Acute Patients Hospital	Myoglobin (single test)	6 hours after onset of pain		955	100%	79%	89%	7.18182	1	0.235955	0

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Paper	Population	Biomarker	Reported timing of test (presentation or onset of symptoms)	Natural History of test (when does it peak)	N (Population)	Prevalence	Sensitivity	Specificity	Positive Likelihood ratio	Positive predictive Value	Negative Likelihood ratio	Negative predictive Value
Zimmerman	Acute Patients Hospital	Myoglobin (single test)	10 hours after onset of pain		955	100%	87%	90%	8.7	1	0.144444	0
Zimmerman	Acute Patients Hospital	Myoglobin (single test)	14 hours after onset of pain		955	100%	62%	88%	5.16667	1	0.431818	0
Zimmerman	Acute Patients Hospital	Myoglobin (single test)	18 hours after onset of pain		955	100%	58%	89%	5.27273	1	0.47191	0
Zimmerman	Acute Patients Hospital	Myoglobin (single test)	22 hours after onset of pain		955	100%	43%	91%	4.77778	1	0.626374	0
Balk	Acute Patients Emergency department	Troponin I (serial)	Not reported		1393	8%	95%	90%	9.5	0.43515	0.055556	0.995516
Balk	Acute Patients Emergency department	Troponin T (serial)	Not reported		904	28%	93%	85%	6.2	0.70684	0.082353	0.968968
Balk	Acute Patients Emergency department	CK (serial)	Not reported		786	35%	83%	76%	3.4375	0.649241	0.230263	0.889689
Balk	Acute Patients Emergency department	CK-MB (serial)	Not reported		11625	18%	79%	96%	19.75	0.812571	0.21875	0.954182
Zarich	Acute Patients Emergency department	CK-MB (serial)	Presentation		267	32%	95%	87%	7.30769	0.77472	0.057471	0.973667
Balk	Acute Patients Emergency department	Myoglobin (serial)	Not reported		1277	26%	89%	87%	6.84615	0.706349	0.126437	0.957466

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Paper	Population	Biomarker	Reported timing of test (presentation or onset of symptoms)	Natural History of test (when does it peak)	N (Population)	Prevalence	Sensitivity	Specificity	Positive Likelihood ratio	Positive predictive Value	Negative Likelihood ratio	Negative predictive Value
Balk	Acute Patients Emergency department	CK-MB and myoglobin (single)	2 studies at presentation, 1 study 2 hours after presentation		2283	29%	83%	82%	4.61111	0.647639	0.207317	0.923671
Balk	Acute Patients Emergency department	CK-MB and myoglobin (serial)	Not reported		291	17%	100%	83%	5.88235	0.546448	0	1
Chui	Acute Patients Emergency department or CCU	Triple test (single test) (troponin I, CK-MB, myoglobin)	0-4 hours after onset of pain		87	100%	62%		0.62	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	Triple test (single test) (troponin I, CK-MB, myoglobin)	4-8 hours after onset of pain		87	100%	96%		0.96	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	Triple test (single test) (troponin I, CK-MB, myoglobin)	8-12 hours after onset of pain		87	100%	98%		0.98	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	Triple test (single test) (troponin I, CK-MB, myoglobin)	12-24 hours after onset of pain		87	100%	97%		0.97	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	Triple test (single test) (troponin I, CK-MB, myoglobin)	24-48 hours after onset of pain		87	100%	99%		0.99	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	Triple test (single test) (troponin I, CK-MB, myoglobin)	48-72 hours after onset of pain		87	100%	99%		0.99	1	#DIV/0!	0
Chui	Acute Patients Emergency department or CCU	Triple test (single test) (troponin I, CK-MB, myoglobin)	>72 hours after onset of pain		87	100%	95%		0.95	1	#DIV/0!	0

### 1.1.2 Summary table showing highest sensitivities and specificities of each biomarker evaluated in the study

#### Single testing

Paper	Troponin I		Troponin T		CK-MB		CK		Myoglobin		CK-MB & Myoglobin		Triple test	
	Sensitivity	Specificity	Sensitivity	Specificity	Sensitivity	Specificity	Sensitivity	Specificity	Sensitivity	Specificity	Sensitivity	Specificity	Sensitivity	Specificity
Balk (time if given)	39%	93%	39%	93%	42%	97%	37%	87%	49%	91%	83%	82%		
Ebell (time if given)	90%	95% 6 hours	93%	80% 10 hours										
Guo (time if given)			95%	94%										
Kost (time if given)	90%	100%	89%	90%	89%	90%			80%	75%				
Alp (time if given)	76%	92% 6 hours												
Chiu (time if given)	97%	12-24 hours			98%	8-12 hours			92%	4-8 hours			99%	72 hours
Eggers (time if given)	100%	95% 12 hours			81%	95% 6 hours			63%	95% 0 hours				
Falahati (time if given)	100%	96%			88%	93%	74%	85%						
Fesmire (time if given)					91%	78% 2 hours			75%	80% 2 hours				
Gust (out of hospital) (time if given)			25%	98%										
Planer (out of hospital) (time if given)			83%	100%										

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Zarich (time if given)			87%	94%	47%	83%				
				0 hours		0 hours				
al Harbi (time if given)	93%	79%								
		16 hours								
Vatansever (time if given)			97%	90%			79%	90%	97%	90%
				2 hours				2 hours		2 hours
Zimmerman (time if given)	96%	93%	87%	96%	96%	90%			87%	90%
		18 hours		10 hours		10 hours				10 hours

Serial testing

Paper	Troponin I Sensitivity Specificity	Troponin T Sensitivity Specificity	CK-MB Sensitivity Specificity	CK Sensitivity Specificity	Myoglobin Sensitivity Specificity	CK-MB & Myoglobin Sensitivity Specificity
Balk (time if given)	95% 90%	93% 85%	79% 96%	83% 76%	89% 87%	100% 83%
Zarich (time if given)			95% 87%			