Appendices

Appendix - Table 1. NTS urgency levels and response times

NTS Urgency level	Definition	Response time	Medical help	
U0 – Resuscitation	Loss of vital functions	Immediately	Ambulance	
U1 – Life	Unstable vital functions	Immediately, within 15	Ambulance	
threatening		minutes		
U2 – Emergent	Vital functions in danger	As soon as possible,	Home visit by GP or	
	or organ damage	within 1 hour	appointment at OHS-PC	
U3 – Urgent	Possible risk of damage,	A few hours (<3 hours)	Home visit by GP or	
	human reasons		appointment at OHS-PC	
U4 – Non-urgent	Marginal risk of damage	24 hours	Appointment at OHS-PC or	
			telephone advice	
U5 – Advice	No risk of damage	Advice, no time related	Telephone advice	
GP: general practitioner	r			
NTS: Netherlands Triage Standard				
OHS-PC: out-of-hours services in primary care				

OHS-PC: out-of-hours services in primary care

Appendix-table 2: Combinations of NTS triage criteria that generate an U1 level within the NTS main complaints that can be used for patients with chest discomfort.

Main compla	int 'Chest pain'	
→ ABCD stable	AND severe chest	pain (Numeric Rating Scale (NRS)-score ≥ 8) lasting less than 12 hours
→ ABCD	AND mild (NRS ≤	AND one of the following:
stable	4) to moderate	- retrosternal located pain
	(NRS 5-7) chest	- tightening or pressing
	pain lasting for	- radiation to jaw, arm or upper back
	less than 12	- progressive pain intensity in short time
	hours	- past or present autonomous nervous system-related symptoms
		- dizziness
Main compla	int 'Collapse'	
→ ABCD	AND collapse	AND chest pain of any severity
stable		
Main compla	int 'Back complaints'	
→ ABCD	AND severe	AND past or present autonomous nervous system-related
stable	upper back pain	symptoms
	(NRS ≥8)	

ABCD: acronym for Airway, Breathing, Circulation and Disability. When the triage nurse starts the telephone triage with the NTS, the system requires a mandatory 'ABCD-check'; i.e. the triage nurse has to ask questions to assess whether the patient has life-threatening problems concerning the Airway, Breathing, Circulation and Disability for which an ambulance should be sent immediately.

Appendix-table 3. Characteristics of patients divided among the seven OHS-PC locations.

Characteristics	Location A	Location B	Location C	Location D	Location E	Location F	Location G
	N=205	N=355	N=544	N=262	N=164	N=412	N=250
	(9.4%)	(16.2%)	(24.8%)	(12.0%)	(7.5%)	(18.8%)	(11.4%)
Prevalence of ACS (n,%)	32 (15.4%)	31 (8.7%)	59 (10.8%)	35 (13.4%)	15 (9.1%)	53 (12.9%)	26 (10.4%)
Male sex (n,%)	77 (37.6%)	154 (43.4%)	256 (47.1%)	108 (41.2%)	84 (51.2%)	188 (45.6%)	112 (44.8%)
Mean age in years (SD)	62.3 (19.6)	58.6 (19.8)	58.0 (19.4)	56.6 (18.9)	61.8 (20.5)	61.6 (19.4)	56.1 (18.7)

Appendix – table 4. Overview of the percentages of missing predictors, divided into patients with and without the diagnosis ACS.

Characteristics	ACS, n=251	No ACS, n=1941
N=2192	(11.5%)	(88.5%)
	Missing (%)	Missing (%)
Mean in years age (SD)	0	0
Female sex	0	0
Median call duration in min (IQR)	0	0
Mean patient's introduction in min (IQR)	0	0
Triage nurse consulted the GP	0	0
Someone else called on behalf of patient	0	0
The person who calls expressed concerns	144 (57.3)	943 (48.6)
Cardiovascular disease or risk factor combined	30 (12.0)	318 (16.4)
History of coronary artery disease	107 (42.6)	933 (48.1)
Diabetes	162 (64.5)	1126 (58.0)
Hypertension	172 (68.5)	1126 (58.0)
Hypercholesterolemia/statin use	172 (68.5)	1196 (61.6)
Cardiac arrhythmia	176 (70.1)	1110 (57.2)
Chest pain	4 (1.6)	72 (3.7)

Shortness of breath	68 (27.1)	429 (22.1)
Chest pain duration <12 hours	28 (11.2)	242 (13.1)
Pain intensity severe (NRS >7 in range 1-10)	149 (59.4)	1124 (57.9)
Pressing heavy pain*	63 (25.1)	502 (25.9)
Stabbing chest pain*	63 (25.1)	502 (25.9)
Chest pain located retrosternal**	87 (34.7)	538 (27.7)
Chest pain located left or right on thorax**	87 (34.7)	538 (27.7)
Radiation of chest pain to any location	38 (15.1)	476 (24.5)
Radiation to the arm ***	94 (37.5)	1019 (52.5)
Radiation to the shoulder blades ***	94 (37.5)	1019 (52.5)
Radiation to the jaws ***	94 (37.5)	1019 (52.5)
Sweating	79 (31.5)	747 (38.5)
Nausea or vomiting	133 (53.0)	1071 (55.2)
Pallor or ashen skin	158 (62.9)	1361 (70.1)
(Near) fainting	33 (13.1)	217 (11.2)
Palpitations	225 (89.6)	1615 (83.2)
Patient recognizes symptoms from previous cardiac	139 (55.4)	1137 (58.6)
event		

^{*}Pain described by patient. Pressing heavy pain: pressing, heavy or tightening pain vs. other types of pain (stabbing, burning, cramping, tearing) Stabbing pain: stabbing vs. other types of pain (pressing, heavy, tightening, burning, cramping)

^{**} Retrosternal location vs. other pain locations. Left or right side thorax vs. other pain locations

^{***} Radiation location vs. no radiation and radiation other pain

Appendix-table 5. Patient and call characteristics of 3,204 patients with chest discomfort calling OHS-PC between 2014-2017, comparing patients with and without information on the study outcome.

Characteristics	Follow-up	No follow-up	P-value	
N=3,204	n= 2192 (68.4%)	n= 1012 (31.6%)		
Patient characteristics				
Median age in years (IQR) (n=3,204)	59.1 (19.5)	57.3 (20.4)	0.020	
Female sex (n=3,204)	1213 (55.3)	565 (55.8)	0.794	
Call characteristics			<u> </u>	
Median total call duration in min (IQR) (n=3,204)	6:51 (4:59-9:23)	6:56 (5:04-9:15)	0.836	
Mean patient's introduction in min (IQR) (n=3,204)	0:17 (0:11-0:25)	0:17 (0:11-0:26)	0.052	
Triage nurse consulted the GP (n=3,204)	1147 (52.3)	519 (51.3)	0.583	
Someone else called on behalf of patient (n=3,204)	1114 (50.8)	479 (47.3)	0.066	
Person who calls expressed concerns (n=1,478)	988 (89.7)	430 (90.1)	0.804	
Medical history and risk factors			l	
Cardiovascular disease or CV risk factor (n=2,672)	1192 (64.6)	515 (62.3)	0.254	
History of coronary artery disease (n=1,663)	389 (33.8)	166 (32.4)	0.573	
Diabetes (n=1,283)	180 (19.9)	68 (18.0)	0.432	
Hypertension (n=1,274)	323 (36.1)	121 (31.9)	0.150	
Hypercholesterolemia/statin use (n=1,176)	212 (25.7)	88 (25.1)	0.842	
Cardiac arrhythmia (n=1,326)	230 (25.4)	102 (24.3)	0.684	
Symptoms				
Chest pain (n=3,079)	1981 (93.6)	894 (92.8)	0.417	
Shortness of breath (n=2,505)	1094 (64.5)	520 (64.3)	0.911	
Chest pain duration <12 hours (n=2,793)	1403 (73.2)	610 (69.6)	0.052	
Severe pain (NRS >7 in range 1-10) (n=1,351)	337 (36.6)	185 (43.0)	0.024	
Pressing, heavy pain* (n=2,347)	1023 (62.9)	444 (61.6)	0.538	
Stabbing chest pain* (n=2,349)	366 (22.5)	177 (24.5)	0.280	
Chest pain located retrosternal** (n=2,298)	641 (40.9)	294 (40.2)	0.736	
Chest pain located left or right on thorax** (n=2,299)	627 (40.0)	294 (40.2)	0.945	
Radiation of chest pain to any location (n=2,437)	1077 (64.3)	458 (60.1)	0.047	
Radiation to the arm ***(n=1,521)	452 (42.2)	179 (39.8)	0.373	

Radiation to the shoulder blades *** (n=1,519)	326 (30.5)	136 (30.2)	0.924
Radiation to the jaws *** (n=1,984)	124 (11.6)	41 (9.1)	0.156
Sweating (n=1,758)	559 (40.9)	259 (42.0)	0.638
Nausea or vomiting (n=1,474)	489 (49.5)	229 (47.1)	0.381
Pallor or ashen skin (n=1,007)	322 (47.8)	136 (40.8)	0.038
(Near) fainting (n=2,855)	143 (7.4)	72 (7.8)	0.678
Palpitations (n=501)	284 (80.7)	125 (83.9)	0.396
Patient recognizes symptoms from previous cardiac	250 (27.3)	102 (26.7)	0.819
event (n=1,298)			
Urgency allocation			
High urgency (U1 or U2) (n=3,204)	1496 (68.2)	661 (65.3)	0.100
U1	974 (44.5)	390 (38.6)	
U2	522 (23.8)	271 (26.8)	
Low urgency (U3 or U4 or U5)	696 (31.8)	351 (34.7)	

Appendix - Table 6. Base model with age and sex for predicting the diagnosis ACS.

Predictors	Regression coefficients (standard error)
Intercept	- 14.671 (3.442)
Age	0.289 (0.079)
Age'	- 0.379 (0.123)
Age"	1.017 (0.386)
Female sex	2.155 (5.385)
Age * Female sex	- 0.084 (0.123)
Age' * Female sex	0.175 (0.190)
Age" * Female sex	- 0.532 (0.589)

Apparent c-statistic 0.72 (95% CI 0.70-0.75)

Adjusted c-statistic 0.72 (95% CI 0.68-0.75)

Calibration slope 0.977 (95% CI 0.617-1.338)

Calibration 0.016 (95% CI-0.702-0.734)

 $R^2 0.065$

Knots for cubic spline functions placed at 5, 35, 65 and 95

Appendix - Table 7. Full model including all candidate predictors for predicting the diagnosis ACS

Predictors	Regression coefficients (standard error)			
Intercept	-15.914 (3.55)			
Age	0.288 (0.081)			
Age'	-0.388 (0.126)			
Age"	1.058 (0.396)			
Female gender	2.459 (5.519)			
Age * Female sex	-0.094 (0.126)			
Age' * Female sex	0.187 (0.195)			
Age" * Female sex	-0.554 (0.606)			
Chest pain	-0.064 (0.365)			
Acute chest pain (< 12 hours)	0.258 (0.200)			
Shortness of breath	-0.141 (0.200)			
Sweating	0.459 (0.183)			
Retrosternal located pain	0.178 (0.177)			
Radiation of chest pain	0.617 (0.180)			
Pressing heavy feeling	0.619 (0.272)			
Stabbing pain	-0.200 (0.353)			
History of cardiovascular disease*	-0.039 (0.247)			
History of coronary artery disease	0.108 (0.234)			
Someone else calls instead of the patient	0.197 (0.160)			
Patient calls during the night (0am-9am)	0.495 (0.152)			
Apparent c-statistic 0.79 (95% CI 0.76-0.81)				
Adjusted c-statistic 0.77 (95% CI 0.74-0.80)				
Calibration slope 0.818 (95% CI 0.650-0.986)				
Calibration -0.238 (-0.621-0.145)				
R ² 0.107				
Knots for cubic spline functions placed at 5, 35, 65 and 95				

Appendix – Figure 1. Callibration of full model with internal external validation

