Supplemental Online Content

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eAppendix. ICD Codes Defining Sepsis, Infection, and Organ Dysfunction **eTable 1.** SOFA Score Adapted for Use Outside of Intensive Care **eTable 2.** The Linder-Mellhammar Criteria of Infection

This supplemental material has been provided by the authors to give readers additional information about their work.

eAppendix. ICD Codes Defining Sepsis, Infection, and Organ Dysfunction

		ICD-1
Sepsis	Explicit sepsis codes	A02.1
		A20.0
		A20.7
		A21.7
		A22.7
		A24.1
		A26.7
		A28.2
		A32.7
		A39.1
		A39.2
		A39.3
		A39.4
		A40.
		A41.
		A42.7
		A48.3
		A49.9
		A54.8
		B00.7
		B37.6
		B37.7
		B49
		075.3
		O85
		R65.0
		R65.1
		R57.2
	Severe sepsis	R65.1
	Septic shock	R57.2
Infections		J01
		J02
		J03
		J04
		J06
		J05
		J09
		J10
		J11
		J12
		J13
		J14

J15

J16

J17

J18

J20

J21

J22

J44.0

J44.1

J86

J85

...

A15

A16

U69.00

A36

A37

B38

B39

P23

P24.0

P24.8

P24.9

. _ ...

A00

A01

A02

A03

A04

A05

A06

A07

A08

, ...

A09

K35

K37

K36 K5702

K5703

K5712

K57.13

1137.13

K57.22

K57.23

K57.32

K5733

K5742

K5743

K5752

K5753

K5782

K5783

K5792

K5793

K61

K65

K67

K63.0

K63.1

K75.0

.., 5.0

K75.1

K81.0

K77.0

U69.40!

P38

P77

P78.1

A46

B47

L03

L04

L08

L05

B00

B07

B08

DOO

B09

H05.0

H60.2

H70.0

J36

J39.0

J39.1

L02

N10

N15.1

N15.9

N34

N30

N39.0

N41

N45

N48.2

N49

N70

N71

N72

N73

N74*

N75

N76

N77

N61

N98.0

A59

A55

A56

A39

G00

G01

G02

G03

G04

G05

G06

G07

G08

A17

A81

A83

A84

A85

A86

A87

A88

A89

132

133

139

140

141

180

138

198.1

T82.6

T82.7

T83.5

T83.6

T84.5

T84.6

T84.7

T85.7

A18 A19

A20

A21

A22

A23

A24

A25

A26

A27

A28

A32

A38

A42

A43

A44

A48

, , , ,

A49

A54

A69.0

A69.1

A69.2

A69.8

A69.9

A90

A91

A97

B37

B40

B41

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B42

B43

B44

B45

B46

B48

B49

B50

B51

B52

B53

B54

M00

M01

14101

M86

T80.2

T81.4

T88.0

A50

A65

A74

A75

A77

A78 A79

A80

A92

A93

A94

A95

A96

A98

A99

B01

B02

B03

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P35

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P39

075.3

085

0030

0035

O040

0045

0050

0055

0060

0065

0070

0075

0.800

086

023

041.1

088.3

098

091

Organ dysfunction

195.9 R57.8 R57.9 R57.2 J96. J80 J98.4 R06.0 R06.8 F05 G93.1 G93.4 R40 N17. N19 E87.2 D65 D68.8 D68.9 D69.5 D69.6 K72.0 K72.7 K72.9 K76.2 K76.3

eTable 1. SOFA Score Adapted for Use Outside of Intensive Care

SOFA score

Organ system	1	2	3	4
Respiration PaO ₂ /FIO ₂ mmHg	< 400	< 300	<200	<100
corresponding SpO ₂ without O ₂ support	<95%	<91%	with respira	tory support
Coagulation Platelets x 10^9/L	< 150	< 100	< 50	< 20
Liver Bilirubin µmol/L	>19	>32	>101	>204
Cardiovascular Hypotension	MAP < 70mmHg	Dopamine ≤ 5 μg/kg/min or dobutamine	Dopamine > 5 or epinephrine or norepinephrine ≤ 0,1 µg/kg/min	Dopamine > 15 or epinephrine or norepinephrine >0.1 µg/kg/min
		for at least one hour		
Central Nervous System				
Glascow Coma Scale	13-14	10-12	6-9	3-5
Renal Creatinine μmol/L	>110	>170	>300	>440
or urine output mL/day			< 500	< 200

eTable 2. The Linder-Mellhammar Criteria of Infection

For each focus, 1-4 points are given for the following entities: symptoms and signs, radiological findings, laboratory findings, and microbiologic findings in relation to the suspected infection. These entities are grouped together as rows in a table for each infection focus. The highest scores from all rows are added together to give a total score for each focus. The level of evidence of infection is graded according to the scores:

0-1 points, no infection

2 points, possible infection

3 points, probable infection,

4 or more points, proven infection

Table 1. LOWER RESPIRATORY TRACT INFECTION

Clinical entity	1	2	3	4
Symptoms	 Localized symptoms Purulent sputum or change in character of sputum New onset of cough or worsening of cough Pleural pain Generalized symptoms Fever >38°C or history of fever Rigors Leukocytosis or leukopenia 			
Signs	Possible fine crackles without other likely cause	Distinct crackles or bronchial breath sounds		
Radiological findings	Possible pneumonic infiltrate		Probable pneumonic infiltrate	Cavitation Abscess Empyema
Microbiological findings	 Detection of viral pathogens in respiratory tract Detection of pneumococciantigen in urine 	 Growth of likely pathogen from respiratory secretions Pneumocysti s detected in immunocom promised patient Detection of legionella-antigen in urine 	 Detection of Chlamydophila, Mycoplasma, Chlamydia, Legionella antigen, DNA or RNA in respiratory tract Growth of pathogens in specimen from bronchoscopy Growth in blood culture of pneumonia- pathogens without any other source in association with focal symptoms or signs 	Detection of pathogens in pleural fluid culture, not from already present drain

Table 2. URINARY TRACT INFECTION

	1	2	3	4
Clinical entity				
Symptoms	 Localized symptoms Mild urinary tract symptoms¹ Generalized symptoms Fever >38°C or history of fever Rigors Leukocytosis or leukopenia 	Distinct urinary tract symptoms ¹		
Signs	Localized tenderness (suprapubic or surrounding kidney)			
Radiologic findings				Abscess or inflammation
Laboratory findings	 Positive dipstick for leukocyte esterase or nitrate or pyuria in microscopic exam not from existing catheter 			
Microbiological findings	>10 ⁵ CFU/mL of a single primary urinary pathogen from existing catheter (>7 days)	>10 ⁵ CFU/mL of a single pathogen not from existing catheter	Growth in blood culture of uropathogen without any other source	 Growth in other specimen than urine from the bladder or upper urinary tract Growth of same pathogen in blood culture and urine

^{1.} Symptoms relevance graded by clinician, can include urgency, frequency, dysuria

Table 3. ABDOMINAL INFECTION

	1	2	3	4
Clinical entity				
Symptoms and signs	Localized symptoms and signs abdominal pain tenderness jaundice Generalized symptoms and signs Fever >38°C or history of fever Rigors Leukocytosis or leucopoenia	Peritonitis		Surgical confirmation
Radiological findings		Inflammatory findings		Abscess Perforation
Laboratory findings		Inflammatory peritoneal fluid		
Microbiological findings			Growth in blood culture with pathogens compatible of abdominal infection in association with focal symptoms or signs	Detection in specimen from the affected site, not from remaining drain

Table 4. GASTROINTESTINAL INFECTION

Clinical entity	1	2	3	4
Symptoms and signs		Indistinct symptoms and signs¹ Generalized symptoms and signs • Fever >38°C or history of fever • Rigors • Leukocytosis or leukopenia Localized symptoms	Distinct symptoms and signs¹ Generalized symptoms and signs • Fever >38°C or history of fever • Rigors • Leukocytosis or leukopenia Localized symptoms	

	 Diarrhea Abdominal pain Tenderness Nausea Vomiting 	 Diarrhea Abdominal pain Tenderness Nausea Vomiting 	
Microbiological findings	Enteric pathogen or clostridial toxin detected from stool or rectal swab	J	Growth in blood culture without any other source with pathogens compatible of gastrointestinal infection ²

^{1.} Symptoms and signs relevance graded by clinician

Table 5. SKIN & SOFT-TISSUE INFECTION

Clinical entity	1	2	3	4
Symptoms	Localized symptoms Pain Tenderness Swelling Heat Generalized symptoms Fever >38°C or history of fever Rigors Leukocytosis or leukopenia		Toxic shock (shock, generalized erythema w/wo vomiting) in association with focal symptoms or signs	
Signs			 Erythema (new localized) drainage of pus lymphangitis crepitations 	Surgical confirmation
Radiological findings			Presence of inflammation or abscess	
Laboratory findings		Positive antigen test for group A streptococci from suspected site of infection		
Microbiological findings		Detection in superficial specimen from suspected focus of infection of relevant pathogens	Growth in blood culture without any other source with pathogens compatible of SSTI in association with focal symptoms or signs	Detection in specimen from the affected site, not from remaining drain or superficial specimen

Table 6. BONE AND JOINT INFECTION

	1	2	3	4
Clinical entity				
Symptoms and signs	Localized symptoms and signs Swelling Tenderness Limitation of motion Heat Generalized symptoms Fever > 38°C or history of fever Rigors Leukocytosis or leukopenia			Surgical confirmation
Radiological findings		Radiological evidence		
Laboratory findings		Joint fluid analysis compatible of infection, without other explanation		
Microbiological findings			Growth in blood culture without any other source with pathogens compatible of bone and joint infection in association with focal symptoms or signs	Detection in specimen from the affected site

Table 7. CENTRAL NERVOUS SYSTEM INFECTION

	1	2	3	4
Clinical entity				
Symptoms and signs	Localized symptoms and signs • headache • irritability • mild neck stiffness Generalized symptoms • Fever >38°C or history of fever • Rigors • Leukocytosis or leukopenia	Distinct nuchal rigidity		Surgical confirmation
Radiological findings			Radiological evidence	Abscess
Laboratory findings			Increased white blood cells, protein and/or decreased glucose in CSF	
Microbiological findings			Growth in blood culture without any other source with pathogens compatible of CNS infection in association with focal symptoms or signs	Detection of pathogens in CSF Growth in specimen from the affected site

Table 8. PRIMARY BLOODSTREAM INFECTIONS

Clinical entity	1	2	3	4
Microbiological findings				Growth in blood culture of pathogens without any other source of infection. Growth in blood culture with possible skin contaminants¹ the same in two or more cultures drawn from separate venous punctures

^{1.} For example Coagulase-negative staphylococci, Propionibacterium cuti, Bacillus species, diphteroids or micrococci

Primary bloodstream infection is only considered when score is higher than in any other infection focus.

Table 9. CATHETER-RELATED INFECTION

Clinical entity	1	2	3	4
Microbiological findings		Growth in blood culture from catheter exit site		
Microbiological findings		Growth in culture of tip		
Microbiological findings		Growth in blood culture		

For a proven infection, the same microorganism needs to be found in different specimen For a proven infection based on blood culture and blood culture from catheter, growth has to be recorded >120 minutes earlier from catheter

Table 10. EAR NOSE THROAT INFECTION

Clinical entity	1	2	3	4
Symptoms	Localized symptoms Pain Generalized symptoms Fever >38°C or history of fever Rigors Leukocytosis or leukopenia			Surgical confirmation
Signs	Erythema of pharynx	 Purulent drainage Inflammation in the ear canal Retraction or decreased mobility of eardrum Visible abcess or swollen soft palate 		
Radiological findings			Radiological evidence	
Microbiological findings	Positive antigen test for group A streptococci		 Detection in specimen from the affected site Growth in blood culture without any other source with pathogens compatible of ENT infection in association with focal symptoms or signs 	

Table 11. UPPER RESPIRATORY TRACT INFECTION

Clinical entity	1	2	3	4
Symptoms	 Localized symptoms Hoarseness New onset of cough or worsening of cough Purulent sputum Dyspnea Generalized symptoms Fever >38°C or history of fever Rigors Leukocytosis or leukopenia 			
Signs	Wheezing			
Radiological findings				
Microbiological findings		Detection of viral pathogens in respiratory tract		

Table 12. NEUTROPENIC FEVER

Clinical entity	1	2	3	4
Symptoms, signs and findings	Fever >38°C sustained for 1 h or repeatedly or >38.3			
Laboratory findings	Neutrophil count <0.5 x 10 ⁹ /L			

Table 13. REPRODUCTIVE TRACT INFECTION

	1	2	3	4
Clinical entity				
Symptoms and signs	Localized symptoms Tenderness Dysuria Purulent discharge Generalized symptoms Fever >38°C or history of fever Rigors Leukocytosis or leukopenia			Surgical confirmation
Radiological findings			Radiological evidence	Abscess
Laboratory findings		Positive antigen test for group A streptococci		
Microbiological findings			 Detection in specimen from the focus of infection Growth in blood culture without any other source with pathogens compatible of RTI infection, in association postpartum or with focal symptoms or signs 	