

**Title:** Decreased high density lipoprotein cholesterol is an independent predictor for persistent organ failure, pancreatic necrosis and mortality in acute pancreatitis

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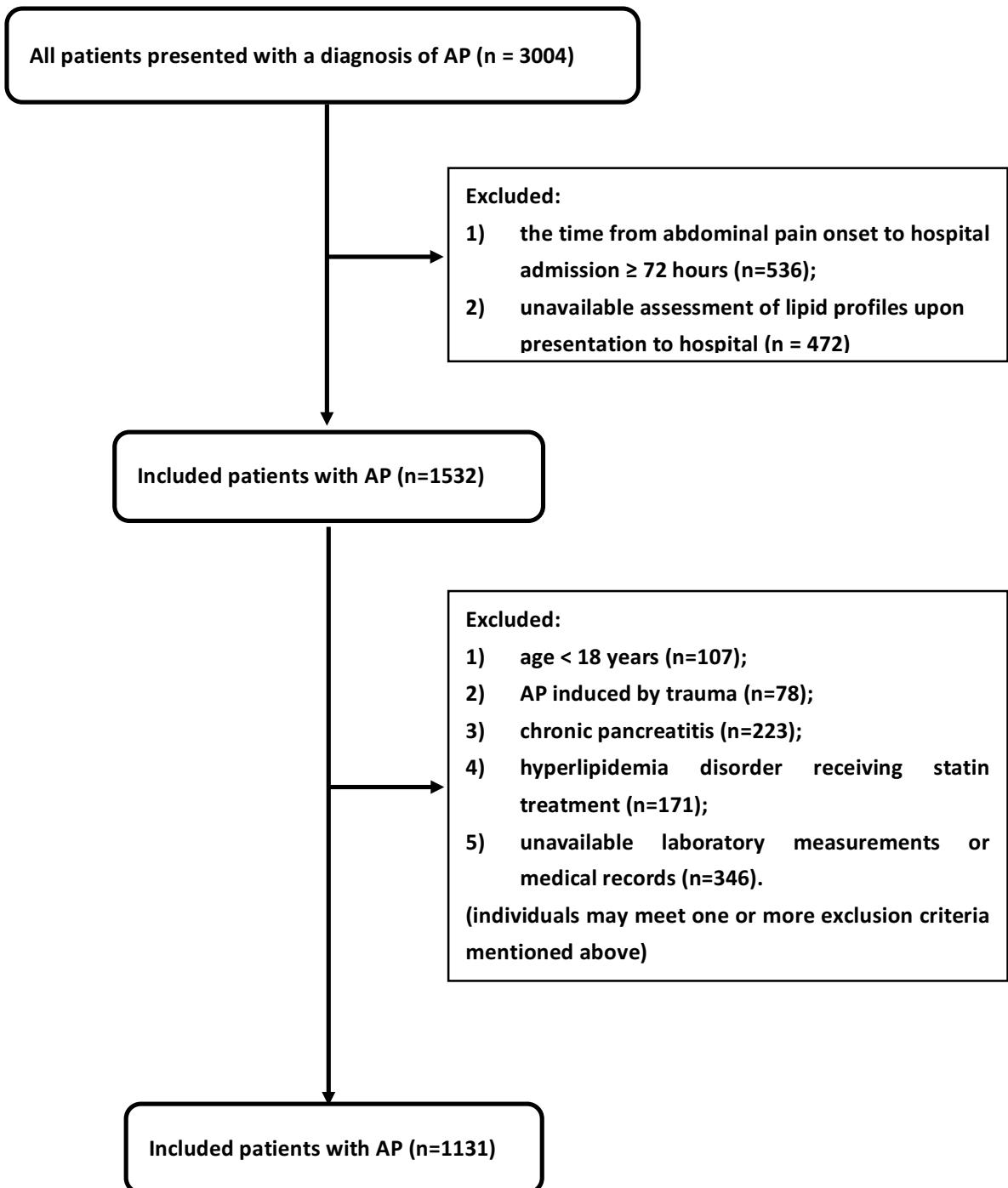
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**Supplementary figure**

**Supplementary figure 1. Flow chart illustrating patient selection**

Abbreviations: AP = acute pancreatitis



## Supplementary tables

**Supplementary table 1. Subgroup analysis: incidence of POF according to HDL-C level quartiles**

Variables	1st Quartile	2nd Quartile	3rd Quartile	4th Quartile	P for trend	P for interaction
<b>Subgroup (n)</b>						
<b>HRa (95% CI)</b>						
<b>n (number of POF)</b>						
<b>Women (n=537)</b>						
HRa (95% CI)	1.00	1.32 (0.72,	0.17 (0.07,	0.24 (0.12,	< 0.001	0.762
n	(reference)	2.42)	0.41)	0.48)		
	32	30	7	14		
<b>Men (n=804)</b>						
HRa (95% CI)	1.00	0.17 (0.10,	0.11 (0.06,	0.08 (0.04,	< 0.001	
n	(reference)	0.29)	0.20)	0.18)		
	76	22	14	8		
<b>Age ≥50 (n=542)</b>						
HRa (95% CI)	1.00	0.44 (0.24,	0.08 (0.04,	0.11 (0.06,	< 0.001	0.001
n	(reference)	0.82)	0.18)	0.23)		
	43	32	12	15		
<b>Age &lt;50 (n=799)</b>						
HRa (95% CI)	1.00	0.26 (0.15,	0.13 (0.06,	0.10 (0.04,	< 0.001	
n	(reference)	0.45)	0.27)	0.22)		
	65	20	9	7		
<b>Daily drinker (n=569)</b>						
HRa (95% CI)	1.00	0.14 (0.08,	0.16 (0.08,	0.04 (0.01,	< 0.001	0.181
n	(reference)	0.26)	0.31)	0.16)		
	68	15	11	2		
<b>No daily drinker (n=772)</b>						
HRa (95% CI)	1.00	0.95 (0.55,	0.12 (0.06,	0.21 (0.12,	< 0.001	
n	(reference)	1.64)	0.25)	0.39)		
	40	37	10	20		
<b>Current smoker (n=664)</b>						
HRa (95% CI)	1.00	0.18 (0.10,	0.08 (0.04,	0.04 (0.01,	< 0.001	0.688
n	(reference)	0.32)	0.18)	0.12)		
	70	19	8	3		
<b>No current smoker (n=677)</b>						
HRa (95% CI)	1.00	0.83 (0.48,	0.21 (0.11,	0.28 (0.15,	< 0.001	
n	(reference)	1.44)	0.42)	0.51)		
	38	33	13	19		
<b>Pre-existing co-morbidities (n=410) HRa (95% CI)</b>						
n	1.00 (reference)	0.28 (0.14,	0.08 (0.04,	0.17 (0.08,	< 0.001	< 0.001
	40	0.57)	0.18)	0.36)		
		25	11	15		
<b>No pre-existing co-morbidities (n=410) HRa (95% CI)</b>	1.00	0.35 (0.21,	0.12 (0.06,	0.07 (0.03,	< 0.001	

n	(reference) 68	0.58) 27	0.24) 10	0.17) 7		
Total leukocyte count $\geq 12 \times 10^9/L$ (n=616) HRa (95% CI)	1.00 (reference)	0.32 (0.17, 0.60)	0.38 (0.18, 0.82)	0.17 (0.07, 0.40)	< 0.001	< 0.001
n	61	35	17	12		
Total leukocyte count $< 12 \times 10^9/L$ (n=725) HRa (95% CI)	1.00 (reference)	0.31 (0.14, 0.68)	0.08 (0.02, 0.24)	0.33 (0.13, 0.85)	< 0.001	
n	47	17	2	10		
Serum glucose $\geq 7.1 \text{ mmol/L}$ , (n=751) HRa (95% CI)	1.00 (reference)	0.32 (0.19, 0.57)	0.13 (0.07, 0.27)	0.20 (0.10, 0.42)	< 0.001	< 0.001
n	90	40	16	18		
Serum glucose $< 7.1 \text{ mmol/L}$ , (n=590) HRa (95% CI)	1.00 (reference)	0.41 (0.15, 1.11)	0.25 (0.08, 0.86)	0.26 (0.07, 0.95)	< 0.001	
n	18	12	5	4		
Serum calcium $< 2 \text{ mmol/L}$ , (n=406) HRa (95% CI)	1.00 (reference)	0.40 (0.23, 0.71)	0.19 (0.09, 0.43)	0.25 (0.10, 0.64)	< 0.001	< 0.001
n	90	41	13	11		
Serum calcium $\geq 2 \text{ mmol/L}$ , (n=935) HRa (95% CI)	1.00 (reference)	0.25 (0.11, 0.60)	0.15 (0.06, 0.40)	0.21 (0.09, 0.53)	< 0.001	
n	18	11	8	11		

Abbreviations: CI, confident interval; HDL-C, high-density lipoprotein cholesterol; HR, hazard ratio; POF, persistent organ failure. HRa adjustments include sex, age, smoking status, current drinker, pre-existing co-morbidities including history of chronic pulmonary disease, chronic renal disease and cardiovascular disease, total leukocyte count, serum glucose, calcium; but exclude the exposure variable for the individual sub-groups.

**Supplementary table 2. Subgroup analysis: incidence of PNec according to HDL-C level quartiles**

Variables	1st Quartile	2nd Quartile	3rd Quartile	4th Quartile	P for trend	P for interaction
<b>Subgroup (n)</b>						
<b>HRa (95% CI)</b>						
<b>n (number of PNec)</b>						
<b>Women (n=537)</b>						
<b>HRa (95% CI)</b>	1.00 (reference)	1.77 (0.97, 3.24)	0.20 (0.09, 0.47)	0.42 (0.23, 0.78)	< 0.001	< 0.001
<b>n</b>	31	35	8	22		
<b>Men (n=804)</b>						
<b>HRa (95% CI)</b>	1.00 (reference)	0.27 (0.17, 0.42)	0.18 (0.11, 0.28)	0.14 (0.08, 0.25)	< 0.001	
<b>n</b>	101	52	35	22		
<b>Age ≥50 (n=542)</b>						
<b>HRa (95% CI)</b>	1.00 (reference)	0.69 (0.38, 1.28)	0.16 (0.08, 0.32)	0.29 (0.15, 0.53)	< 0.001	0.982
<b>n</b>	37	35	16	25		
<b>Age &lt;50 (n=799)</b>						
<b>HRa (95% CI)</b>	1.00 (reference)	0.45 (0.29, 0.69)	0.24 (0.14, 0.39)	0.15 (0.09, 0.27)	< 0.001	
<b>n</b>	95	52	27	19		
<b>Daily drinker (n=569)</b>						
<b>HRa (95% CI)</b>	1.00 (reference)	0.24 (0.15, 0.39)	0.26 (0.15, 0.43)	0.16 (0.08, 0.31)	< 0.001	< 0.001
<b>n</b>	95	41	29	14		
<b>No daily drinker (n=772)</b>						
<b>HRa (95% CI)</b>	1.00 (reference)	1.50 (0.87, 2.56)	0.19 (0.10, 0.38)	0.38 (0.22, 0.65)	< 0.001	
<b>n</b>	37	46	14	30		
<b>Current smoker (n=664)</b>						
<b>HRa (95% CI)</b>	1.00 (reference)	0.33 (0.21, 0.52)	0.20 (0.12, 0.33)	0.14 (0.08, 0.26)	< 0.001	< 0.001
<b>n</b>	94	49	30	19		
<b>No current smoker (n=677)</b>						
<b>HRa (95% CI)</b>	1.00 (reference)	1.01 (0.59, 1.74)	0.21 (0.11, 0.42)	0.38 (0.21, 0.67)	< 0.001	
<b>n</b>	38	38	13	25		
<b>Pre-existing co-morbidities (n=410)</b>						
<b>HRa (95% CI)</b>	1.00 (reference)	0.56 (0.28, 1.08)	0.20 (0.09, 0.41)	0.38 (0.19, 0.79)	< 0.001	0.008
<b>n</b>	32	28	15	19		
<b>No pre-existing co-morbidities (n=410)</b>						
<b>HRa (95% CI)</b>	1.00 (reference)	0.50 (0.34, 0.76)	0.20 (0.12, 0.33)	0.16 (0.09, 0.25)	< 0.001	
<b>n</b>						

	100	59	28	25		
<b>Total leukocyte count <math>\geq 12 \times 10^9/L</math> (n=616) HRa (95% CI)</b>	1.00 (reference) n 73	0.73 (0.43, 1.22) 58	0.61 (0.33, 1.14) 29	0.55 (0.29, 1.05) 26	0.003	< 0.001
<b>Total leukocyte count <math>&lt; 12 \times 10^9/L</math> (n=725) HRa (95% CI)</b>	1.00 (reference) n 59	0.37 (0.19, 0.71) 29	0.16 (0.07, 0.34) 14	0.32 (0.15, 0.68) 18	< 0.001	
<b>Serum glucose <math>\geq 7.1 \text{ mmol/L}</math>, (n=751) HRa (95% CI)</b>	1.00 (reference) n 108	0.34 (0.22, 0.54) 63	0.14 (0.08, 0.23) 29	0.20 (0.12, 0.34) 33	0.003	<0.001
<b>Serum glucose <math>&lt; 7.1 \text{ mmol/L}</math>, (n=590) HRa (95% CI)</b>	1.00 (reference) n 24	1.02 (0.54, 1.93) 24	0.42 (0.21, 0.85) 14	0.28 (0.13, 0.59) 11	< 0.001	
<b>Serum calcium <math>&lt; 2 \text{ mmol/L}</math>, (n=406) HRa (95% CI)</b>	1.00 (reference) n 101	0.50 (0.29, 0.87) 51	0.44 (0.22, 0.87) 24	0.64 (0.28, 1.46) 18	0.174	< 0.001
<b>Serum calcium <math>\geq 2 \text{ mmol/L}</math>, (n=935) HRa (95% CI)</b>	1.00 (reference) n 31	0.67 (0.37, 1.21) 36	0.24 (0.12, 0.47) 19	0.34 (0.18, 0.64) 26	< 0.001	

Abbreviations: CI, confident interval; HDL-C, high-density lipoprotein cholesterol; HR, hazard ratio; PNec, pancreatic necrosis. HRa adjustments include sex, age, smoking status, current drinker, pre-existing co-morbidities including history of chronic pulmonary disease, chronic renal disease and cardiovascular disease, total leukocyte count, serum glucose, calcium; but exclude the exposure variable for the individual sub-groups.

**Supplementary table 3. Subgroup analysis: in-hospital mortality according to HDL-C level quartiles**

Variables	1st Quartile	2nd Quartile	3rd Quartile	4th Quartile	P for trend	P for interaction
<b>Subgroup (n)</b>						
<b>HRa (95% CI)</b>						
<b>n (number of in-hospital death)</b>						
<b>Women (n=537)</b>						
HRa (95% CI)	1.00 (reference)	1.35 (0.63, 2.89)	- 0	0.16 (0.05, 0.48)	< 0.001	0.207
n	16	16		4		
<b>Men (n=804)</b>						
HRa (95% CI)	1.00 (reference)	0.12 (0.04, 0.36)	0.27 (0.12, 0.60)	0.09 (0.02, 0.37)	< 0.001	
n	26	4	8	2		
<b>Age ≥50 (n=542)</b>						
HRa (95% CI)	1.00 (reference)	0.63 (0.31, 1.28)	0.08 (0.03, 0.25)	0.13 (0.05, 0.34)	< 0.001	< 0.001
n	22	18	4	6		
<b>Age &lt;50 (n=799)</b>						
HRa (95% CI)	1.00 (reference)	0.10 (0.02, 0.44)	0.24 (0.08, 0.71)	- 0	< 0.001	
n	20	2	4			
<b>Daily drinker (n=569)</b>						
HRa (95% CI)	1.00 (reference)	0.08 (0.02, 0.35)	0.37 (0.14, 0.94)	- 0	< 0.001	0.574
n	22	2	6			
<b>No daily drinker (n=772)</b>						
HRa (95% CI)	1.00 (reference)	0.93 (0.46, 1.86)	0.06 (0.01, 0.25)	0.15 (0.06, 0.38)	< 0.001	
n	20	18	2	6		
<b>Current smoker (n=664)</b>						
HRa (95% CI)	1.00 (reference)	0.19 (0.06, 0.56)	0.21 (0.07, 0.64)	- 0	< 0.001	0.019
n	20	4	4			
<b>No current smoker (n=677)</b>						
HRa (95% CI)	1.00 (reference)	0.69 (0.34, 1.39)	0.13 (0.04, 0.37)	0.17 (0.06, 0.42)	< 0.001	
n	22	16	4	6		
<b>Pre-existing co-morbidities (n=410)</b>						
HRa (95% CI)	1.00 (reference)	0.48 (0.22, 1.04)	0.10 (0.03, 0.31)	0.21 (0.08, 0.57)	< 0.001	< 0.001
n	20	14	4	6		
<b>No pre-existing co-morbidities (n=410)</b>						
HRa (95% CI)	1.00 (reference)	0.28 (0.11, 0.71)	0.19 (0.06, 0.56)	- 0	< 0.001	
n						

	22	6	4				
<b>Total leukocyte count <math>\geq 12 \times 10^9/L</math> (n=616) HR<sub>a</sub> (95% CI)</b>	1.00 (reference)	0.64 (0.26, 1.56)	0.47 (0.14, 1.52)	0.32 (0.09, 1.15)	0.035	0.002	
<b>n</b>	22	16	6	4			
<b>Total leukocyte count <math>&lt; 12 \times 10^9/L</math> (n=725) HR<sub>a</sub> (95% CI)</b>	1.00 (reference)	0.23 (0.07, 0.80)	0.15 (0.03, 0.80)	0.18 (0.03, 0.96)	< 0.001		
<b>n</b>	20	4	2	2			
<b>Serum glucose <math>\geq 7.1 \text{ mmol/L}</math>, (n=751) HR<sub>a</sub> (95% CI)</b>	1.00 (reference)	0.39 (0.17, 0.87)	0.25 (0.09, 0.72)	0.19 (0.06, 0.64)	0.035	< 0.001	
<b>n</b>	36	14	6	4			
<b>Serum glucose <math>&lt; 7.1 \text{ mmol/L}</math>, (n=590) HR<sub>a</sub> (95% CI)</b>	1.00 (reference)	1.09 (0.26, 4.65)	0.46 (0.07, 3.16)	0.46 (0.07, 3.09)	0.075		
<b>n</b>	6	6	2	2			
<b>Serum calcium <math>&lt; 2 \text{ mmol/L}</math>, (n=406) HR<sub>a</sub> (95% CI)</b>	1.00 (reference)	0.39 (0.19, 0.83)	0.25 (0.08, 0.76)	0.10 (0.02, 0.48)	0.025	< 0.001	
<b>n</b>	40	16	6	2			
<b>Serum calcium <math>\geq 2 \text{ mmol/L}</math>, (n=935) HR<sub>a</sub> (95% CI)</b>	1.00 (reference)	2.24 (0.26, 19.19)	0.87 (0.07, 10.62)	1.50 (0.14, 15.71)	0.699		
<b>n</b>	2	4	2	4			

Abbreviations: CI, confident interval; HDL-C, high-density lipoprotein cholesterol; HR, hazard ratio.

HR<sub>a</sub> adjustments include sex, age, smoking status, current drinker, pre-existing co-morbidities including history of chronic pulmonary disease, chronic renal disease and cardiovascular disease, total leukocyte count, serum glucose, calcium; but exclude the exposure variable for the individual sub-groups.

**Supplementary table S4. Receiving operator curve analyses of lipid files and Ranson score in predicting outcomes**

	AUC (95%CI)	Sensitivity	Specificity	PPV	NPV
<b>POF</b>					
TG	0.606 (0.562-0.650)	0.394	0.801	0.302	0.852
T-CHO	0.526 (0.472-0.579)	0.261	0.919	0.414	0.850
HDL-C	0.778 (0.740-0.815)	0.729	0.704	0.350	0.922
LDL-C	0.658 (0.615-0.700)	0.557	0.700	0.289	0.878
TG/HDL-C	0.681 (0.639-0.723)	0.581	0.706	0.302	0.885
Non-HDL-C	0.533 (0.472-0.579)	0.305	0.875	0.348	0.852
Ranson score	0.678 (0.638-0.719)	0.562	0.749	0.329	0.887
<b>PNec</b>					
TG	0.669 (0.632-0.706)	0.451	0.841	0.513	0.805
T-CHO	0.565 (0.523-0.607)	0.297	0.918	0.572	0.779
HDL-C	0.734 (0.700-0.769)	0.723	0.661	0.442	0.867
LDL-C	0.656 (0.619-0.693)	0.575	0.690	0.408	0.814
TG/HDL-C	0.717 (0.683-0.752)	0.598	0.749	0.469	0.834
Non-HDL-C	0.610 (0.570-0.655)	0.349	0.910	0.584	0.788
Ranson score	0.701 (0.668-0.734)	0.529	0.776	0.467	0.816
<b>In-hospital mortality</b>					
TG	0.550 (0.485-0.615)	0.947	0.216	0.080	0.983
T-CHO	0.602 (0.515-0.689)	0.395	0.911	0.242	0.954
HDL-C	0.768 (0.710-0.827)	0.632	0.777	0.170	0.967
LDL-C	0.678 (0.609-0.747)	0.658	0.652	0.120	0.964
TG/HDL-C	0.630 (0.567-0.694)	0.290	0.907	0.183	0.947
Non-HDL-C	0.562 (0.477-0.647)	0.263	0.963	0.339	0.948
Ranson score	0.745 (0.694-0.795)	0.724	0.723	0.158	0.973

Abbreviations: AUC: area under the curve; CI: confidence intervals; HDL-C, high-density lipoprotein cholesterol; HR, hazard ratio; LDL-C, low-density lipoprotein cholesterol; NPV: negative predictive value; PNec, pancreatic necrosis; POF, persistent organ failure; PPV: positive predictive value; ROC, receiver operating curve; T-CHO, total cholesterol; TG, triglycerides.