# **BMJ Open** Acute coronary syndromes presentations and care outcomes in white, South Asian and Chinese patients: a cohort study

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## ABSTRACT

**Objectives** Successful treatment of acute coronary syndrome (ACS) relies on its rapid recognition. It is unclear whether the accepted presentation of chest pain applies to different ethnic groups. We thus examined potential ethnic variations in ACS symptoms and clinical care outcomes in white, South Asian and Chinese patients.

**Design** Cross-sectional survey.

**Setting** Participants were hospitalised at 1 of 12 Canadian centres across four provinces.

**Participants** 1334 patients with ACS (630 white; 488 South Asian; 216 Chinese).

Main outcome measures ACS presentation symptoms (classic/typical midsternal pain/discomfort with or without radiation to the left neck, shoulder or arm) were assessed by self-report. Clinical care outcomes (time to emergency room [ER] presentation, cardiac catheterisation; receipt of cardiac catheterisation, percutaneous coronary intervention [PCI] or coronary artery bypass grafting [CABG]) were obtained by health record audit.

Results The mean age of the sample was 62 years and 30% had ST-elevation myocardial infarction (STEMI). The most common presenting symptom was midsternal pain/ discomfort of any intensity regardless of ethnic status. Yet, a substantial proportion of patients reported atypical symptoms (33% white, 19% South Asian, 20% Chinese; p<0.006). After adjustment for age, sex, education, current smoking, extent of coronary artery disease, presence of diabetes or chronic kidney disease and STEMI vs non-STEMI/unstable angina, South Asians were more likely to present with at least moderate intensity midsternal pain/ discomfort (adjusted OR [AOR] 1.44; 95% Cl 1.05 to 1.98), whereas Chinese were less likely to present with radiating symptoms (AOR 0.53; 95% CI 0.38 to 0.74) compared with whites. South Asians with atypical pain (relative to those with midsternal pain/discomfort) took significantly longer to present to the ER (p=0.037), and were less likely to receive PCI (p=0.008) or CABG (p=0.041). **Conclusions** Atypical presentations were associated with

greater delays in arrival to the emergency department and reduced invasive cardiovascular care in South Asians.

# INTRODUCTION

The burden of acute coronary syndrome (ACS) is substantial and it is now the leading

# Strengths and limitations of this study

- This was a rigorously designed study which included participants who spoke a variety of languages and used highly systematic and validated translation and data collection processes.
- A number of mechanisms were used to obtain a data regarding participants' acute coronary syndrome (ACS) symptoms.
- Only participants who survived their ACS event were included in this study. It is not possible to know if patients who succumbed to their ACS event or did not seek care had different symptoms than those who were diagnosed and survived.
- There was a noteworthy proportion of South Asian and Chinese patients approached to participate who refused.

cause of death and disability worldwide.<sup>1 2</sup> Life-saving therapies for ACS are dependent on early and timely recognition of symptoms. Atypical presentation of ACS symptoms leads to delays in recognising symptoms by both patients and healthcare providers, and can result in misdiagnosis, delayed treatment, receipt of fewer evidence-based therapies and higher in-hospital morbidity and mortality.<sup>3</sup>

The 'classic' ACS presentation, based on the presentation of men of European descent, includes moderate to severe intensity central chest with or without left arm, neck or jaw radiating pain or discomfort.<sup>4 5</sup> This description has been adopted internationally, though some studies suggest that ACS symptoms may differ by ethnicity.<sup>6-11</sup> Although these studies yielded discordant findings, some noted that non-white patients may present with additional symptoms (eg, gastrointestinal, dyspnoea, nausea)<sup>6-9 11</sup> and South Asians may have more diffuse pain and more frequent back pain<sup>10</sup> than whites. It is difficult to draw robust conclusions from the

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Dr Kathryn King-Shier; kingk@ucalgary.ca current evidence given variable methods of data collection, inadequate cross-cultural language adaptation, small sample sizes and insufficient adjustment for other factors (eg, sex, age, clinical factors) that may influence ACS presentation.

South Asians have a higher incidence of ACS and earlier onset relative to other ethnic groups.<sup>12 13</sup> Though Chinese currently have a lower incidence of acute myocardial infarction (AMI) relative to whites and South Asians,<sup>14 15</sup> cardiovascular events in China are expected to rise by more than 50% by 2030.<sup>16</sup> A rigorous investigation of ACS symptom presentation, time to arrival to the emergency department and clinical care outcomes between these ethnic groups is warranted to ensure appropriate public health messaging and/or inappropriate stereotyping of presenting symptoms. We thus aimed to examine potential ethnic variations in ACS symptom guestionnaire and visual identifiers in a multicentre cohort of white, South Asian and Chinese patients hospitalised in Canada.

#### **METHODS**

#### Study design

We conducted a cross-sectional survey of a cohort of white (ie, of European descent), South Asian (ie, descended from India, Pakistan, Bangladesh, Sri Lanka) and Chinese (ie, descended from China, Hong Kong, Taiwan, Macau) patients hospitalised with physician-confirmed ACS (unstable angina, ST-elevation myocardial infarction [STEMI] or non-STEMI<sup>5</sup>).

#### Setting and sample

Following institutional ethics review at each participating hospital, participants were recruited consecutively from 12 urban hospitals in four Canadian provinces (>90% of Canada's immigrants reside in urban areas and the majority reside in these provinces<sup>17</sup>) between September 2010 and December 2015. Inclusion criteria were age  $\geq 19$ years; admission to hospital with a confirmed diagnosis of ACS (as identified on health record); self-reported ethnicity for either white (European), South Asian or Chinese; and speaking English, Punjabi, Tamil, Urdu, Hindi or Gujarati (among the most common languages of Canadian residents from South Asia),<sup>17</sup> Cantonese or Mandarin. Nearly 95% of South Asians and all Chinese residing in Canada would have been able to speak one or more of the languages used in this study.<sup>17</sup> Exclusion criteria were multiple ethnic origins (ie, mixed race) or known cognitive deficits (ie, Alzheimer's disease, dementia, uncontrolled psychiatric disorder), as identified on the health record. All participants provided informed consent to participate in this study.

# Patient and public involvement

Patients were not involved in the conception, design or interpretation of this study. However, we have a detailed plan to disseminate this information widely to healthcare providers as well as potential patients, through traditional and social media.

# **Data collection**

Data were collected while participants were in hospital (within the first 5 days), in the patient's preferred language (as listed above) by highly trained like-speaking research assistants.

As reported elsewhere,<sup>18</sup> rigorous translation processes were undertaken for study materials.

Questionnaires were developed based on the current literature and our earlier work.<sup>10 19-21</sup> We, like Teoh et al,<sup>10</sup> asked participants to mark, on a gender-neutral torso silhouette (pictograph), all the locations of the pain/discomfort that brought them to hospital. Each participant used the same type of pen to 'colour in' the areas. Then, the participant was prompted to identify if he/she experienced other symptoms such as nausea, diaphoresis, dyspnoea, dizziness or weakness. Next, they were asked to identify their 'chief' or 'main' symptom, describe its nature by pointing to pictorial identifiers (ie, stabbing, heavy, shooting, burning, squeezing)<sup>9 21</sup> and finally identify its intensity (Likert-type scale, 0-10 where 0 was no pain). Pain/discomfort severity rated as <5 was considered mild and  $\geq 5$  or higher was considered moderate to severe pain.

A health record audit was undertaken to collect additional demographic and as well as clinical data, once the participant had been discharged from hospital.

The presence of classic (typical) ACS presentation was defined as having midsternal or having midsternal with radiating left neck, shoulder or arm pain/discomfort of any or at least moderate intensity. The clinical care outcomes of interest were time to emergency room (ER) presentation and cardiac catheterisation, as well as receipt of cardiac catheterisation, percutaneous coronary intervention (PCI) or coronary artery bypass grafting (CABG).

There was potential for bias when using this design and data collection method. This was a self-selected group of patients (ie, selection bias) who agreed to participate in the study. Also, the retrospective nature of the design could lead to recall bias given that patients were asked about their symptoms up to 5 days following hospital admission.

#### **Statistical analysis**

Descriptive statistics and  $\chi^2$  tests were used to identify differences in participant characteristics, as well as symptoms and location(s) of pain/discomfort between ethnic groups. Each variable had a 'not entered' category. In binary variables of present/not present, 'not entered' was assumed to be not present. To analyse presenting symptoms data, a grid was placed over the original torso silhouette to identify the location(s) of pain/discomfort in a standardised manner.<sup>10</sup> These data were also presented as proportions of each participant group who identified any pain/discomfort in the various areas as well as moderate to severe pain/discomfort rated as  $\geq 5/10$  on a Likert-type

	White (n=630)	South Asian (n=488)	Chinese (n=216)	P value
Age, mean (SD), years	65.9 (13.0)	62.2 (12.5)	65.1 (12.6)	<0.0001
Male	276 (43.8%)	355 (72.8%)	154 (71.3%)	<0.0001
English Language Survey	630 (100%)	440 (90.2%)	161 (74.5%)	<0.0001
Immigration status				<0.0001
Canadian-born	455 (72.1%)	11 (2.3%)	10 (4.6%)	
Immigrant <20 years	15 (2.4%)	189 (38.8%)	81 (36.6%)	
Immigrant >20 years	160 (25.4%)	288 (59.0%)	123 (56.9%)	
None documented	1 (0.25)	0	2 (0.9%)	
Marital status				<0.0001
Married/Common law	362 (57.4%)	383 (78.5%)	174 (80.6%)	
Other	269 (42.7%)	105 (21.6%)	42 (19.5%)	
Highest level education				0.0001
None	8 (1.3%)	31 (6.4%)	7 (3.2%)	
<high school<="" td=""><td>94 (14.9%)</td><td>83 (17.0%)</td><td>41 (19.0%)</td><td></td></high>	94 (14.9%)	83 (17.0%)	41 (19.0%)	
≥High school	529 (83.9%)	374 (76.7%)	168 (77.8%)	
None documented	0	0	0	
Current smoker	123 (19.5%)	53 (10.9%)	27 (12.5%)	0.0002
Extent of coronary artery disease				<0.0001
Normal/<50% disease	92 (14.6%)	50 (10.3%)	36 (16.7%)	
1–2-vessel disease	148 (23.5%)	91 (18.7%)	33 (15.3%)	
3-vessel disease/LAD	272 (43.2%)	293 (60.0%)	109 (50.5%)	
Left main	11 (1.8%)	4 (0.8%)	10 (4.6%)	
Not entered/missing	107 (17.0%)	50 (10.3%)	28 (13.0%)	
Diabetes	193 (30.6%)	224 (45.9%)	64 (29.6%)	<0.0001
Chronic renal disease	74 (11.7%)	45 (9.2%)	18 (8.3%)	0.229
Acute coronary syndrome				0.187
Unstable angina	205 (32.5%)	132 (27.1%)	61 (28.2%)	
Non-STEMI	228 (36.2%)	208 (42.6%)	83 (38.4%)	
STEMI	188 (29.8%)	145 (29.7%)	68 (31.5%)	
Unspecified	9 (1.5%)	3 (0.6%)	4 (1.8%)	

LAD, left anterior descending artery; STEMI, ST-elevation myocardial infarction.

scale. Crude and adjusted OR (AOR) with 95% CI were examined using logistic regression for South Asians and Chinese relative to whites having any midsternal pain/ discomfort or any midsternal with radiating left neck, shoulder or arm pain/discomfort, as well as moderate to severe (≥5/10) midsternal pain/discomfort or midsternal with radiating left neck, shoulder or arm pain/discomfort. Finally, crude and AOR (as above) with 95% CI were examined to determine the likelihood of receiving cardiac catheterisation, PCI and CABG for each ethnic group, based on having any midsternal pain/discomfort or any midsternal with radiating left neck, shoulder or arm pain/discomfort.

Models were adjusted for demographic and clinical characteristics associated with atypical symptoms (age, sex, education, current smoker, extent of coronary artery disease [based on cardiac catheterisation], presence of diabetes or chronic kidney disease [CKD; as defined by the Charlson Comorbidity Index<sup>22</sup>], and STEMI vs non-STEMI/unstable angina). All data were analysed using SAS V.9.4.

#### **Pictographic analysis**

We used a heat-mapping technology to portray where participants identified that they pain/discomfort on the torso silhouettes.<sup>10</sup> Given that the numbers of participants varied by ethnic group, the mapping was based on proportion of participants who coloured areas on the torsos versus the absolute number.

Table 2 Reported acute coronary syndrome symptoms by ethnicity				
	White (n=630)	South Asian (n=488)	Chinese (n=216)	P value
Pain/Discomfort location				
Midsternal	559 (88.7%)	438 (89.9%)	194 (89.8%)	0.828
Left shoulder	310 (49.1%)	239 (49.0%)	72 (33.3%)	0.0006
Left arm	206 (32.7%)	140 (28.7%)	22 (10.2%)	<0.0001
Left jaw	122 (19.3%)	67 (13.7%)	21 (9.7%)	0.003
Left neck	114 (18.1%)	69 (14.1%)	19 (8.8%)	0.0006
Midsternal with radiating left neck, shoulder or arm pain/discomfort	435 (69.0%)	316 (63.8%)	123 (57.0%)	<0.0001
Midsternal pain/discomfort with intensity 🐵 >5	482 (76.5%)	394 (80.7%)	172 (79.6%)	0.0435
Midsternal with radiating left neck, shoulder or arm pain/discomfort with intensity $\odot$ >5	385 (61.6%)	285 (58.4%)	111 (51.4%)	<0.0001
Nature of pain/discomfort				
Pressure	312 (49.5%)	174 (35.7%)	93 (43.1%)	<0.0001
Squeezing	118 (18.7%)	120 (24.6%)	57 (26.4%)	0.016
Burning	82 (13.0%)	71 (14.6%)	25 (11.6%)	0.533
Stabbing	48 (7.6%)	68 (14.0%)	17 (7.9%)	0.001
Shooting/Moving	30 (4.8%)	39 (8.0%)	11 (5.1%)	0.065
None	40 (6.4%)	16 (3.3%)	13 (6.0%)	0.059
Not documented	0	0	0	-
Other symptoms				
Shortness of breath	386 (61.2%)	303 (62.1%)	132 (61.1%)	0.952
Diaphoresis	313 (49.7%)	239 (49.0%)	103 (47.7%)	0.877
Dizziness	329 (52.2%)	221 (45.3%)	99 (45.8%)	0.047
Nausea or vomiting	235 (37.3%)	145 (29.7%)	56 (25.9%)	0.002

# RESULTS

Of 3243 persons screened for eligibility, 1042 did not meet eligibility criteria and 903 patients declined enrolment in the study. Of the final 1334 patients (58.8% men), 47.2% were white, 36.6% were South Asian and 16.2% were Chinese (see table 1).

# **Baseline characteristics**

Over 72% of whites were Canadian-born whereas the majority of South Asians and Chinese had immigrated to Canada >20 years earlier. Among these three ethic groups, whites were most likely to be a current smoker and have 1–2-vessel disease, South Asians were most likely to have 3-vessel disease and diabetes and the greatest proportion of each patient group (36.2% whites, 42.6% South Asian, 38.4% Chinese) were admitted with non-STEMI.

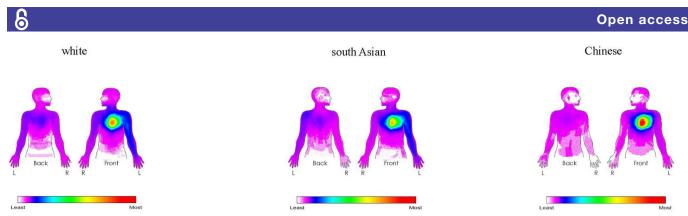
## **ACS symptom presentation**

The most common presenting symptom across ethnic groups was midsternal pain/discomfort of any intensity (89.3%) followed by left shoulder pain/discomfort (46.6%) (table 2). Most of the pain or discomfort was rated as moderate to severe ( $\geq 5/10$ ) across ethnicities. More than 78% of all patients reported the classic presentation of at least moderate intensity midsternal pain/

discomfort, and more than 58% of all patients reported moderate to severe intensity midsternal chest with any of radiating left neck, shoulder or arm pain/discomfort. In these unadjusted analyses, whites were less likely to report having the classic presentation of at least moderate intensity midsternal pain/discomfort while the Chinese were less likely to report having at least moderate intensity midsternal pain/discomfort with radiation to left neck, shoulder or arm. Extent of coronary artery disease was not associated with having classic symptoms in whites (p=0.973), South Asians (p=0.562) or Chinese (p=0.304) patients.

The majority of patients described their pain or discomfort as pressure, squeezing or burning (81.3% whites, 74.8% South Asians, 81.0% Chinese), although south Asians reported significantly more stabbing pain than their counterparts. Other symptoms were also prevalent among all ethnic groups: shortness of breath (62.4%), diaphoresis (49.8%), dizziness (49.3%) and nausea/ vomiting (33.1%). There were significant differences between groups in reporting these symptoms with whites reporting more dizziness and nausea or vomiting.

The pictographic analysis of the torsos (figure 1) shows that the profiles were very similar for whites and South



**Figure 1** Pictographic for each ethnic group. The colour gradient (white/purple to red) indicates no/few participants to most (>65%) of participants identified pain/discomfort in that location.

Asians, whereas the Chinese tended to report more central as well as less back or left arm pain/discomfort.

In the analysis stratified by ethnic group (table 2), ethnicity was not significantly associated with reporting midsternal pain/discomfort of any intensity. However, in adjusted (age and sex; age sex, education, current smoker, extent of coronary artery disease, diabetes, CKD and STEMI vs non-STEMI/unstable angina) models (table 3), South Asians were significantly more likely to report having midsternal pain/discomfort of moderate to severe intensity. The Chinese participants were less likely to report having radiating ACS symptoms (ie, midsternal, with left neck, shoulder or arm pain or discomfort) of any as well as moderate to severe intensity, relative to whites in crude and adjusted models.

# **Clinical care outcomes**

The mean time to ER presentation, for those who had a distinct time of symptom onset, was 5.53 to 7.41 hours (table 4). South Asians with atypical symptoms had significantly longer delays in arrival to the emergency department than those who had any typical symptoms. The mean time to receipt of cardiac catheterisation following ER presentation was 3–3.93 hours. Whites with atypical symptoms had significantly longer delays in time to receipt of cardiac catheterisation than those with typical symptoms. South Asian participants who had any typical midsternal pain/discomfort were more likely to receive PCI but less likely to receive CABG than those with atypical symptoms. Table 4 also shows that South Asian participants who had any typical midsternal with left neck, shoulder or arm pain/discomfort were more likely to receive PCI than those who had atypical symptoms.

However, after adjustment, South Asians with atypical symptoms were least likely to receive PCI than those with typical symptoms (table 5).

# DISCUSSION

The classic (typical) presentation of moderate to severe intensity midsternal chest pain or discomfort was the most common presenting ACS symptom across ethnic groups. Yet, a substantial proportion of patients, up to one-third, in each group studied had non-classic (atypical) ACS presentations. Differences in ACS symptom presentation among whites, South Asian, and Chinese patients were statistically and clinically significant, as South Asians were more likely to present with classic ACS symptoms and Chinese were less likely to present with radiating symptoms in all

Table 3 Crude and adjusted OR for ACS symptom presentation						
	Any intensity			Moderate to sever	re intensity	
Ethnic group	Crude OR (95% CI)	Adjusted* OR (95% CI)	Adjusted <sup>†</sup> OR (95% CI)	Crude OR (95% Cl)	Adjusted* OR (95% CI)	Adjusted† OR (95% CI)
At least moderate intensity midsternal pain/discomfort						
White	1.0	1.0	1.0	1.0	1.0	1.0
South Asian	1.11 (0.76 to 1.63)	1.04 (0.70 to 1.56)	1.04 (0.68 to 1.57)	1.29 (0.96 to 1.72)	1.44 (1.06 to 1.96)	1.43 (1.04 to 1.97)
Chinese	1.12 (0.68 to 1.86)	1.12 (0.67 to 1.89)	1.18 (0.70 to 2.00)	1.20 (0.82 to 1.75)	1.40 (0.95 to 2.07)	1.41 (0.95 to 2.09)
At least moderate midsternal, with left neck, shoulder or arm pain/discomfort						
White	1.0	1.0	1.0	1.0	1.0	1.0
South Asian	0.86 (0.68 to 1.10)	0.83 (0.64 to 1.06)	0.81 (0.62 to 1.05)	0.94 (0.74 to 1.19)	0.95 (0.74 to 1.21)	0.92 (0.71 to 1.75)
Chinese	0.44 (0.32 to 0.61)	0.45 (0.32 to 0.62)	0.46 (0.33 to 0.63)	0.50 (0.36 to 0.69)	0.53 (0.38 to 0.74)	0.53 (0.38 to 0.75)

\*Adjusted for age and sex.

†Adjusted for age, sex, education, current smoker, extent of coronary artery disease (no disease vs low risk or high risk), diabetes, chronic kidney disease and ST-elevation myocardial infarction (STEMI) vs non-STEMI/unstable angina. ACS, acute coronary syndrome.

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Table 4 Clinical care outcomes by symptoms and ethnicity	by symptoms an	d ethnicity							
	White			South Asian			Chinese		
	Typical (n=482)	Atypical (n=148)	P value	Typical (n=394)	Atypical (n=94)	P value	Typical (n=172)	Atypical (n=44)	P value
Any midsternal pain/discomfort	n=451	n=116		n=359	n=71		n=144	n=32	
Time to ER presentation (hours; if distinct time of onset) Mean (SD)	6.43 (5.77)	6.11 (5.53)	0.591	5.69 (5.35)	7.17 (5.94)	0.037	5.92 (5.49)	7.41 (6.10)	0.176
Median (IQR)	3.0 (6.5)	3.0 (6.0)	0.872	3.0 (6.0)	5.0 (6.5)	0.127	3.0 (6.0)	6.0 (6.5)	0.318
Time to CATH (hours) Mean (SD)	3.35 (1.76)	3.93 (1.57)	0.001	3.10 (1.80)	3.10 (1.72)	0.975	3.09 (1.82)	3.58 (1.72)	0.134
Median (IQR)	4.0 (2.0)	5.0 (1.0)	0.002	3.0 (2.0)	2.0 (2.0)	0.862	3.0 (2.0)	5.0 (1.5)	0.109
	n (%)	u (%)		n (%)	n (%)		N (%)	n (%)	
CATH	402 (83.4%)	124 (83.8%)	0.913	359 (91.1%)	81 (86.2%)	0.148	151 (87.8%)	38 (86.4%)	0.826
PCI	228 (47.3%)	60 (40.5%)	0.149	211 (53.6%)	36 (38.3%)	0.008	81 (47.1%)	19 (43.2%)	0.777
CABG	44 (9.1%)	17 (11.5%)	0.396	38 (9.6%)	16 (17.0%)	0.041	13 (7.6%)	5 (11.4%)	0.636
Any midsternal with left neck, shoulder, or arm pain/discomfort	n=330	n=237		n=234	n=196		n=68	n=108	
Time to ER presentation (hours; if distinct time of onset) Mean (SD)	6.41 (5.82)	6.31 (5.57)	0.836	5.53 (5.35)	6.42 (5.60)	0.092	6.0 (5.65)	6.31 (5.62)	0.726
Median (IQR)	3.0 (6.5)	3.0 (3.0)	0.926	2.0 (6.0)	3.0 (6.5)	0.037	3.0 (6.0)	3.0 (6.0)	0.649
Time to CATH (hours) Mean (SD)	3.40 (1.76)	3.61 (1.68)	0.154	3.0 (1.77)	3.2 (1.79)	0.241	3.28 (1.81)	3.14 (1.81)	0.615
Median (IQR)	4.0 (2.0)	5.0 (1.5)	0.192	3.0 (2.0)	3.0 (2.0)	0.202	4.0 (2.0)	3.0 (2.0)	0.635
	n (%)	n (%)		n (%)	n (%)		n (%)	n (%)	
CATH	306 (86.0%)	220 (80.3%)	0.058	235 (91.1%)	205 (89.1%)	0.469	65 (82.3%)	124 (90.5%)	0.126
PCI	172 (48.3%)	116 (42.3%)	0.135	143 (55.4%)	104 (45.2%)	0.024	41 (51.9%)	59 (43.1%)	0.361
CABG	33 (9.3%)	28 (10.2%)	0.690	30 (11.6%)	24 (10.4%)	0.675	4 (5.1%)	14 (10.2%)	0.307
CABG, coronary artery bypass graft surgery; CATH, catheterisation; El	urgery; CATH, cath	leterisation; ER, en	nergency room	; PCI, percutaneou	R, emergency room; PCI, percutaneous coronary intervention.	ntion.			

Table 5 Crude and adjusted OR for clinical care outcomes					
Atypical versus typical	Crude OR (95% CI)	Adjusted* OR (95% CI)			
Any midsternal pa	in/discomfort				
CATH					
White	1.03 (0.62 to 1.69)	1.15 (0.53 to 2.49)			
South Asian	0.61 (0.31 to 1.20)	0.66 (0.22 to 1.98)			
Chinese	0.89 (0.33 to 2.35)	2.77 (0.64 to 11.90)			
PCI					
White	0.76 (0.52 to 1.10)	0.73 (0.45 to 1.19)			
South Asian	0.54 (0.34 to 0.85)	0.57 (0.32 to 1.00)			
Chinese	0.87 (0.45 to 1.69)	1.10 (0.45 to 2.68)			
CABG					
White	1.29 (0.71 to 2.34)	1.59 (0.81 to 3.09)			
South Asian	1.92 (1.02 to 3.62)	1.90 (0.96 to 3.77)			
Chinese	1.63 (0.56 to 4.73)	1.76 (0.53 to 5.80)			
Any midsternal, w	th left neck, shoulder o	or arm pain/discomfort			
CATH					
White	0.67 (0.44 to 1.02)	0.90 (0.47 to 1.73)			
South Asian	0.80 (0.44 to 1.46)	1.37 (0.55 to 3.41)			
Chinese	2.08 (0.92 to 4.68)	4.94 (1.41 to 17.31)			
PCI					
White	0.79 (0.57 to 1.08)	0.84 (0.56 to 1.26)			
South Asian	0.66 (0.46 to 0.95)	0.74 (0.47 to 1.18)			
Chinese	0.69 (0.40 to 1.21)	0.49 (0.23 to 1.05)			
CABG					
White	1.11 (0.66 to 1.89)	1.24 (0.69 to 2.21)			
South Asian	0.89 (0.50 to 1.56)	0.92 (0.50 to 1.70)			
Chinese	1.82 (0.63 to 5.31)	2.10 (0.65 to 6.85)			

\*Adjusted for age, sex, education, current smoker, extent of coronary artery disease (no disease vs low risk or high risk), diabetes, CKD and STEMI vs non-STEMI/unstable angina. CABG, coronary artery bypass graft surgery; CATH, catheterisation; PCI, percutaneous coronary intervention.

models relative to whites. Our study extends the findings from previous work by employing rigorous cross-cultural and language adaptation of study questionnaires, and including visual diagrams across multiple centres.

Findings from other studies have some similar conclusions. Like the south Asian and Chinese participants in our study, Blacks have been reported to have a similar prevalence of chest pain relative to whites,<sup>6–9</sup> while other studies suggest that non-white patients with ACS are more likely to have chest pain than whites. A study from a single hospital in the UK reported South Asians are more frequently to have 'classic chest pain with radiation' than whites (90 vs 82%).<sup>10</sup> A study of 390 patients with ACS residing in the Asia-Pacific rim revealed patients from India, China and Korea were significantly more likely to report typical pain than their white counterparts.<sup>11</sup> A small study of whites, blacks (US residents) and Koreans (Korea residents) revealed Koreans reported significantly more frequent and greater radiation of chest pain.<sup>8</sup> Yet, our earlier study from a single health region revealed that South Asian and Chinese patients with AMI were less likely to report classic symptom presentation.<sup>19</sup> These inconsistencies may be explained by the controlled analyses we used, and the lack of cross-cultural (and language) adaptation in other studies.

We found significant differences among ethnic groups in the intensity and nature of the pain/discomfort as well as the presence of additional symptoms. Ethnicity and culture may influence patients identifying, recognising and acknowledging urgent symptoms as such, as well as making the decision to seek care. Though not consistent, study findings suggest that people who represent ethnic minorities (who are not residing in their country of origin) have greater pain sensitivity in both clinical and experimental conditions relative to the ethnic majority (eg, whites). Though most of this research has been undertaken in African Americans and Hispanics relative to whites,<sup>23</sup> what little research that has focused on south Asians or Chinese suggests that both groups may report less pain tolerance relative to whites.<sup>24–25</sup> Single-centre studies have revealed that south Asians and Chinese with AMI tended to report atypical symptoms and did not recognise or accept their symptoms as 'urgent'.<sup>19 26</sup> Research findings suggest that expression of symptoms, and language and semantic differences of symptom descriptors may also account for discrepancies in reporting symptoms. Given that pain/discomfort are subjective experiences, how they are portraved are culturally bound.<sup>23-30</sup> For example, the Worchester Heart Attack Study revealed that south Asian participants were significantly less accepting of overtly expressing pain,<sup>21</sup> and both south Asians and Chinese are viewed as more stoic in their presentations.<sup>29</sup>

We identified little overall ethnic variation in time to emergency room presentation, though south Asians who had atypical symptoms were more likely to have a longer time to presentation to the emergency room (mean of 1.48 hours longer) relative to those who had typical symptoms. In a study of 440 398 registry patients, Ting *et al*<sup>11</sup> demonstrated that delays from symptom onset to ER presentation were associated with reduced likelihoods of receiving any reperfusion therapy in the general population. Authors of other American studies have demonstrated ethnic (eg, African Americans, Hispanics, Pacific Islanders) differences relative to whites in longer time to ER presentation and reduced access to treatment (reperfusion therapies) for ACS events.<sup>30-33</sup>

Overall, and after adjustment, South Asians with atypical symptoms were least likely of the ethnic groups to receive PCI than those with typical symptoms. The reasons for this difference are not clear. It is possible that South Asians with atypical presentations arrive too late for emergent coronary revascularisation, there are biases from healthcare providers or this population may be more likely to refuse invasive cardiovascular procedures.

#### **Study limitations**

There were some limitations to this study. First, we studied participants who survived their ACS event. It is not possible to know if patients who succumbed to their ACS event or did not seek care had different symptoms than those who were diagnosed and survived. Second, we studied only those who agreed to participate. The proportion of south Asian and Chinese patients approached to participate and who refused was noteworthy. Third, we studied only patients in Canada who were admitted to selected hospitals. Finally, the cross-sectional design renders only observations and no basis for attributing causation. We undertook the following to balance these limitations: we employed a rigorous approach to comparing symptom presentation between cultural groups in this large cross-sectional study; we included participants who spoke a variety of languages, using highly systematic and validated translation and data collection processes; and had a large sample size.

## **CONCLUSIONS**

A substantial proportion of white, south Asian and Chinese patients with ACS do not report classic ACS symptoms. Differences in ACS symptoms presentation among whites, South Asians and Chinese patients are significant with south Asians being more likely to report midsternal pain/discomfort and Chinese being less likely to report radiating pain/discomfort, relative to whites. On the whole, the mean time to ER presentation was unacceptably long for each ethnic group and especially so for south Asians. These delays can reduce the opportunity to receive evidence-based reperfusion therapies. From a public health perspective, we need to include non-chest pain symptoms for screening and public awareness campaigns for ACS among South Asian, Chinese and white persons.

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#### REFERENCES

- World Health Organization. Media Centre Cardiovascular Diseases (CVDs). Available at www.who.int/mediacentre/factsheets/fs317/en/ (Accessed on 12 Feb 2016).
- Vedanthan R, Seligman B, Fuster V. Global perspective on acute coronary syndrome: a burden on the young and poor. *Circ Res* 2014;114:1959–75.
- El-Menyar A, Zubaid M, Sulaiman K, et al. Atypical presentation of acute coronary syndrome: a significant independent predictor of inhospital mortality. J Cardiol 2011;57:165–71.
- 4. Anderson JL, Adams CD, Antman EM, et al. ACC/AHA 2007 guidelines for the management of patients with unstable angina/non-ST-Elevation myocardial infarction: a report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines (Writing Committee to Revise the 2002 Guidelines for the Management of Patients With Unstable Angina/Non-ST-Elevation Myocardial Infarction) developed in collaboration with the American College of Emergency Physicians, the Society for Cardiovascular Angiography and Interventions, and the Society of Thoracic Surgeons endorsed by the American Association of Cardiovascular and Pulmonary Rehabilitation and the Society for Academic Emergency Medicine. J Am Coll Cardiol 2007;50:e1–157.
- Théroux P, Fuster V. Acute coronary syndromes: unstable angina and non-Q-wave myocardial infarction. *Circulation* 1998;97:1195–206.
- Lee H, Bahler R, Chung C, et al. Prehospital delay with myocardial infarction: the interactive effect of clinical symptoms and race. Appl Nurs Res 2000;13:125–33.
- Klingler D, Green-Weir R, Nerenz D, et al. Perceptions of chest pain differ by race. Am Heart J 2002;144:51–9.
- 8. Lee H, Bahler R, Park OJ, *et al.* Typical and atypical symptoms of myocardial infarction among African-Americans, whites, and Koreans. *Crit Care Nurs Clin North Am* 2001;13:531–9.
- Hravnak M, Whittle J, Kelley ME, et al. Symptom expression in coronary heart disease and revascularization recommendations for black and white patients. Am J Public Health 2007;97:1701–8.
- Teoh M, Lalondrelle S, Roughton M, *et al.* Acute coronary syndromes and their presentation in Asian and Caucasian patients in Britain. *Heart* 2007;93:183–8.
- Greenslade JH, Cullen L, Parsonage W, et al. Examining the signs and symptoms experienced by individuals with suspected acute coronary syndrome in the Asia-Pacific region: a prospective observational study. Ann Emerg Med 2012;60:777–85.
- Rana A, de Souza RJ, Kandasamy S, et al. Cardiovascular risk among South Asians living in Canada: a systematic review and metaanalysis. CMAJ Open 2014;2:E183–91.
- Khan NA, Grubisic M, Hemmelgarn B, et al. Outcomes after acute myocardial infarction in South Asian, Chinese, and white patients. *Circulation* 2010;122:1570–7.
- Zaman MJ, Philipson P, Chen R, et al. South Asians and coronary disease: is there discordance between effects on incidence and prognosis? *Heart* 2013;99:729–36.

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- Jin K, Ding D, Gullick J, et al. A Chinese immigrant paradox? Low coronary heart disease incidence but higher short-term mortality in western-dwelling chinese immigrants: a systematic review and metaanalysis. J Am Heart Assoc 2015;4:e002568.
- Moran A, Gu D, Zhao D, et al. Future cardiovascular disease in China: markov model and risk factor scenario projections from the coronary heart disease policy model-china. Circ Cardiovasc Qual Outcomes 2010;3:2432–52.
- Canada S. Immigration and Ethnocultural Diversity in Canada. National Household Survey, Catalogue no. 99-010-X2011001. 2011. Available at www12.statcan.gc.ca/nhs-enm/2011/as-sa/99-010-x/ 99-010-x2011001-eng.pdf (Accessed 8 Mar 2016).
- King KM, Khan N, Leblanc P, et al. Examining and establishing translational and conceptual equivalence of survey questionnaires for a multi-ethnic, multi-language study. J Adv Nurs 2011;67:2267–74.
- King KM, Khan NA, Quan H. Ethnic variation in acute myocardial infarction presentation and access to care. *Am J Cardiol* 2009;103:1368–73.
- King-Shier KM, Singh S, LeBlanc P, et al. The influence of ethnicity and gender on navigating an acute coronary syndrome event. Eur J Cardiovasc Nurs 2015;14:240–7.
- Milner KA, Vaccarino V, Arnold AL, et al. Gender and age differences in chief complaints of acute myocardial infarction (Worcester Heart Attack Study). Am J Cardiol 2004;93:606–8.
- Sundararajan V, Henderson T, Perry C, *et al.* New ICD-10 version of the Charlson comorbidity index predicted in-hospital mortality. *J Clin Epidemiol* 2004;57:1288–94.
- Rahim-Williams B, Riley JL, Williams AK, et al. A quantitative review of ethnic group differences in experimental pain response: do biology, psychology, and culture matter? Pain Med 2012;13:522–40.

- 24. Campbell CM, Edwards RR. Ethnic differences in pain and pain management. *Pain Manag* 2012;2:219–30.
- Rowell LN, Mechlin B, Ji E, et al. Asians differ from non-Hispanic Whites in experimental pain sensitivity. Eur J Pain 2011;15:764–71.
- Chan C-W, Lopez V, Chung JWY. Chest pain description and recognition among Chinese people with coronary problems. *World Crit Care Nurs* 2008;6:66–8.
- 27. Office of the Surgeon General (US); Center for Mental Health Services (US); National Institute of Mental Health (US). *Mental health: culture, race, and ethnicity: a supplement to mental health: a report of the surgeon general*. Rockville (MD): Substance Abuse and Mental Health Services Administration (US), 2001.
- Kleinman AM. Depression, somatization and the "new cross-cultural psychiatry". Soc Sci Med 1977;11:3–9.
- 29. Galanti G-A. *Caring for patients from different cultures.* 5th edn. Pittsburgh, PA: University of Pennsylvania Press, 2015.
- Nayak S, Shiflett SC, Eshun S, et al. Culture and gender effects in pain beliefs and the prediction of pain tolerance. Cross-Cultur Res 2000;34:135–51.
- Ting HH, Bradley EH, Wang Y, et al. Delay in presentation and reperfusion therapy in ST-elevation myocardial infarction. Am J Med 2008;121:316–23.
- 32. Bradley EH, Herrin J, Wang Y, *et al.* Racial and ethnic differences in time to acute reperfusion therapy for patients hospitalized with myocardial infarction. *JAMA* 2004;292:1563–72.
- Vaccarino V, Rathore SS, Wenger NK, et al. Sex and racial differences in the management of acute myocardial infarction, 1994 through 2002. N Engl J Med 2005;353:671–82.