# RESEARCH ARTICLE

# Evaluation of a cardiac troponin process flow at the chest pain center with the shortest turnaround time

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### Abstract

**Background:** Early diagnosis of myocardial infarction is crucial in chest pain management and cardiac troponin (cTn) test is an important step in it. Process improvement to shorten the test turnaround time (TAT) may improve patients' outcomes. The cTn test at chest pain center (CPC) of Zhongshan Hospital had the shortest TAT ever reported, but its process flow was not fully evaluated.

**Methods:** We performed a stepwise evaluation of CPC cTn TAT and explored the potential factor that might cause delay. The performance of CPC cTn test was also compared with cTn test and human chorionic gonadotropin (HCG) test ordered from emergency department (ED).

**Results:** At least 95% of CPC cTn tests were completed in 60 min, while 62% in 30 min. The medians of monthly order-to-collect time, collect-to-received time, and receivedto-result time were ~7 min, ~3 min, and ~13 min, respectively. The samples collected at the bedside had longer collect-to-received time than the ones collected at the blood draw site next to the laboratory. Compared to ED cTn test and ED HCG test, CPC cTn test took less time in each step. A combination of the sample type switch and the centrifugation time reduction contributed the most to the shortening of TAT, which was reflected in the received-to-result time.

**Conclusions:** The current process flow of CPC cTn test satisfied the requirements of chest pain management, giving an example of how to implement process improvement for emergency medicine to shorten TAT of laboratory tests.

Hao Wang and Xinyue Wang contributed equally to this work.

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### KEYWORDS

cardiac troponin, emergency medicine, myocardial infarction, process improvement, turnaround time

# 1 | INTRODUCTION

Ischemic injury is a key cause of death and other adverse events in patients suffering myocardial infraction (MI).<sup>1-4</sup> Reopening the obstructed artery via percutaneous coronary intervention (PCI) may effectively eliminate ischemia.<sup>2</sup> A lot of efforts have been made to shorten the time from the patient seeking emergency care to inflation of the catheter balloon, which is also called door-to-balloon time, and duration of ischemia.<sup>3</sup> One of them is the establishment of chest pain centers (CPCs), where specified processes are implemented to facilitate diagnosis and treatment for MI patients.<sup>1</sup> In China, the establishment and accreditation of CPCs have started since 2011.

An accurate diagnosis of MI, which includes evaluation of cardiac biomarker, is the rate-limiting step in the management of chest pain.<sup>5-8</sup> Nowadays, cardiac troponin (cTn) has become the primary cardiac biomarker in the management of MI and the central in the definition of non-ST-elevation MI (NSTEMI).<sup>4,9-11</sup> The recommended cTn delivery turnaround time (TAT) at a CPC or an emergency department (ED) for chest pain is 60 min, to offer timely treatments to NSTEMI patients and to minimize door-to-balloon time, which is essential for the very high-risk patients.<sup>4,12</sup> Meanwhile, a short cTn TAT also helps to minimize patient length of stay at ED and to ease crowding.<sup>10</sup> However, it is a challenge for a clinical laboratory to achieve the TAT goal.

Although point-of-care testing, which may reduce assay TAT, is a recommended replacement, its reliability and expense still could not satisfy clinicians, and laboratory testing remains the preferred option.<sup>4-8</sup> Therefore, several studies have investigated how process improvement may help to reduce laboratory TAT for cTn tests.<sup>5-8</sup> Their improvements included barcoding, floorplan design, sample processing priority and so on, and the shortest monthly order-to-result time (median) is around 50 min reportedly. At Zhongshan Hospital, the establishment of CPC was completed in 2017, and previous studies were used for reference in the determination of the laboratory testing processes for cTn. A monthly review in 2020 showed that most of CPC cTn tests at Zhongshan Hospital were reported to patients in 50 min, which was much better than previous reports, meeting the requirements in the clinical guidance documents.<sup>4,13</sup>

In this study, we made a systematic evaluation of CPC cTn TAT at Zhongshan Hospital. The performance of CPC cTn was also compared to that of ED cTn, which was ordered for a heart disease patient without chest pain, and ED human chorionic gonadotropin (HCG), which had a similar process flow with previous cTn (before establishment of CPC), to investigate how process improvements shortened CPC cTn TAT. The findings may give ideas on the further improvements of laboratory testing and chest pain management processes, for both Zhongshan Hospital and other medical institutions.

# 2 | MATERIALS AND METHODS

### 2.1 | The establishment of CPC

Before 2017, the patients with chest pain were admitted at ED of Zhongshan Hospital and treated following similar process flows of other patients. In 2017, to ensure that a patient suffering from a heart attack could be quickly identified and admitted for further services, Zhongshan Hospital established CPC based on ED. CPC and ED shared the spaces, instruments and laboratory. However, CPC only took care of individuals with chest pain and, had specified instruments and procedures for cTn test to reduce TAT. CPC of Zhongshan Hospital provided services since 2018.

The final process flow of CPC cTn test was a combination of stepwise process improvements. The improvements included (1) waiving physician assessment, (2) prioritizing order module, (3) waiving prior payment, (4) increasing blood sampling priority, (5) changing the sample type, (6) receiving samples immediately, (7) using an individual centrifuge, (8) shortening centrifugation time, (9) applying automatic numbering, (10) using rapid testing reagents, and (11) applying autoverification, which could be explained by a comparison among CPC cTn test, ED cTn test, ED HCG test, and previous cTn test (Table 1).

### 2.2 | Study design

This study was to evaluate the cTn TAT at CPC after around 2 years of operation and identify opportunities for further improvement. The order-to-result time of CPC cTn was a key metric in evaluating TAT. Based on the order-to-result time of each CPC cTn sample, we calculated and evaluated the monthly on-time percentages. The evaluation also included stepwise TAT analysis and comparison with tests at ED. Zhongshan Hospital Research Ethics Committee approved this study (Ethics certificate number, B2021-524R) and waived informed consents from patients.

### 2.3 | Laboratory settings

Emergency department of Zhongshan Hospital had its own laboratory, where CPC tests were also performed, and its patient blood draw site was next to the laboratory. The entrance, cashier, patient blood draw site, and laboratory were all on the ground floor of ED. All the immunoassay tests, including cTn and HCG, were performed on Roche E411 analyzer (made in Germany). CPC cTn test had an individual centrifuge, whereas ED cTn and HCG tests shared a centrifuge with other tests. CPC and ED cTn samples were analyzed using 9-min electrochemiluminescence cardiac troponin T (cTnT) reagents. TABLE 1 Stepwise process improvement

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Previous cTn	CPC cTn	ED cTn	ED HCG						
Door-to-order (Step 1, not included in the evaluation)									
Ordered by a physician after assessment	Ordered by a nurse or physician without assessment	Ordered by a physician after assessment	Ordered by a physician after assessment						
Routine order module in the workstation	Prioritized order module for cTn in the workstation	Routine order module in the workstation	Routine order module in the workstation						
Order-to-collect (Step 2)									
Payment for test	No payment required before blood sampling	Payment for test	Payment for test						
Waiting for blood draw	Priority for blood sampling	Waiting for blood draw	Waiting for blood draw						
Collect-to-received (Step 3)									
5-ml gold-cap tube for serum	4-ml green-cap tube for plasma	4.5-ml light green-cap tube for plasma	4.5-ml light green-cap tube for plasma						
Queueing for received	Directly received	Queueing for received	Queueing for received						
Received-to-result (Step 4)									
Standing still for 10 mins	No standing for clotting	No standing for clotting	No standing for clotting						
Shared centrifuge	Centrifuge for CPC cTn only	Shared centrifuge	Shared centrifuge						
10-min centrifugation	1-min centrifugation	10-min centrifugation	10-min centrifugation						
Manual numbering after centrifugation	Automatic numbering when received	Manual numbering after centrifugation	Manual numbering after centrifugation						
18-min testing	9-min rapid testing	9-min rapid testing	18-min testing						
Manual verification	Auto-verification	Auto-verification	Auto-verification						

Meanwhile, ED HCG samples were analyzed using 18-min electrochemiluminescence HCG reagents.

# 2.4 | Data collection and analysis

# 2.4.1 | Data

The order, collection, received, and result time of each sample were stored at the Network Center of Zhongshan Hospital. Time data generated from CPC cTn test, ED cTn test, and ED HCG test between February 2018 and May 2020 were retrieved. Samples were grouped by month and, order-to-collect time, collect-to-received time, and received-to-result time of each sample were calculated. According to order-to-collect time, collect-to-received time, and received-to-result time, samples with outlier values were eliminated.

# 2.4.2 | On-time percentage

The order-to-result time of each CPC cTn sample was calculated. Four standards, 30, 40, 50, and 60 min, were used to evaluate the monthly order-to-result on-time percentages. The correlation among monthly on-time percentages and sample size was also evaluated.

# 2.4.3 | Stepwise CPC cTn TAT

Monthly data distribution of order-to-collect time, collect-to-result time, collect-to-received time, and received-to-result time was analyzed.

### 2.4.4 | Identification of the delay-causing factor

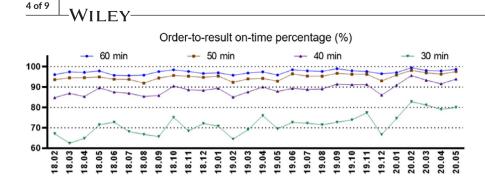
Since monthly data distribution of collect-to-received time appeared as bimodal shapes in the diagram, a cut-off line, showing the trough between two peaks, was determined. The samples were divided into two groups according to the cut-off (samples without collection location information were not included in the analysis). The monthly bedside sample amount and lab sample (sample collected at the patient blood draw site next to the laboratory) amount was compared between the two groups.

### 2.4.5 | Effects of the delay-causing factor

The samples were divided into laboratory samples and bedside samples. The collect-to-received time and order-to-collect time were compared between the two sample groups.

FIGURE 1 The order-to-result on-time

percentages at different standards



# 2.4.6 | Stepwise evaluation of process improvements

The monthly order-to-collect time, collect-to-received time and received-to-result time of cTn (it also used rapid testing reagents, but ED process flow) and HCG (its process flow and reagent types were similar to those of previous cTn) at ED were compared with those of cTn at CPC, respectively.

# 2.5 | Statistics

The diagram creation and outlier value identification were performed using GraphPad Prism software. The correlation among monthly ontime percentages and sample size was analyzed using Spearman's correlation efficient. Categorical variables were analyzed using the Chi-square test. A *p*-value smaller than 0.05 was considered as statistically significant.

### 3 | RESULTS

# 3.1 | An overview of CPC cTn TAT

Twenty thousand, one hundred forty-three samples were included in this study. The smallest monthly sample size was 389, and the largest one was 1274. For each month, at least 95% of samples were completed (order-to-result) in 60 min, 91% in 50 min, 84% in 40 min, and 62% in 30 min (Figure 1). In another word, the median of monthly order-to-result time of CPC cTn was consistently below 30 min. The monthly sample size curve did not coincide with each monthly ontime percentage curve and on-time percentages were not correlated with sample size, suggesting that patient volume didn't challenge the CPC cTn testing performance (Figure S1).

### 3.2 | Time cost in each step

According to the patient involvement, the order-to-result process could be divided into the patient-dependent order-to-collect step and the patient-independent collect-to-result step. The median of monthly order-to-collect time ranged from 4.55 min to 8.09 min, whereas the median of monthly collect-to-result time ranged from 16.53 min to 17.95 min (Figure 2). Interestingly, compared to the order-to-collect time, whose distribution was symmetric or near symmetric, the collect-to-result time had data distribution of an overt bimodal shape, indicating that there were some factors separating samples into two distinct groups and prolonging the collect-to-report time (Figure 2).

To further investigate whether the collect-to-received step or the received-to-report step contributed to the prolonged collect-toreport time, their data distributions were analyzed. More obvious bimodal shapes appeared in the diagram for collect-to-received time, rather than that for received-to-result time (Figure 3). The cut-off line, which showed the trough between two peaks, represented 140 s (Figure 3). These results suggested that factors in the collectto-received step separated samples.

### 3.3 | Sampling location factor in CPC cTn test

The collect-to-received time was the sample transportation time. Since some chest pain patients were not able to walk to the patient blood draw site next to the laboratory, nurses had to draw their blood at the bedside and transport samples to the lab, which took a longer time. According to the collect-to-received time, samples were divided into two groups, above the cut-off line and below the cut-off line. The chi-square test results indicated that most of the patients with a prolonged collect-to-received time were sampled at the bedside (Table 2).

The samples collected at the bedside were compared to those collected next to the laboratory (lab) via analyzing the location-specific collect-to-received time. In either group only one peak was detected in the distribution of each monthly collect-to-received time, and the medians of bedside-specific collect-to-received time were 3–4 min longer (Figure 4, top). The medians of bedside-specific order-to-collect time were also, to some extent, longer, probably because of material preparation for blood sampling at the bedside (Figure 4, bottom).

### 3.4 Comparison with ED tests

The order-to-collect time for CPC cTn test was the shortest among 3 tests, indicating that removal of payment and waiting for blood

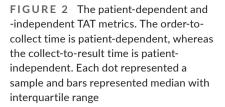
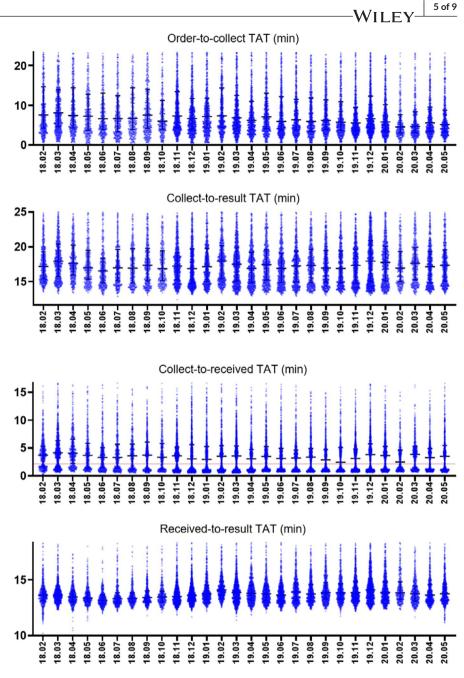


FIGURE 3 The location-dependent and -independent TAT metrics. The collect-to-received time is locationdependent, whereas the received-toresult time is location-independent. Each dot represented a sample and bars represented median with interquartile range. A red line showed the trough between two peaks



draw helped (Figure 5, top). The maximum difference of median was 4.86 min in September 2019 (Figure 5, top).

Compared to ED HCG test, whose samples were mostly collected at the patient blood draw site, CPC cTn test had a reduced collectto-received time (reduction ranged from 1.14 min to 7.66 min), showing the importance of immediate reception (Figure 5, middle). Meanwhile, the collect-to-received time for ED cTn test was much longer than that for ED HCG test (8.80–12.92 min longer), which was probably caused by more samples collected at the bedside or longer transportation time (Figure 5, middle).

The most significant reduction occurred in the received-to-result time. The differences between ED cTn and ED HCG tests ranged from 3.33 min to 8.97 min, showing the contribution of 9-min rapid testing reagents (Figure 5, bottom). The maximum reduction from ED cTn test to CPC cTn test was 25.00 min, while the minimum was 20.29 min, indicating the importance of sample type change, short centrifugation time and automatic numbering (Figure 5, bottom). The sample type change eliminated 10-min standing for clotting (Table 1). A smaller sample volume of the E411 analyzer allowed 1-min centrifugation to generate enough plasma (on the top layer) for instrumental analysis. In addition, the specified centrifuge and automatic number significantly reduced the waiting time.

# 4 | DISCUSSION

The reduction of cTn TAT is always an important goal to achieve in emergency medicine. With the clinical application of high-sensitivity

Month	Location	Total	<140s	≥140s	p Value	Month	Location	Total	<140s	≥140s	p Value
18.02	Lab	314	216	98	<0.0001	19.04	Lab	374	359	15	< 0.0001
	Bedside	382	10	372			Bedside	429	10	419	
18.03	Lab	228	219	9	<0.0001	19.05	Lab	286	272	14	<0.0001
	Bedside	591	18	573			Bedside	470	23	447	
18.04	Lab	132	122	10	<0.0001	19.06	Lab	354	332	22	<0.0001
	Bedside	322	5	317			Bedside	436	12	424	
18.05	Lab	149	141	8	<0.0001	19.07	Lab	332	319	13	<0.0001
	Bedside	278	9	269			Bedside	446	11	435	
18.06	Lab	181	169	12	<0.0001	19.08	Lab	265	256	9	< 0.0001
	Bedside	291	17	274			Bedside	420	16	404	
18.07	Lab	184	174	10	<0.0001	19.09	Lab	349	336	13	<0.0001
	Bedside	294	7	287			Bedside	360	10	350	
18.08	Lab	153	148	5	<0.0001	19.10	Lab	348	332	16	< 0.0001
	Bedside	255	10	245			Bedside	352	12	340	
18.09	Lab	183	176	7	<0.0001	19.11	Lab	447	429	18	<0.0001
	Bedside	348	16	332			Bedside	512	14	498	
18.10	Lab	150	141	9	<0.0001	19.12	Lab	537	520	17	<0.0001
	Bedside	239	8	231			Bedside	712	14	698	
18.11	Lab	326	317	9	<0.0001	20.01	Lab	352	337	15	<0.0001
	Bedside	649	20	629			Bedside	530	18	512	
18.12	Lab	462	449	13	<0.0001	20.02	Lab	192	183	9	<0.0001
	Bedside	559	19	540			Bedside	194	7	187	
19.01	Lab	363	351	12	<0.0001	20.03	Lab	185	176	9	<0.0001
	Bedside	435	13	422			Bedside	339	10	329	
19.02	Lab	313	302	11	<0.0001	20.04	Lab	301	296	5	< 0.0001
	Bedside	450	16	434			Bedside	406	19	387	
19.03	Lab	413	407	6	<0.0001	20.05	Lab	305	295	10	<0.0001
	Bedside	646	18	628			Bedside	390	8	382	

TABLE 2 Location factor of prolonged collect-to-received time

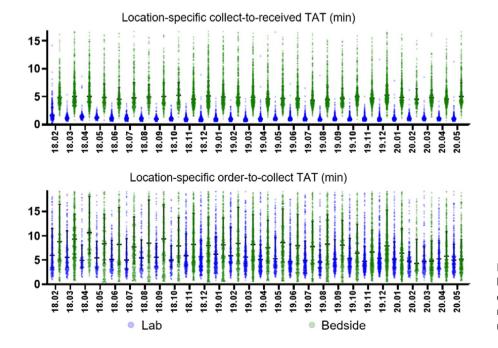
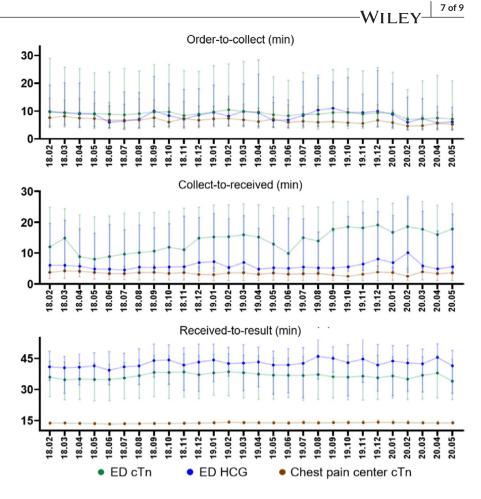


FIGURE 4 The contribution of location factor to TAT metrics. Each dot represented a sample and bars represented median with interquartile range

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FIGURE 5 The comparison of cTn TAT metrics at the chest pain center with cTn and HCG TAT metrics at ED. Each dot represented median and bars represented interquartile range



cTn tests, more and more strategies have been developed to maximize the value of cTn testing. In 2015 European Society of Cardiology guidelines, a 0/1-h algorithm of cTn results was recommended for earlier diagnosis of NSTEMI.<sup>14</sup> Similar algorithms were proposed by several other studies and, in all these studies, the increase of cTn results between two sequential tests may give a hint of NSTEMI earlier than a traditional one-time cut-off.<sup>9,11,14-17</sup> In principle, a patient taking two serial cTn tests should stay at ED/CPC for 2 h with a cTn TAT of 60 min, whereas the length of stay is 1.5 h with a TAT of 30 min. The 25% reduction of the length of stay indicates that the use of cTn algorithms adds an economic value to the time control of cTn testing.<sup>10,18</sup>

Lean methodology is a way to optimize the efficiency of a system via eliminating the unnecessary waste of people, resources, efforts and so on. During the establishment of CPC at Zhongshan Hospital, the process flow of cTn testing was determined according to the principles of lean methodology, thereby effectively controlling the TAT.<sup>5-7</sup> In this study, a stepwise evaluation gives us an insight into the performance of each step and opportunities for further improvement. From more to less, the time was spent on the received-to-result step, the order-to-collect step and the collect-to-received step successively, which is consistent with other studies (Figures 2 and 3).<sup>6,7</sup> Meanwhile, the order-to-collect step had the widest quartile range, while the received-to-result step had the narrowest one, indicating that the sample analysis at the laboratory

was optimized significantly and the order-to-collect step should be further explored. Considering the entrance and the patient blood draw site were on the same floor and the walk time between them was about 2 min, a possible explanation for the delayed sample collection is that patients wasted some time in finding the blood draw site. Another possibility is that some patients took electrocardiography, another key examination for MI, before blood draw. More signposts and education for patients may help to improve this situation. The location factor identified in this study suggests that the floorplan of ED still could be optimized and a short distance between the patient observation room and the laboratory might reduce the transportation time for bedside samples (Figure 4 and Table 2).

A key measure to shorten cTn TAT at CPC of Zhongshan Hospital is the combination of a sample type switch and a centrifugation time reduction. Since the manufacturer informed that, for its cTn tests, plasma had similar performance with serum in early diagnosis of MI, we switched the sample type to plasma to eliminate 10-min clotting time (Table 1). Moreover, considering the E411 analyzer draws the plasma from the top and the required sample volume is 50  $\mu$ l, onemin centrifugation could remove most of the blood cells on the top layer and generate a sufficient volume of plasma for the instrumental analysis. As a result, the queuing and processing time for centrifugation decreased from 10-20 min to 1-2 min, without clinical concerns on the assay performance. A previous study has suggested <sup>8 of 9</sup> WILEY

the use of whole blood and a hematocrit-based conversion equation to remove the centrifugation step.<sup>19</sup> However, whole blood is not a recommended sample type and our procedure is more suitable for the E411 analyzer.

There were still some limitations in this study. Firstly, the process improvements were implemented simultaneously, rather than progressively. Hence, we could not evaluate the process improvements step by step to find out the valueless attempts. Secondly, the time data for cTn test before 2017 could not be retrieved and the replacement, ED HCG test, did not fully reflected its performance. Thirdly, the contributions of sampling location factor were influenced by floorplan and the floorplan for each hospital was unique. All these should be investigated in further studies.

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### CONFLICT OF INTEREST

All authors report no conflict of interest.

### AUTHOR CONTRIBUTIONS

HW, XW, KW, and WG conceptualized and designed the study. KW and WJ acquired the data. HW, XW, KW, XD, and BP analyzed and interpreted the data. HW and XW drafted the manuscript. HW, BT, BP, BW, and WG revised the manuscript. HW, BW, BP, and WG acquired funding.

### DATA AVAILABILITY STATEMENT

All the data are included in this manuscript.

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# SUPPORTING INFORMATION

Additional supporting information may be found in the online version of the article at the publisher's website.

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