IMPORTANCE OF ACUTE CARDIAC CARE REGISTRIES AT THE NATIONAL LEVEL

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SUMMARY - Improving organization and patient care quality in intensive care units is increasingly important as intensive care unit diagnostic and therapeutic procedures account for a growing proportion of hospital services. We identified the lack of comprehensive national and international registries available in the contemporary literature. This paper aims to describe and analyze cardiac intensive care unit (CICU) network at the national level in Croatia and its comparison with more developed countries. Thirty-four representatives from all Croatian acute hospitals (response rate of 100%) filled in a web based questionnaire on CICU organization and competence during September and October 2016. Organization and available technical procedures for health care in general, and especially in very expensive CICU treatment, highly depends on gross domestic product (GDP) per capita. That is why one could expect that Croatia, with the second lowest GDP among European Union countries and 4.7 CICU per million inhabitants will have worse results in this field in comparison with most of these countries. Results such as one nurse responsible for a mean of 2.7 CICU patients, 52% of cardiologists among physicians during working hours but 37% during night shifts, 24/7 transesophageal echocardiography in only 26.5% of CICUs, one-third without therapeutic hypothermia, and 23.5% without extracorporeal membrane oxygenation treatment are some of these results, revealing much room for improvement. This representative, nationwide sample of Croatian CICUs also demonstrated considerable variation of key elements of structures with respect to hospital size, academic status and financial issues, as well as a trend towards current guidelines. This kind of investigation is very important for proposing standards, reimbursement master plan, or quality assessment of the national health system.

Key words: Intensive cardiac care; Cardiology; Registry; Heart; Questionnaire

Introduction

Intensive care unit (ICU) and cardiac intensive care unit (CICU) are dedicated to critical medical care, defined as the diagnosis and management of acute lifethreatening conditions treated by specifically trained health professionals. Two major advances in management followed by distinctive changes in the organization of CICU have been described¹, and attributed to early invasive strategy for treating acute coronary syndromes, but also rapid aging of the population. In the next near half-decade, another paper recognized an increased demand for cardiovascular critical care². A meta-analysis of 13 non-randomized studies showed that ICUs managed by intensive care specialists provided superior outcomes, resulting in a reduced length of stay³, reduced length of mechanical ventilation⁴ and associated costs, and finally, reduced mortality⁵⁻⁷. In these studies, mandatory intensivist consultation or a closed ICU was superior to low-intensity staffing, defined as no intensivist or elective intensivist consultation⁸. Also, adding a trained intensivist in a community teaching hospital was temporally associated with improved clinical and educational outcomes in that type of hospital⁹.

According to Morrow *et al.*², we have been witnessing evolution of CICU from point of rapid resuscita-

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Fig. 1. Gross domestic product per capita (US dollars) 2014 or latest year, ESC member countries. ESC Atlas of Cardiology based on data from the World Bank (WB); http://data.worldbank.org/indicator.

tion to preventive intervention, and finally comprehensive critical care. The same authors made a proposal for categorization of CICU, ranging from level 1 as the highest to level 3 as the lowest category. A level 1 CICU has all forms of invasive and noninvasive monitoring capabilities, and advanced technologies, management of patients with refractory shock or resuscitated cardiac arrest by full-time intensivists, either cardiac intensivists or general intensivists working collaboratively with cardiologists, nurse-to-patient ratios 1:1 or 1:2, and access to interventional cardiology and cardiac surgical support. A level 1 CICU would be the likely paradigm for most large tertiary medical centers. A level 2 CICU is capable of providing initial evaluation and management of most acute cardiovascular conditions, all invasive and noninvasive monitoring is available, but mechanical hemodynamic support is limited to non-ventricular assist devices. Management



Fig. 2. Change in cardiovascular disease crude mortality rates between 2010 and latest year available among females and males aged <70 years in ESC member countries vs. GDP per capita (US\$, 2017). According to European Society of Cardiology: Cardiovascular Disease Statistics 2019 European Heart Journal, ebz859, https:// doi.org/10.1093/eurheartj/ebz859.

is generally by cardiologist, but the unit may be combined with a general medical or surgical ICU, and nurse-to-patient ratios are usually 1:1 to 1:3. A level 3 CICU has capacity to manage respiratory failure, administer vasopressors and inotropes for hypotension but may be focused on the care of patients with suspected acute coronary syndrome, heart failure without shock, and hemodynamically stable arrhythmias. A level 3, like level 2, may also be combined with a general medical ICU. Invasive monitoring and mechanical hemodynamic support are not usually provided in this unit, and nurse-to-patient ratios are usually 1:2 to 1:3.

In summary, critical care costs are expected only to rise because of the development of new technologies, including complex modes of mechanical ventilation, invasive hemodynamic monitoring tools, induction of therapeutic hypothermia, renal replacement therapies, bedside vascular procedures with imaging guidance, and mechanical circulatory support. New technologies are also an answer to the rising prevalence of non-cardiovascular diagnoses including sepsis, acute kidney injury, and respiratory failure¹⁰. ICUs have been a center of interest concerning quality assessment, given the high mortality rate, potential for iatrogenic complications, and increased resource utilization. Quality assessment initiatives in the ICU have traditionally focused on outcomes, particularly on mortality rates. Nevertheless, in order to investigate outcomes, it is crucial to implement evaluation of structure (organization), and process (health care delivery), and this was the main reason for us to engage in this research. Surveys with questionnaires play a crucial role in policy making¹¹, in addition to the states described by Markov processes.

The gross domestic product (GDP) *per capita* is one of the standard indicators of financial development. There is considerable diversity of national resources across Europe; GDP *per capita* ranges from <\$4000 in some Eastern European countries to >\$40000 in the West¹² (Fig. 1). Political and organizational aspects of health care show similar variation, the net result being huge disparities in the population cardiovascular disease (CVD) risks and the resources available for prevention and treatment.

Acute cardiac care conditions in Croatia are usually diagnosed in out-of-hospital setting by emergency doctors or less often by general practitioners, and in hospital emergency departments and ambulances. All patients with acute cardiac conditions are urgently hospitalized in the nearest hospital. The Croatian primary percutaneous coronary intervention (PCI) network was introduced earlier^{13,14}, and showed a growing trend of 540-550 patients with acute ST-segment elevation myocardial infarction (STEMI) per million inhabitants. Each of the eleven PCI centers in Croatia with 24/7 work schedule had a catchment population of 384,000 inhabitants, approaching the number of at least 600 primary PCIs per million inhabitants. On the contrary, we found that only 380-390 non-ST-segment elevation myocardial infarction (NSTEMI) patients per million inhabitants received PCI in the first 72 hours¹⁵. Croatia has achieved a decreasing mortality trend in coronary heart disease, as well as in the overall cardiovascular pathology in the past ten years (53% vs. 47%). Even though we have witnessed significant improvement in cardiovascular mortality in the previous

decade, Croatia is still positioned among European countries with high cardiovascular mortality. According to the newest European Society of Cardiology: Cardiovascular Disease Statistics 2019 that includes trends in CVD crude mortality rates between 2010 and the latest year available, there still is room for significant improvement (Fig. 2). Acute cardiac care conditions are reimbursed by the National Health System based on obligatory health insurance of all employed persons, but also retired and unemployed (the latter paid by state). Quoted *per* patient reimbursement with a system of flat rate packages built from pre-established lists of pathologies, technical interventions and some comorbidities are also used in Croatia. Croatia belongs to the group of transition countries, and to the best of our knowledge, observational studies involving organization of acute cardiac care have not been systematically reviewed in this kind of setting. We compared CICU and ICU of this economically less developed European country with developed European countries.

Patients, Materials and Methods

During September and October 2016, we conducted an online questionnaire involving all 34 Croatian acute hospitals. We promoted and distributed a national web-based survey in 34 CICUs, either strictly cardiac ICU or combined with other internal medicine fields or anesthesiology ICU. We developed two types of questionnaire depending on the type of ward (only cardiac/internal medicine patients or combined cardiac/internal medicine/anesthesiology unit). The first type of questionnaire contained 17 items and the second one contained 15 items. Questions were either binary or multiple-choice. In addition, there was a possibility of making comments at the end of the questionnaire. E-mail addresses were obtained from the National Cardiac Care Society and the questionnaire was sent to chief physicians of CICUs in Croatia. Percentages reported are proportions of responses and Fisher exact test was used to calculate p-values for categorical data. Statistical significance was set at p<0.05.

Results

Our investigation included all Croatian acute hospitals (local, regional and tertiary level hospitals) and



Fig. 3. Types of intensive care units.



Fig. 4. Number of cardiac intensive care unit beds.



Among services that were consistently used were primary PCI, where all acute hospitals urgently transferred STEMI/high risk NSTEMI patients to PCI centers, or performed PCI themselves. Additionally, 50% of patients requiring intra-aortic balloon pump





Fig. 6. Number of beds per nurse and physician during working hours.

Fig. 5. Percentage of specialists during working hours and night shifts.

were transferred to other hospitals. Dialysis was performed in 90% of investigated units, in 29% of them without help from nephrologists. Mechanical ventilation was performed in almost 60% of centers without help from other specialists, and in only one hospital, these patients were transferred to another hospital (Fig. 9). With regard to other specific forms of intensive treatment, Croatian hospitals showed large differences. Even though more than one-fourth performed therapeutic hypothermia alone or with the help of other specialists, one-third of Croatian centers did not implement hypothermia, and other centers transferred patients to other hospitals or other departments in their hospitals. Extracorporeal membrane oxygenation (ECMO) is predominantly a multidisciplinary proce-



Fig. 7. Total number of admissions per year.

dure, in Croatia reserved for large academic hospitals, with two exceptions.

Discussion

According to this and other investigations¹⁵, there is a respectable access to tertiary hospitals for urgent cardiac patients in Croatia. Tertiary hospitals are distributed across all Croatian regions and coupled with the nearest regional and county hospitals in the organizational network. Local policies on referral and selection criteria generally do not differ among Croatian regions and reflect concordance with clinical guidelines. In order to take a more realistic view on the state-of-the-art, comparison with other countries according to development of economy is crucial. Many



Fig. 8. Transthoracic echocardiography (TTE) and transesophageal echocardiography (TEE) availability.



Fig. 9. Technical procedures in acute cardiac care. ECMO = extracorporeal membrane oxygenation

studies suggest that outcome of ICU treated patient is related to the number of ICU beds or number of patients treated. An interesting retrospective cohort study demonstrated that both hospital volume of ICU patients and ICU to hospital bed ratio were significantly associated with mortality¹⁶. The number of ICU beds in two large Canadian jurisdictions is comparable to those found in this investigation, and so is the nurse to patient ratio¹⁷. Another international, multicenter, observational study found that a nurse to patient ratio of less than 1:1.5 was independently associated with a lower risk of in-hospital death¹⁸. This ratio was achieved in less than one-third of the ICUs investigated in this survey. A nationwide analysis of ICUs in Germany showed that the mean number of beds per ICU was 10.4, with significant differences depending on hospital size¹⁹. Our investigation revealed less sized ICUs on average. In a German study, 81% of the ICUs provided around-the-clock physician presence (range: 66%-98%). We identified 100% physician around-theclock presence, however, emphasizing that a significant percentage of non-cardiologists during night shifts is of utmost importance, reflecting low availability of cardiologists, especially in smaller hospitals. In addition, our results indicated physician heterogeneity during night shifts, which reflects the obvious need for a greater number of cardiologists. Shift-wise, one nurse was responsible for a mean of 2.7 patients (morning 1:2.3, afternoon 1:2.6, night 1:3.3 patients) with significant variation according to hospital size (smaller hospitals 1:2.9, university hospitals 1:2.1; p<0.001). Interesting results of a survey in Germany showed an increase in interdisciplinary ICUs (74%), and a divergent trend in human resource development with an increase of physicians and decrease of nurses²⁰. A European survey from 1998 found that at that time, 67% of the ICUs had between 6 and 12 beds, and 54% were mixed medical-surgical units. Among British units, 79% had more than three full-time nursing equivalents per ICU bed, while in Sweden 75% of units had less than two full-time nursing equivalents per ICU bed²¹.

Croatia has 4.7 CICUs *per* million inhabitants with a range of 1-9 (mostly 5-6) beds. A more precise and contemporary population-based cross-sectional study established the overall age- and gender-standardized number of ICU patients *per* 1000 personyears for the 5-year period. The result was 4.3 patients ranging from 3.7 to 5.1 patients *per* 1000 person-years in five regions of Denmark and from 2.8 to 23.1 patients *per* 1000 person-years in 98 municipalities²². We recognized the need for a similar study in Croatia for more precise comparison.

According to most studies, TTE was widely available in ICU, but in one study from Switzerland it was not routinely used, and in most cases, TTE was not performed by intensive care specialists²³. Our study showed that TTE was performed not only by intensivists but often by internal medicine specialists. By far most experience in the use of TEE in CICU stems from critical care units with a high proportion of cardiac, including coronary and/or cardiac surgical patients (53%). Given this heterogeneity of settings, it is not unusual to find the reported findings to vary considerably²⁴. To the best of our knowledge, although the present body of evidence supporting the use of TEE lacks prospective, randomized controlled studies, current evidence, as well as clinical experience points to the significant benefits gained by the availability of TEE in CICUs. To conclude, CICUs in Croatia with 26.5% around-the-clock availability emphasize the need for systematic training among physicians and staff.

According to Danish Heart Registry, the number of primary PCI during the last ten years ranged between 1200 and 1300 *per* million inhabitants²⁵, and there were 7650 coronary angiographies for NSTE-MI/unstable angina pectoris indication. The figure of almost 600 primary PCIs *per* million inhabitants positioned Croatia along the most developed European economies. On the contrary, with only 380-390 NSTEMI patients *per* million inhabitants receiving PCI in the first 72 hours, we recognize much room for improvement.

An Italian ICU survey from 2013 showed that therapeutic hypothermia was still underused (55.1%), however, with a significant increase in the adoption of hypothermia in the past 5 years²⁶. The use of target temperature management after cardiac arrest has been introduced in most of the German ICUs²⁷. These results do not correlate with ours (only 26.4%) and more efforts should be invested to increase the utility of this method.

Extracorporeal membrane oxygenation, either supporting gas exchange or hemodynamics in patients with severe respiratory or cardiac failure, has shown considerable evolution over the last decade, with a constant rise since 2009 in the number of ECMO-treated patients and number of centers providing ECMO support. In addition, a growing amount of evidence on treating such patients²⁸ provides better treatment and care.

Scottish adult cardiothoracic surgical services are provided by three ECMO regional centers covering a population of 5.2 million individuals and achieving acceptable functional recovery despite a significant complication rate²⁹. According to our best knowledge, there is only one study³⁰ to show recent epidemiological data of ECMO utilization and associated mortality in a high-income country. Although mortality decreased with increasing experience, it has remained high in both venovenous and venoarterial ECMO. In a high-income country such as Germany, the use of ECMO has been rapidly increasing since 2007 for both respiratory and cardiac support, from 0.1:100,000 in 2007 to 0.7:100,000 in 2012, and to a maximum of 3.5:100,000 in 2014 for cardiac support. Since the introduction of ECMO program in Croatia in 2008, there is a positive trend of its utilization rates, but according to our investigation data, 23.5% of ICUs do not use ECMO and do not transfer patients with indication for ECMO to other centers with the possibility of such treatment.

A questionnaire developed in adult ICUs in Switzerland found that overall utilization rate for noninvasive ventilation (NIV) was 26% of all mechanical ventilations, also showing that the preferred location of NIV initiation in the acute hospital was the ICU (86%), not the emergency department³¹. We showed that only one ICU did not implement this method.

Online survey in the UK revealed that a nephrologist was never or rarely consulted about patients with acute renal injury in over 40% of the units. Continuous renal replacement therapy was managed almost exclusively by intensivists, while intermittent hemodialysis was managed predominantly by nephrologists³². In our study, nine of ten ICUs provided hemodialysis and one of three without the help from nephrologists.

Future research should focus on additional elements of health care organization. Comprehensive staffing models evaluating all staff involved may be one approach. Another approach may be to evaluate other factors such as adherence to new evidence-based practices or processes (for example, therapeutic hypothermia, ECMO). If these procedures are evaluated as effective also in real-world studies and registries, then all hospitals should be able to implement similar models of treatment in daily routine. Structures in intensive care medicine comprise human, as well as material resources, organization and management, which are related to processes thereby affecting patient outcomes.

In conclusion, this representative, nationwide sample of Croatian CICUs demonstrated considerable variation of key elements of structures with respect to hospital size, but also a trend towards current guidelines. However, we can conclude that our clinical practice offers room for improvement in this field, which will be tightly associated with the country's financial capabilities. Results of this kind of investigation have to be taken into account when proposing standards, reimbursement plan, development, or quality assessment of CICU network.

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Sažetak

ZNAČENJE REGISTARA AKUTNE SKRBI KARDIOLOŠKIH BOLESNIKA NA NACIONALNOJ RAZINI

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Unaprjeđenje organizacije i kvalitete skrbi o bolesnicima u jedinicama intenzivne skrbi postaje sve važnije zbog rastućeg opsega dostupnih metoda liječenja i skrbi. Suvremena literatura ukazuje na nedostatak odgovarajućih nacionalnih i internacionalnih registara. Cilj ovog istraživanja bio je analiza organizacije jedinica intenzivne kardijalne skrbi (JIKS) u Hrvatskoj i usporedba s ekonomski razvijenijim zemljama. Liječnici iz 34 hrvatske akutne bolnice (100%-tni odgovor) ispunili su e-poštom poslan upitnik o organizaciji skrbi i metodama liječenja akutnih kardioloških bolesnika. Odgovori su prikupljeni i analizirani tijekom rujna i listopada 2016. godine. Hrvatska ima oko 5 JIKS na milijun stanovnika (raspon od 1 do 9, većinom 5-6 kreveta). Jedna medicinska sestra skrbi za prosječno 2,7 bolesnika (u jutarnjim satima za 2,3 bolesnika, u poslijepodnevnim satima 2,3 bolesnika, noću 3,3 bolesnika) uz varijabilnost ovisno o veličini bolnice (u manjim bolnicama prosječno za 2,9 bolesnika, u sveučilišnim bolnicama za 2,1 bolesnika, p<0,001). Gotovo dvije trećine [IKS sadrži manje od 4 kreveta na jednog liječnika, dok kardiolozi čine 52% liječnika tijekom radnog dana, ali samo 37% liječnika tijekom dežurstva. Utvrdili smo značajnu varijabilnost u dostupnosti ultrazvuka srca tijekom radnog dana u odnosu na dežurstvo (76,5% JIKS ima 24-satnu dostupnost transtorakalne ehokardiografije, ali samo 26,5% za transezofagusnu ehokardiografiju). Trećina ispitivanih centara nije uvela terapijsku hipotermiju, a 23,5% centara ne radi izvantjelesnu membransku oksigenaciju niti premješta bolesnike u odgovarajuće ustanove. Organizacija i dostupne metode liječenja u IIKS ovise i o bruto društvenom proizvodu. Ovo istraživanje, prvo takve vrste u Republici Hrvatskoj, ukazuje na značajnu varijabilnost ključnih dijelova zdravstvene skrbi akutnih kardioloških bolesnika ovisno o veličini bolnice, ali i ukupni trend prema postojećim smjernicama. Ukazujući na prostor za napredak, ono može poslužiti kao polazna točka u postizanju željenog standarda, planiranju financija te procjeni i praćenju kvalitete nacionalnog zdravstvenog sustava.

Ključne riječi: Intenzivna kardiološka skrb; Kardiologija; Registar; Srce; Upitnik