An Asian Perspective of the Management of COVID-19: the Asian National Cancer Centers Alliance Led Regional Comparison

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Objective: To describe how the Asian National Cancer Centers Alliance (ANCCA) members preserve high standards of care for cancer patients while battling the COVID-19 pandemic and to propose new strategies in the Asian Cancer Centers' preparedness to future pandemics.

Methods: A 41-question-based survey was developed using an online survey tool and conducted among 15 major Asian National Cancer Centers, including 13 ANCCA members. Direct interviews of several specialists were conducted subsequently to obtain additional answers to key questions that emerged during the survey analysis.

Result: Institution/country-specific results provided a strong insight on the diverse ways of managing the pandemic around Asia, while maintaining well-balanced cancer care. Pragmatic strategies were put in place in each NCC hospital, including zoning and intensive triage depending on the pandemic impact. Distancing strategies and telemedicine were implemented

in different capacity depending on the national healthcare system. In addition, there was a diverse impact on the manpower and financial aspect of cancer care across surveyed NCCs relating to magnitude of the pandemic impact on the country.

Conclusion: The priorities nevertheless remain on maintaining cancer care delivery while protecting both patients and health care workers from the risk of COVID-19 infection. The role of a think-tank such as ANCCA to help share experiences in a timely manner can enhance preparedness in future pandemic scenarios.

Introduction

The current coronavirus pandemic has created a global crisis. The severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) virus, which caused Coronavirus disease 2019 (COVID-19), is a novel coronavirus first detected in Wuhan, China in December, 2019 [1]. As of 13-July 2020, over 570,000 people have died from COVID-19 related disease with more than 13 million people being infected globally. Of them, 3 million cases and over 70,000 deaths have been described in the Asia region, representing a global share of 23.21% and 12.42% for incidence and mortality, respectively [2]. Report from WHO-China Joint Mission on COVID-19 showed that comorbid conditions such as cardiovascular disease, diabetes, hypertension, chronic respiratory disease, and cancer lead to poor clinical outcomes and higher mortality rates compared to patients without comorbid concerns [3]. The immunocompromised status of cancer patients caused by both the malignancy and anticancer therapies (chemotherapy, surgery, and radiotherapy) increases the likelihood to develop severe complications of COVID-19. Growing evidence from several countries particularly China, Italy, and United Kingdom have reported cancer as a major risk factor for adverse outcomes of and death from COVID-19 [4-6].

Rapid spread and epidemiological novelty of coronavirus infection have brought unprecedented challenges to the healthcare systems globally. Healthcare professionals have been urged to reorganize healthcare systems, sometimes without sufficient scientific evidence available at the beginning of the pandemic. Cancer providers need to alter care delivery models in order to handle the COVID-19 crisis as well as to protect patients without compromising cancer outcomes. Over time, various strategies have been initiated and proposed, including the countrywide strategic preparedness and response plan suggested by WHO; access to resources specific for cancer patients with COVID-19 provided by several organizations including the Union for International Cancer Control (UICC); and research recommendation and initiatives were made available on International Agency for Research on Cancer (IARC) and the United States National Cancer Institute (NCI) websites.

The Asian National Cancer Centers Alliance (ANCCA) was established in 2005 as an initiative to embark together to fight against cancer. With 14 leading cancers institutions in Asia (mostly National Cancer Centers (NCC) recognized by their respective Ministries of Health), ANCCA is the official group of leading cancer centers in Asia that serves as hub and driver to promote collaboration among ANCCA members as well as with multiple stakeholders in cancer care and prevention [7]. The COVID-19 pandemic triggered a regional collaborative effort led by ANCCA member countries, namely NCC-China, NCC-Indonesia, NCC-Japan, NCC- Korea, NCC-Mongolia, NCC-Singapore, NCC-Vietnam, as well as Institut Kanser Negara (NCI-Malaysia), National Cancer Institute (NCI)-Thailand, NCI-Turkey and Hacettepe Oncology Institute (NCI-Turkey-HOI), Tata Memorial Centre (India-TMC), Nepal Kathmandu Cancer Center (Nepal-KCC), Shaukat Khanum Memorial Cancer Hospital & Research Centre (Pakistan-SKCC), and with the collaboration of two other Asian institutions from Iran and Myanmar (Iran Tehran Cancer Institute (Iran-TCI) and Myanmar Yangon General Hospital (Myanmar-YGH), to summarize, compare and discuss the different strategies in managing cancer-care that were tailored based on country's pandemic preparedness, state of epidemic curve, political and economic status of each country.

Among the ANCCA member countries, the emergence of COVID-19 varied in extent and time, represented by various dates of national emergency declaration, depicted from 24-January 2020 for China to 7-April 2020 for Japan (Figure 1).

Figure 1:Transition of COVID-19 Case Based on WHO Data among Responders and Date of National Declaration of Emergency.

One trend that became apparent was the prompt governmental action by early responders (January – February) to the pandemic (by time of declaration: China, Mongolia, Malaysia, Singapore, Iran, and Korea), highlighted by the date of emergency declaration and occurrence of COVID-19 cases in each of these countries. In general, the course of COVID-19 pandemic and the actual encounter of COVID-19 patients in NCCs defined the context in which each institution developed and implemented its plans (Table 2). Adding to the complexity to manage the circumstances and adapt its strategies to re-organize the delivery of cancer care while going through the pandemic management were other factors such as the geographic location; status of emergency state declaration; country population; and population density or land area.

With the NCCs needing to handle challenges in adopting new ways of cancer delivery, the ANCCA facilitated the sharing of strategies to preserve high standards of care through regular video conferences among members and making available resources/materials related to COVID-19 on the ANCCA website. This paper is an extension of the sharing efforts and describes actions taken by thirteen ANCCA members and two collaborative members to re-organize cancer services by balancing the COVID-19 specific implementation strategies.

Materials and Methods

Through regular interactions, ANCCA members decided to join forces in response to the COVID-19 pandemic by designing and distributing an online survey to 14 ANCCA members, and to 5 non-members cancer institutes in the Asia region. The aim of the survey was to assess the short- and longer-term impacts of the pandemic on Asian cancer centers' activities as well as implemented strategies and countermeasures through a set of 41 questions divided into 6 categories.

The online survey tool was used to collect both open-ended and closed-ended questions (multiple choice and rating scale choice questions) to congregate relevant data from respondents. The qualitative and quantitative data were then used to analyze for trends of strategies adopted by ANCCA members and other Asian cancer centers.

Questions were distributed into 6 major categories: impact at the national and at the institutional level; hospital preparedness; strategies in place or countermeasures; triage; zoning, and repurposing. Survey topics also included the impact of the pandemic on cancer treatment and cancer care, such as distancing measures, online systems and quality of life (QOL) management. The area of clinical trial was also included for cancer centers running clinical trials before the pandemic to understand the challenges the centers experienced in conducting clinical trials (including those related to COVID-19). The specific aspect of the financial impact of COVID-19 on each institution was also ascertained from all participating members.

Quantitative and qualitative results were analyzed and sorted to allow most meaningful comparison among cancer centers. The contribution of specialists with specific knowledge and expertise from a wide range of departments, including infection control, human resources, finances, and oncology field played an essential role in extracting in-depth information on challenges and ways of addressing the pandemic.

Thirteen ANCCA members and 2 non-ANCCA Asian major cancer centers responded to the voluntary survey, providing thorough insight on the impact of the pandemic, their countermeasures as well as short- and long-term strategies. Response collection occurred between 22-May 2020 and 2-June 2020 for ANCCA members and between 8-June 2020 and 14-June 2020 for other cancer centers in Asia, representing in total centers from 15 countries in the region. Subsequent direct interviews of specialists were conducted to obtain additional answers to open-ended questions that required for input as assessed during the survey analysis.

Results

Pandemic in Asia: Overall status

Among participating countries, the emergence of COVID-19 varied in magnitude and with time, reflecting the different timing of each government's decision to declare COVID-19 as national emergency (Figure 1). Two trends could be extracted from WHO [8] data comparing COVID-19 weekly incidence of participating countries as of 19-June 2020 (cut-off date) (Figure 1). The first was based on the number of weekly cases, segregating countries into four categories: "high" incidence (>20,000 weekly cases at peak); "higher middle" (between 4,000 and 8,000 cases weekly); "lower middle" (between 500 and 3,000 cases weekly at peak); and "low" (below 100 cases weekly at peak). The second trend observed could also be classified in 3 different patterns: one single peak observed (per time of peak occurrence: China, Korea, Thailand, and Japan, seemingly having overcome the first wave of COVID-19); one peak followed by less prominent decrease (per time of first peak occurrence: Turkey and Singapore still struggling with rising curve); and a third group of countries with later start, a peak that did not reach at cut-off date (per time of occurrence start: Indonesia, India, Pakistan, and Nepal currently facing an ostentatious growth in the number of COVID-19 patients). The second trend was not applicable for countries with low COVID-19 incidence (i.e. Mongolia, Myanmar and Vietnam). The course of COVID-19 pandemic in each country (Figure 1) as well the actual encounter of COVID-19 patients at the cancer centers (Table 2) define the context and circumstances in which each institution took countermeasures to reorganize the delivery of cancer care while going through the pandemic.

Regional leadership and strategies in place or implemented(including staff management)

Despite the unpredictability of the pandemic, most participating cancer center adapted with immediate actions, partly due to availability of infection control measures and management practices, with 13 (87%) cancer centers confirmed the existence of a specific infection control dedicated department in place, and 8 (53%) confirmed the presence of dedicated staff. As a concrete action plan, most centers developed or updated guidelines or contingency measures in response to the pandemic. Eleven (73%) centers developed or updated guidelines related to infection management as countermeasures to the pandemic. NCC-Singapore's programs are worth highlighting: high level of preparedness through the existence of the Disease Outbreak Response (DORS) taskforce, regular review of its operational manual according to national and international guidelines, as well as conducting pandemic response exercises within the campus. Two other concrete examples are from NCC-Korea with a system termed Drug Utilization Review (DUR) which allows the legal access of people's overseas travel history, as well as a screening processes (with instructions on screening clinics and booths at every entrance and every visitor) in place; and Myanmar-YGH's new quidelines for the use of blood and blood products.

In addition, as a response to the pandemic, 11 (73%) centers took prompt and sustained educational initiatives directed at healthcare professionals and cancer patients, showing leadership in the cancer field in the Asia region with an obligation for educating the staff and public. India-TMC and NCC-Indonesia started regular webinars for healthcare professionals on 21-March 2020 and 22-April 2020 respectively [9-10]. Pakistan-SKCC launched a resource library and conducted

information sharing sessions for healthcare providers starting from February 2020 [11] and published a COVID-19 guideline on 10-April 2020 [12]. NCC-Vietnam organized a weekly online consultation program ("Things to know for cancer patients during the COVID-19 pandemic"), as of 14-April 2020 [13]. NCC-China enhanced the health education for cancer patients and the public on personal protection measures and principles of cancer care during the pandemic. In Korea, the Korean Cancer Society and NCC-Korea made a press release on 29-April 2020 on the "Recommendations for cancer patients based on COVID-19 situation" [14].

As another evidence of regional leadership, 8 cancer centers (53%, India-TMC, NCC-Japan, NCC-Vietnam, Iran-TCI, Myanmar-YGH, Nepal-KCC and Pakistan-SKCC) actively participated by contributing their eminent speakers to share their experience and expertise in virtual international meetings and symposium to combat COVID-19 for the benefit of other institutions or countries.

Staff management (including social distancing) was also a key focus of most centers, with respectively 9 (60%) and 11 (73%) centers implementing "working from home" and "virtual meetings" (Table 1).

Institu tion	S ta ff virtual meeti ngs	ng from		surger ies	cancer	chemo	Resch radiot herapy	ts of tr eatme	Refuse new patien t visits	Total
NCC- China										10
NCC-In donesi a										11
NCC- Japan										11
NCC- Korea										10
NCC-M ongolia										1
NCC-Si ngapor e										8
NCC-Vi etnam										9
NCI-Th ailand										7
NCI-Tu rkey- HOI										6
India- TMC										7
Nepal- KCC										10
Pakista n- SKCC										7
Iran- TCI										7
Myanm ar-YGH										7

Total 14	13	9	13	13	10	9	8	8	6	6	1	1	

Table 1: Distancing Strategies in the Cancer Centers to Decrease Physical Contact among the Staff and the Patients.

Decrease of staff was the norm, while increase of staff was also observed, partly due to staff reallocation related to COVID-19 preparedness and management of triage; pre-screening; screening activities; as well as the treatment of COVID-19 patients, in about half the respondents (7 responders (54%)).

It is noteworthy that NCC-Indonesia implemented workforce rearrangement strategy in order to protect staff (especially > 60-year-old and with comorbidities) and to ensure successful implementation of social and physical distancing. India-TMC also took similar exemplary measures to address the psychological impact of the pandemic on healthcare workers through timely communication with all cadres of staff regarding preparedness, paid-leave for high-risk staff members (elderly people, people with co-morbidities or with immunosuppressive treatment and pregnant women); rotation of staff (2/3rd on-site working and 1/3rd working from home to ensure availability of enough numbers in case of mass exposure and quarantine); and arrangement of transport for staff during the lockdown. To secure proper care, Polymerase Chain Reaction (PCR) tests for staff suspected with COVID-19 was mandatory at 9 centers (60%). NCC-Indonesia, NCI-Turkey-HOI, and Nepal-KCC implemented serologic/immunity test as a screening tool to select patients and healthcare workers who must take PCR-tests [15].

Triage (including guidelines), zoning, repurposing and material shortage - Hospital level

To combat the pandemic, all cancer centers adopted ad hoc short- and long-term measures from pre-screening to treatment or re-orientation of their cancer patients, while managing with the shortage of material (personal protective equipment (PPE), including masks, hand sanitizers, gowns, as well as oncology drugs, COVID-19 related drugs, other drugs, blood products, medical devices, and laboratory testing). Pre-screening measures included the travel history information system (NCC-China, NCC-Korea, NCC-Vietnam) and the massive COVID-19 screening and triage at the entrance (NCC-China, India-TMC, NCC-Korea, NCC-Indonesia and NCC-Singapore). As concrete examples, NCC-China closed all entrances except the main entrance, and NCC-Singapore established a fever screening clinic with COVID-19 swabbing capabilities and redirecting COVID-19 cancer outpatients to Singapore General Hospital. At NCC-Vietnam, early implementation of contingency plan and social distancing, as well as screening regulations (based on strategies taken by Central Government) eased the burden, facilitating COVID-19 prevention in the hospital.

More than half of responding centers (8 centers, 58%) had actually encountered and mostly treated COVID-19 patients, while 6 centers (43%) were COVID-19-free at the time of the survey (Table 2).

	Institution		institution	other	# Tre ated general population	Zoning plan for COVID-19 patients		
						Single entry / exit	Separate entry / exit	Dedicated ward
No encounter with COVID-19 patients	NCC-China	0	N/A	N/A	0			

	NCC-Korea	0	N/A	N/A	0		
	NCC- Mongolia	0	N/A	N/A	0		
	NCC- Vietnam	0	N/A	N/A	0		
	NCI- Thailand	0	N/A	N/A	0		
	Nepal-KCC	0	N/A	N/A	0		
Encounter with COVID-19 patient(s)	NCC- Indonesia	5 (100%)	4(80%)	1(20%)	0		
	NCC-Japan	2 (100%)	1(50%)	1(50%)	15		
	NCC- Singapore	3 (100%)	2(67%)	1(33%)	0		
	NCI- Malaysia	1 (100%)	0(0%)	1(100%)	0		
	NCI-Turkey- HOI	7 (100%)	7(100%)	0(0%)	350		
	India-TMC	314 (100%)	264(84%)	50(16%)	0		
	Myanmar- YGH	1 (100%)	1 (100%)	0(0%)	10		
	Pakistan- SKCC	231(100%)	67(29%)	164(71%)	21		

Table 2: Number of COVID-19 Positive Treated in the Cancer Centers and Zoning Plan for COVID-19 Patients.

The centers with major impact of COVID-19 were India-TMC and Pakistan-SKCC, dealing mainly with cancer patients with COVID-19, while NCC-Japan, NCI-Turkey-HOI, and Myanmar-YGH dealt mainly with COVID-19 patients from the general population. NCC-Japan was an outlier NCC, having to open a specific ward and treat 15 non-cancer patients as a response to the "special COVID-19 functioning hospital" by the Ministry of Health and Welfare from 14-April to 29-May 2020, reallocating up to 40 staff at the peak and implementing 4 shifts to avoid contamination, all based on prior experience of infectious disease management. It is also noteworthy that Pakistan-SKCC contributed to the national effort against the coronavirus and made the strategic decision to enable free testing and treatment for COVID-19 cancer and non-cancer patients unable to afford it [16].

The actual material shortage status varied among centers, less impacting for centers not having encountered COVID-19 patients, except for NCC-Mongolia (experiencing shortage of oncology drugs and medical devices due to shipment delay) and Nepal-KCC (with shortage of PPE, cancer and non-cancer drugs as well medical devices and other equipment). For centers experiencing COVID-19 patients, a shortage of PPE, swabs, tubes for reagents, isolation gown, cancer and non-cancer drugs as well as of medical devices was noted at various levels depending on the centers. The highest impact of shortage was on PPE, and also reagents for PCR testing (NCC-Japan, NCC-Indonesia), and Rapid Detection Kit (RDT) (NCC-Indonesia). To overcome the pandemic, Pakistan-SKCC (the most affected institution) as well as India-TMC and NCC-Indonesia took initiatives such as accepting donations from philanthropic organisation or government while NCC-Singapore prevents the shortage of PPE through developing a strict PPE guideline for optimal protection of staff and patients. As a specific case, NCI-Turkey-HOI did not experience any specific shortage despite the high number of COVID-19 patients in the institution.

Patient care: impact on treatment and online systems - Hospital level

Aiming to preserve high standards of care for patients with cancer, participating centers adapted to their healthcare environment and needs in different ways (Table 1). In addition to decreasing contact through exposure to family members and visitors, implemented by all centers (100%),

various aspects of cancer treatment were modified, with a non-negligible impact on both the institution and patients and their families. Worth noting is the postponement of non-urgent surgeries, rescheduled surgeries, cancer screening, chemotherapies or radiation therapies taken respectively by 13 (93%), 10 (71%), 9 (64%), 8 (57%) and 8 (57%) centers, treatment modifications highlighted by several members as case-by-case decision based on prioritization. In order to minimise the impact on cancer treatment, respondents adopted different strategies, such as: for patients preparing to be admitted, NCC-China routinely recorded symptoms potentially associated with COVID-19, such as fever and cough through mandatory routine blood tests and high-resolution computed tomography scans of the lungs. India-TMC did not modify its treatment protocols except rescheduling the post-treatment follow-up and selecting less myelosuppressive chemotherapy for palliative indication; while NCC-Indonesia and NCC-Vietnam rescheduled suitable treatments making sure not to compromise the treatment objectives. Two exceptional cases were NCI-Turkey-HOI, with no modification in cancer treatment, and NCC-Mongolia, with zero local transmission as of 22-May 2020 (time of filling the survey), which only restricted the number of family members and visitors.

Focusing on the "telemedicine" aspect, 13 centers (93% of participants) implemented online consultation in various ways (Table 3).

Institution	Prior system in place for onli ne consultatio n	% Online c onsultation prior to pandemic	% Online c during pandemic		Types of online co nsultation			Notes
				Telephone	Video	Chat-based	E-mails	System in place, free of charge, chat- based system on NCC website
NCC-China		5%	70%					Online consultation available but outside of the scope of insurance coverage
			10%					Approval of online consultation by MHLW on 1-Apr, allowing patients to pick up their drugs at pharmacies closed to their house
NCC-Japan		0%	8%					Online (telephone) consultation legalized on 24-Feb
NCC-Korea		0%	3%					National insurance scheme system in place for online

NCC- Mongolia		0%	0%			COVID-19 transmission Secure system recently developed
						for consultation and charging
NCC- Singapore	0	0%	10%			Common unofficial practice of telephone consultation between patient and physician, not paid therefore not recorded
NCC- Vietnam		0%	90%			
NCI- Thailand		0%	5%			
NCI-Turkey- HOI		0%	25%			
India-TMC		10%	30%			
Nepal-KCC		60%	80%			Online credit card mobile billing application
Pakistan- SKCC		0%	60%			Online consultation applied only at palliative care services, through the use of personal phone number
Iran-TCI		20%	40%	0		Booking ahead, Viber group, Facebook page
Myanmar- YGH		30%	50%			

Table 3: Proportion and Types of Online Consultations at each Cancer Centers before and During the Pandemic.

NCC-Vietnam made the most significant change: from zero before to 90% during the pandemic. NCC-China, India-TMC, Nepal-KCC, Iran-TCI, and Myanmar-YGH had already implemented the online consultation system prior to the pandemic, which allowed for a smoother implementation. In China, the national health and insurance system in place allowed cancer patients from all over China to get free access to online free-of-charge consultations including instructions on taking medication and cancer-related symptoms management. In Japan, the Ministry of Health and Welfare approved Emergency Countermeasures for COVID-19 on 7-April 2020 which allowed NCC-

Japan to implement telemedicine [17], still in application at the time of this manuscript submission. Another fast shift was in NCC-Singapore's implementation of a secure system for consulting and charging.

On the quality of life perspective (Table 4), most centers took initiatives: NCC-Singapore allowed video-conferencing for patients and their families, as well as between medical teams and relatives to update on patient's conditions.

Institution	Specific actions to maintain cancer patient quality of life	
	Outpatient support	Inpatient support
NCC-China		Psychology support for suspected or confirmed cases of COVID-19 patients by clinical psychologist to facilitate patients' acceptance on their illness, reduce patients' anxiety, and provide motivation to improve patient's condition
NCC-Indonesia	1. Provision of online consultation for palliative patients, conducted by palliative doctor as alternative solution for home care/visit that cannot be held during partial lockdown	Psychology support for suspected or confirmed cases of COVID-19 patients by clinical psychologist to facilitate patients' acceptance on their illness, reduce patients' anxiety, and provide motivation to improve patient's condition
	2. Provision of medicine refill through electronic prescription following online consultation	
NCC-Japan	Information sharing about home exercise for physical activity	
	2. Prescriptions made available at local pharmacy after online consultation with physician	
NCC-Korea	1. Provision the home hospice visit service to the patients as usual with medical staffs only (no volunteers, social welfare workers, etc.)	Maintaining the provision of palliative care service while minimizing group meeting among patients and the number of responsible staffs (e.g. excluding volunteers and maintaining only medical staffs)
	2. Maintain the highest sanitary conditions to the visitors by applying strict infection control policy	
NCC-Mongolia	No particular action needed to limited impact of COVID19	Introduction of dedicated hours and restricted access for the visitors and family members at the hospital
NCC-Singapore	1. Support to inpatient medical teams by palliative physician	Video conferencing for patients to stay in contact with their family
	 Video consultation for community care team for outpatients for management of pain and other quality of life issues 	2. Communication between medical team and relatives to update conditions as visitors are not allowed in the wards
NCC-Vietnam	Tele-consultation open session on the hospital website and Facebook about nutrition and psycho-oncology support for patients	
NCI-Turkey-HOI	No need for prescription and coming to hospitals	
India-TMC	1. Provision of food and accommodation	1. Provision of food to the relatives
	2. Travel arrangement	2. Extending the stay for those who were outstation patients
	3. Linking them with the volunteers for psychosocial support	

	4. Support group meetings using platforms like zoom	
Nepal-KCC	Provision of free vehicle facilities for cancer patients and hand sanitizer to all	
Pakistan-SKCC	Everyday telephonic updates for relatives. Video calls between patients and relatives	
	Palliative care team (volunteers) in some region support patients/ virtual consultation by palliative medicine specialist	Virtual tumor boards about difficult patients
Myanmar-YGH	Provision of bed allocation 3 feet apart/by marking waiting area for social distancing, provision of hand washing facility, restriction of patient attendance	Provision of good nutrition support and palliative care for all inpatients

Table 4: Initiatives Related to Quality of Life of the Patients.

NCC-Japan and NCI-Turkey-HOI, smoothened the prescription process to facilitate patients to obtain drugs through online consultation without the burden of an hospital visit. NCC-China, NCC-Vietnam and India-TMC undertook similar approaches to overcome COVID-19 impact on patient's wellbeing by providing food or nutrition support, psychosocial support, as well links with web-based support group meetings. NCC-Japan provided web-support, allowing home exercise for cancer patients.

Clinical trials

Research-wise, the pandemic also highly impacted 8 of the 9 centers (89%) who were actively participating in clinical trials prior to the pandemic, with NCC-Vietnam as an exception, not much impacted due to short social distancing period and controlled outbreak in Vietnam. Most industry-led trials were delayed, with major impact on patient recruitment and visits on 7 centers (78%), and on the overall timeline for 6 centers (67%) (Table 5).

Institution	Patient Recruitment or Enrolment	Providing product to patients	Patients' visits	Overall quality	Overall co mplet ion timeline	Participation to new clinical trial related to COVID-19
NCC-China						N/A
NCC-Japan						Avigan observational study. 14 out of 15 COVID-19 patients at NCC were enrolled. This study is a nation-wide study
NCC-Korea						N/A
NCC-Singapore	0					Collaboration on clinical trial on Virus specific T cells for COVID-19
NCC-Vietnam						N/A
NCI-Turkey-HOI						N/A
India-TMC						Cancer outcomes in patients with

			COVID-19 ongoing
Pakistan-SKCC			WHO COVID-19 Vaccine Trial
Iran-TCI			N/A

Table 5: Impact of COVID-19 Pandemic on Clinical Trials.

*N/A, Not Applicable

Relevant actions and countermeasures included: rapid identification of life-saving clinical trials for immediate approval to continue; deployment of clinical trial coordinators to triage and screening when patient recruitment was reduced (NCC-Singapore); telephone-call based follow-up for trial patients and sending drugs by courier services (India-TMC). By contrast to industry-lead clinical trials, NCC-Japan reported that the impact on investigator-led clinical trials was limited.

Acting as Asian leaders in their field, several ANCCA members also actively contributed to knowledge on COVID-19: NCC-Singapore's collaboration on a clinical trial on Virus specific T cells for COVID-19, India-TMC evaluating cancer outcomes in patients with COVID-19, NCC-Japan and Pakistan-SKCC respectively participating in a nationwide Avigan observational study and a WHO COVID-19 vaccine clinical trial.

Impact on NCCs' departments (including manpower and financial impact)

National Cancer Centers and Institutions from ANCCA are established as leading cancer centers in the region, hosting a broad range of cancer related services to maintain a comprehensive cancer care, cancer control activities, and research on cancer. Clinical and education services and research activities are core activities for all participating centers, besides the other cancer-related departments as depicted in Table 6.

Institution	Hospital	Cancer S cr eening and Prevention	R ese arch Institute	National Can cer Control Departmen t	Public health	Educationa l services	Central La boratories / PCR machines	
NCC-China	**	**	***	*	N/A	**	***	-
NCC- Indonesia	**	***	-	N/A	N/A	***	**	1
NCC-Japan	**	***	*	*	*	***	**	1
NCC-Korea	*	*	*	*	**	**	*	1
NCC- Mongolia	*	*	-	-	-	**	-	-
NCC- Singapore	**	*	***	N/A	N/A	**	**	1
NCC- Vietnam	*	*	-	N/A	**	*	-	1
NCI- Thailand	*	**	-	*	**	*	-	1
NCI-Turkey- HOI	*	*	*	*	*	**	-	1
India-TMC	*	***	**	**	***	*	-	1
Nepal-KCC	*	**	*	*	-	-	-	1
Pakistan- SKCC	**	N/A	-	-	N/A	**	-	-

Iran-TCI	**	**	*	**	**	**	**	\downarrow
Myanmar- YGH	*	*	*	*	*	**	*	1

Table 6: Impact of COVID-19 on each Institution.

*** Had to shut down; ** Highly impacted; * Impacted somehow; -No change; ↑ Increase; ↓ Decrease; N/ANot applicable

All participating centers, even the 6 centers which did not experience COVID-19 patients, were affected by the pandemic. Most highly impacted departments were the hospitals (all centers impacted,respectively highly impacted and impacted somehow by 8 (57%) and 6 (43%) centers), education services and cancer screening and prevention (respectively 10 centers (71%) and 7 centers (50%) had to shut down or were highly impacted). In NCC-Vietnam, the social distancing regulations, even though for a limited period of time, created a huge burden on internal resource requirement and operational coordination, with a drastic increase of patients (new and existing) at the imaging diagnostic department, resulting in subsequent increase of workload for the whole department following the lifting of social distancing measures.

Financial consequences in most departments caused by the pandemic were significant, estimated to be around 30% loss in comparison with the previous year, depending on the institution. Countermeasures were limited or non-existent for most cancer centers at the time of completion of the survey. COVID-19 related expenses were diverse, including reduction of outpatient and inpatient visits, staff and medical equipment shortage after complete or partial lockdown declared in all participating countries, independently from the actual occurrence of cases in the country or the institution. Striking examples are NCC-Vietnam and NCC-Mongolia which were highly financially impacted despite low number of cases in the country. NCC-Vietnam experiencing a 30% financial loss due to triage and reduction of patient visits in addition to the increased budget spent on extra-hours to screen staff and visitors, to quarantine suspected cases and other related measures. Another costly investment worth pointing out was the new fever and triage area in NCC-Singapore, and cost related to the increased lab capacity in NCC-Indonesia (which was appointed on 15-April 2020 by the Regional government to provide PCR tests for COVID-19).

Discussion

NCC Leadership and strategies

National Cancer Centers around the world have the responsibility to balance the risk from delay in cancer diagnosis or treatment against the potential risk of COVID-19 exposure, mitigate the risk of disruptions to cancer care delivery during implementation of social distancing strategy, and manage the proper allocation of limited health care resources. Outbreaks of infection etiology, particularly those caused by a novel virus with no known treatment, may result in the interruption of medical care provided to patients with cancer and put them at risk for undertreatment in addition to the risk of being exposed to infection, which collectively could be a life-threatening event for patients with cancer [18].

ANCCA members have established a roadmap at the end of 2019 [7] with short-, mid-, and long-term goals to halt cancer increase and mortality rates in Asian countries by 2030. The pandemic, through its dramatic impact on society, and even further on cancer care, has opened doors to new collaboration and goals not originally specified. Such pandemic challenge faced by each and every country in the region has allowed ANCCA members to increase the speed and focus the goal of collaboration to decrease the impact of the pandemic on one of the most affected population: cancer patients and healthcare professionals. Shortly after the start of the pandemic, on 22-April

2020, the presidents of 3 NCCs (NCC-China, NCC-Japan and NCC-Korea) virtually gathered to share latest updates, followed by a virtual meeting attended by most members on 20-May 2020, to promote information sharing among ANCCA members through a survey addressed to each institution representative.

Managing cancer care through a pandemic was not new for 10 participating cancer centers (67%) which experienced one or two major coronavirus outbreaks in the region: Severe Acute Respiratory Syndrome (SARS) with a 2003 and Middle East Respiratory Syndrome (MERS), with a 2012 outbreak that have allowed several Asian countries to prepare for the pandemic, from a governmental as well as institution level management of the pandemic [19]. The participating countries of this study were respectively China, India, Indonesia, Korea, Malaysia, Mongolia, Thailand and Vietnam for SARS and Korea, Malaysia, Iran and Turkey for MERS [20] with Korea and Malaysia experiencing their third encounter this time since coronavirus in 2002.

The survey outcomes included the establishment of an infection control department and/or dedicated staff, which turned out to be in place prior to the pandemic for respectively 13 centers (87%) and 8 centers (53%), allowing the distribution of fast and relevant information to the community (healthcare workers, as well as patients) that are key to contribute to national, regional and international efforts to decrease the impact of the pandemic on cancer care.

The results of this study include lessons learned through very stringent actions for NCC-China and NCC- Korea, both tracking the travel history and symptoms of all individuals as well as performing a thorough screening at entrance of all employees.

Caring for staff strategies

Cancer care providers are at increased risk for coronavirus infection as chances of acquiring infection at workplace are high. Previous experiences of SARS and MERS outbreaks had shown that the rate of Coronavirus transmission among healthcare workers were significant, accounting for 21,07% [21] and 19,1% [22-23] of confirmed cases, respectively. Current data on COVID-19 case among healthcare providers are incomplete and very dynamic. However several studies have reported a high rate of COVID-19 transmission among healthcare workers since the onset of the outbreak [23-25]. This situation has led to substantial decrease of available staff due to self-isolation (staff was suspected of COVID-19 or household member developed symptoms).

The pressure of working under pandemic circumstances was augmented by staff's anxiety of the risk of personal contamination and extended work hours [25]. Having contact with confirmed cases of COVID-19 demonstrated as significant factors of high level of distress, experienced by 29-35% healthcare workers delivering care to SARS patients [26] and 26% of doctors involved in MERS care [27]. Past experience provided lessons that healthcare management team should put attention on staff's need of logistic and psychological support [23]. In NCCs of the participating countries, staff protection is one key focus of COVID-19 related strategies, including the implementation of staff virtual meeting and working from home policies to ensure social distancing approach; modification on treatment regimens and rescheduling of non-urgent treatment to reduce hospital visits, and psychological support for staff dealing with suspected or confirmed COVID-19 cases.

Impact on patient care

Cancer center efforts to continue its operations and ensure the safety to resume elective procedures have been guided by standards published by international oncology associations, particularly ESMO, ASCO, NCCN, and American College of Surgeons. Patients prioritization and modification of cancer treatment protocols were implemented to reduce hospital visits and ensure the implementation of social distancing measures in order to minimize the risk of SARS-CoV2 transmission. Similar approaches have been taken by cancer centers worldwide, for instance

oncology institutions under the legal entity of Cancer Core Europe (CCE) [28] and comprehensive cancer centers in US [29-31]. The strategy was effective to reduce the overall number of cancer patients admitted by 70-80% of the normal influx [28] and decrease the on-treatment patient volume (OPTV) by 25% in anticipation of the expected local COVID-19 peak [29].

The extent of social-distancing strategies implemented by NCCs are determined by government approaches to control pandemic at the national level. The population-level physical distancing measures and movement restrictions, often referred as "shut down" and "lock downs" were introduced in many countries, especially where community transition has led to outbreaks with near exponential growth [32]. Two different approaches to control COVID-19, namely suppression and mitigation were introduced and implemented in each country with varying extent. Suppression strategies aim to reach nearly complete suppression (reverse the epidemic spread to reproduction number (R) < 1 and establish population immunity once a vaccine becomes available) [28,33], while mitigation strategies aim to keep COVID-19 incidence consistently at the maximum levels acceptable in order to prevent overwhelming the healthcare systems (establishing population immunity with or without a vaccine) [28]. China, Singapore, and South Korea have demonstrated an example of suppression strategy [28-34], whereas India [35] and Indonesia [36] have been employing a mitigation strategy. Complete suppression is the ideal strategy in cancer care setting, though the potential socio-economic impact of closing down the society have led countries to employ a mitigation approach over suppression strategies.

Strict social-distancing strategies have led to psychological issues in cancer patients due to the uncertainty of treatment continuation. As leading cancer centers in Asia, participating NCCs in our study provide care to cancer patients throughout the countries and serve as main referral cancer hospital in the region [37]. During the pandemic, access to healthcare is difficult in the current restrictive mobilization state and travel limitation [35]. Furthermore, many patients have been fearful of exposing themselves to the risk of infection and have been more reluctant to present to healthcare services. This situation affects the mental health of the patients and their families in addition to anxiety caused by cancer diagnosis and treatment [37]. The ANCCA members and participating countries have implemented impactful initiatives including travel and vehicle arrangement, employing telemedicine to provide psycho-oncology support, cancer education, nutrition support, and video recording for home-exercise. Automated medication refill and delivery have been also provided to ensure treatment continuity (Table 4). As supported from the literature, communication using digital technology can be adopted as a key strategy to continue delivery of cancer care while protecting vulnerable oncology patients and health care workers [38-39]. Telemedicine is also tool to empower patients and caregivers, therefore alleviate their social burden and improve quality of life [37].

The COVID-19 pandemic is rapidly transforming the medical care system and the use of industry 4.0 technologies has the potential to fulfil customised requirements during the crisis [40]. Several implementations such as telemedicine, utilization of travel databases for adequate screening, as well as virtual meetings admittedly are effective approaches to avoid potential risk of COVID-19 exposure. Going forward, further application of technologies, for instance digital imaging, remote medication supply, the use of machine learning and artificial intelligence for autonomous robot for examination, and the use of virtual reality environment for training and education purposes are transformational windows of opportunity for safer oncology care and training delivery [40]. Virtual care delivery should be appropriately documented to facilitate billing [41]. National health coverage and commercial medical insurance should reconsider to revise their payment policy and provide reimbursement for telemedicine activities in response to pandemic [42]. Currently, telemedicine is more of a use to follow up cases and is not adequate for newly diagnosed of cancer or patients under evaluation. There is still a need of face-to-face consultation for optimal evaluation of new patients. Delaying or postponing cancer treatment due presumed increased risk of infection with COVID-19 is a matter of debate and dilemma [37].

Several studies which demonstrated that cancer patients are more vulnerable to COVID-19

complications have encouraged physicians to withhold or postpone cancer treatment during the epidemic [43-47]. The NCCs have been applying strict social distancing measures that aim to decrease hospital visits and elective procedure. Furthermore, cancer control measures have been temporarily put on hold as NCCs' effort to minimize potential exposure of cancer patients to SARS-CoV-2. Our study showed that cancer control activities, in particular cancer screening and prevention have been impacted in varying degrees. Cancer screening has also been suspended, similarly to several countries in Europe [48], UK [49], US [50-51] following a call from the government to prioritize on coronavirus-related treatments. Halting cancer screening procedures will lead to more advanced stage at diagnosis, and in the future, higher economic burden of a delayed diagnosis of cancer will be loaded on public health economy [52-54]. The NCCs need to prepare for the expected increase of patients when government decide to loosen the social distancing measure. Huge workload of patient influx in the case of imaging diagnostic on NCC Vietnam is an example of future burden due to delayed diagnosis and cancer treatment that led to risk of exhausted health system.

In the clinical perspective, the paucity of solid evidence on the benefit of treatment modification or interruption should led to careful decision by oncologists and individualized for every patient [25]. Recent studies by Kuderer et al. [55] and Lee et al. [56] argue that treatment delays would significantly cause more harm to cancer patients [52]. In both studies, risk of death of cancer patients who had confirmed coronavirus infection was significantly associated with age, male sex, former smokers, associated comorbidities, active cancer and poor performance score; but not with administration of immunotherapy, chemotherapy, radiation therapy or targeted therapy within 4 weeks of detection of SARS-CoV-2. These evidences clearly urge the need to facilitate the cancer continuum and government decision to re-open the society.

Cancer research and clinical trials

The pandemic has affected clinical care in a broad range of settings, disrupting all aspects of clinical care, including cancer clinical trials. Numerous challenges with conducting clinical trials highlight opportunities to be evaluated and applied as proposed in the US [57] and also in the Asian region. The experience of the nine centers (60%) actively conducting clinical trials provides additional insight on the dramatic impact of the pandemic in Asia, particularly, on patient recruitment and patients' visits for sponsored clinical trials, while investigator- initiated clinical trials were less impacted. Going forward, the restart of clinical studies, as well as the implementation of COVID-19 related clinical trials are likely to create a significant delay and financial burden on the conduct of clinical trials. A new norm will have to be applied around the world to catch up with such huge impact. Unnecessary visits, and other activities set in study protocols can be removed after discussion among pharmaceutical companies, healthcare professionals and patients, to minimize the burden on patients while optimizing well balanced clinical trials.

On the other front, epidemiological and registry research on the impact of COVID-19 on cancer care in national cancer institutions will be key to address issues, not only within clinical trials, but to cover the full scope of cancer care.

Way Forward/Preparing for the future

The COVID-19 pandemic is likely to disrupt health system in lasting ways. Estimating the extent and severity of novel disease outbreaks is dependent on aggressiveness, accessibility and availability of specific testing. Although there could be differences in population demographics and access to healthcare, COVID-19 death rates (as shown in Appendix 1) could be more reflective of the true incidence and impact of the current pandemic. The current strategies taken by NCCs are likely to evolve over time, depending on the stage and width of the pandemic. The priorities are to focus on maintaining cancer care delivery while protecting both patients and health care workers

from the risk of COVID-19 transmission. With some countries having loosened up the population-level physical distancing measures and movement restrictions (as of 8-July 2020, the time of this writing) and preparing for their "new normal", NCCs in Asia have to continuously monitor and evaluate the effectiveness of their mitigation strategy (Table 7).

Area of Focus	Strategies employed	Experience gained / Lessons learned
Crisis management and preparedness toward sustainability and longer-term preparedness	Contingency plans	-Sharing best practices among members
	• Implementation of new guidelines and standard procedures	-Logistic supports and availability of medicines, material and devices
	Creation and update of guideline: Management of infectious disease on cancer care setting- current momentum	-Telemedicine implementation and national potential applications
	Crisis communication	-Guidelines and criteria for patients requiring admission or self-quarantine to reduce risk of transmission
		-Role of media on awareness program and education
Emergency preparedness	Triage at every entrance	-Tracking of travel history / pre- screening
	• Zoning area (single vs separate entry) with dedicated ward for suspected/confirmed cases	-Screening booths
		-Separate routes for infected patients
		-Dedicated isolation ward (including ICU)
		-Simulation drills
Cancer patients' care (short term)	Social distancing	-Online consultation system
	Patient's QOL management	-Booking by phone one day ahead of outpatient consultation
		-Transportation / travel arrangement for patients
		-Psychological support to improve patient's QOL
		-Transparent information to patients
		-Specific education on principle of cancer care during pandemic
		-Home exercise support
		-Managing patients' expectation based on their experience during pandemic
		-Family/caregivers support to cancer patients with COVID-19 infection
Cancer care continuum	Adaptation of treatment strategies (across cancer types and specific by cancer type)	-Modification on cancer protocol
		-COVID-19 testing to asymptomatic patients' prior treatment/procedure
		-Avoidance of delaying treatment/procedures to COVID-19 negative cancer patients
		-Increased capacity on remote diagnostic and treatment through optimization of digital technology
		-Single-room-one-patient policy for infected patients
Staff management	Caring for staff: address staff shortage and psychological support	-Transportation / travel arrangement for staff
		-Rotation of staff in shifts to avoid cross-

contamination
-Special care for staff with co- morbidities, taking immunosuppressive agents, and > 60 years old (e.g. paid leave)

Table 7:The Way Forward [58-59].

ANCCA will keep playing the role of a think-tank, gathering brains to share expertise, lessons learned and allow win/win scenarios among all participating cancer centers (Table 7)(Table 8).

Country		Cumulative deaths
India	878,254	23,174
Iran	257,303	12,829
Turkey	212,993	5,363
Pakistan	251,625	5,266
China	85,568	4,648
Indonesia	75,699	3,606
Japan	21,868	982
Korea	13,479	289
Malaysia	8,718	122
Thailand	3,220	58
Nepal	16,801	38
Singapore	45,961	26
Myanmar	331	6
Mongolia	230	0
Viet Nam	372	0

Table 8: Appendix 1 - Cumulative COVID-19 cases and deaths (13-July 2020).

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Statement conflict of Interest

No potential conflict of interest was reported by the authors

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