Soft tissue sarcoma in Asia

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Abstract: Sarcoma is an uncommon and heterogeneous group of malignancies linked by their mesenchymal origin. They are rare and account for 1% of adult cancers, and 10–20% of adolescent and young adult (AYA) cancers. While there is good published literature on the incidence and distribution of sarcoma subtypes in the western populations, there is a paucity of data from Asia, particularly on the epidemiology, treatment and outcomes of STS in Asia. Formalized prospective national registries in soft tissue sarcoma (STS) are lacking and little is known about how sarcomas are treated and managed throughout the Asia-Pacific region. Due to geographical and varying affluence levels across over 50 countries, no single uniform guideline exists across Asia to inform of chemotherapeutic options. Any existent guidelines tend to be country-specific, reflecting resource availability and geographical limitation. To understand and improve STS care in Asia, one must appreciate the macroeconomics and healthcare structures in place and to work within the limitations imposed by them. Concurrently, there exists an urgency to develop strong sarcoma centres of excellence (SCE) across Asia to deliver state-of-art care to our patients. Equally important is the need to facilitate sarcoma education to patients and healthcare workers. Development of such centres is vital in improving clinical care as they provide high-quality sub-specialized care to patients within that geographical location, serve as reference centres for knowledge and education as well as nerve centres for care-coordination in a hub-and-spoke model. Key to these centres of excellence is the sarcoma multi-disciplinary team (MDT). Studies have consistently shown that patients managed in high-volume sarcoma centres, by an expert sarcoma MDT, achieve better clinical outcomes. This report serves to highlight the challenges and opportunities of sarcoma care in Asia, map out a vision for the development of SCE across Asia and highlight the areas of potential collaboration between centres to advance the science of sarcoma.

Keywords: Asia sarcoma; adolescent and young adult cancer (AYA cancer); rare cancer in Asia

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Introduction to soft tissue sarcoma (STS)

Sarcoma is an uncommon and heterogeneous group of malignancies linked by their mesenchymal origin. They are ubiquitous and may develop from connective tissues anywhere in the body (1-3). Sarcomas account for 1% of all adult cancers. In 2018, an estimated 13,040 people will be diagnosed with STS in the United States, with approximately 5,150 deaths (4). Based on the current WHO classification, there are now more than 50 different distinct subtypes of STS (4), with the most common aggressive STS being liposarcoma, leiomyosarcoma and undifferentiated pleomorphic sarcoma (3).

While there is good published literature on the incidence and distribution of sarcoma subtypes in the western populations, there is a paucity of data from Asia, particularly on the epidemiology, treatment and outcomes of STS in...
Asia (5). Formalized prospective national registries in STS are lacking and little is known about how sarcomas are treated and managed throughout the Asia-Pacific region (6).

While the general paradigm of sarcoma care may be similar in Asia as it is in the West, exact treatment guidelines are less established. They tend to be country-specific (7), reflecting resource availability within each country (7). This is particularly important and interesting considering the geography and heterogeneity of countries within Asia.

To understand and improve STS care in Asia, one must appreciate the macroeconomics and healthcare structures in place and to work within the limitations imposed by them. Concurrently, there exists an urgency to develop strong sarcoma centres of excellence (SCE) across Asia to deliver state-of-art care to our patients and facilitate sarcoma education to patients and healthcare workers. This report serves to highlight the challenges of sarcoma care in Asia, map out a vision for the development of SCE across Asia and highlight the areas of potential collaboration between centres to advance the science of sarcoma.

The state of STS in Asia

Reviewing the existing literature available for STS trends, there is a lack of outcomes and epidemiological data from Asia (5). There is no prospective data available, with most if not all, being retrospective data and are specific to a country (6,8-10). Ngan et al. had done a systematic review which gave a glimpse into the survival outcomes in STS (6).

In 2015, the STAR study (STS in the Asia-Pacific region) was reported, which described the epidemiology, treatment patterns and clinical outcomes of 635 patients with STS in 5 Asian countries. It demonstrated a median overall survival of 11.7 months in patients who presented with metastatic STS, a result comparable with outcomes from the West (5).

The predisposition to developing STS is not much different in Asia compared to our Western counterparts. Some of these include family cancer syndromes, previous exposure to radiotherapy or chemicals, HIV infection and autoimmune conditions (11-14). Family cancer syndromes include neurofibromatosis, Gardner syndrome, Li-Fraumeni syndrome, retinoblastoma and tuberous sclerosis. Due to the vast disparity in access to medical healthcare and knowledge, it is therefore not unexpected that some of these risk factors may not be known nor even diagnosed some parts of Asia. In general, the prognostic factors are similar to what is expected in the Western population. Malignancy grade, tumour type and size, site, tumour resectability, surgical margins achieved and presence of metastases at diagnosis are important prognostic factors (15-18). There are some tumor-specific retrospective studies, but these also show similar prognostic factors (19).

In the management of advanced STS, standard chemotherapy is still based on anthracyclines as the first-line treatment. Multi-agent chemotherapy is usually with the addition of ifosfamide, when a tumour response is felt to be potentially advantageous in a reasonably fit patient (15). There is now also the possible consideration of adding on an anti-PDGFRA agent, olaratumab to doxorubicin. Tap et al. had shown an improved progression-free survival (PFS) of about 2 months and an improved median overall survival of 12 months in their randomized phase 2 trial (20). Beyond first-line, guidelines suggest treatment that is driven by histology (15). Other recent therapeutic considerations include using palbociclib, a CDK4/6-inhibitor in patients with CDK4-amplified, well-differentiated or dedifferentiated liposarcoma, which showed an improved 10% PFS (4,21).

Medico-socio-economics and the healthcare structure in Asia

There are about 50 countries that make up Asia with huge socio-economic discrepancies (5), both among and within countries. A country with lesser resources will be expected to have only the rudimentary basics of healthcare. Those with richer resources can afford for a more sophisticated healthcare structure. In Southeast Asia (SEA), delivering healthcare services to people dispersed across hundreds of islands or to remote areas is very challenging, made even more difficult when sub-specialized healthcare is needed (22). Recent analyses have drawn attention to the weaknesses of healthcare systems in low-and-middle-income countries (23).

Dr. Anne Mills, a British authority on health economics, published in 2014 a report identifying six levels of constraints that exists within any health care system from the community to the global level (23). In Table 1, we build upon this to reflect how this concept applies to sarcoma care in general and why SCE may help to mitigate some of these issues faced.

SCE

Development of SCE is vital in improving clinical care in patients. These centres provide high quality sub-specialized
sarcoma care to patients within that geographical location, serve as reference centres for knowledge and education as well as nerve centres for care-coordination in a hub-and-spoke model. Key to these centres of excellence is the sarcoma multi-disciplinary team (MDT). Studies have consistently shown that patients managed in high-volume sarcoma centres, by an expert sarcoma MDT, achieve better clinical outcomes (24,25).

**MDT**

A sarcoma MDT should typically consist of specialist sarcoma surgeons, radiation and medical oncologists, preferably a dedicated sarcoma pathologist, radiologist, clinical nurse, as well as palliative care specialists, allied health members and support staff. The MDT serves as a forum for this group of specialists to meet and discuss the management of each patient in a prospective fashion, prior to initiation of treatment, allowing for a coordinated actualization of a well thought-through plan from initial diagnosis to definitive treatment and follow-up. MDTs have become the accepted way of delivering modern cancer care and is key to the provision of high-quality individualized sarcoma care (2). In published studies, improvement in overall-and disease-free survival was observed in patients treated as part of a sarcoma MDT. In a population-based Canadian study, patients with limb STS treated by an MDT within 3 months of diagnosis had improved overall survival and reduced risk of amputation. A UK study also reported better overall survival for pediatric rhabdomyosarcoma treated in pediatric oncology centres compared to those treated in a general hospital (26). A strong sarcoma MDT could also serve as a platform to audit clinical practice, share best practices and enhance national, regional and international collaborations, which would all raise the quality of care in patients.

Of note, STS may be less common than epithelial cancers and there exists a need to justify the time and resources of a

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**Table 1 Constraints in a healthcare system**

<table>
<thead>
<tr>
<th>Level of health care system</th>
<th>Constraints</th>
<th>How/why is sarcoma care affected?</th>
<th>How a sarcoma centre of excellence can mitigate</th>
</tr>
</thead>
<tbody>
<tr>
<td>Community and household</td>
<td>Lack of demand for effective interventions</td>
<td>Less awareness</td>
<td>Create awareness</td>
</tr>
<tr>
<td>Service delivery</td>
<td>No standardization on quality of care</td>
<td>Different standards of care</td>
<td>Strengthen training and supervision</td>
</tr>
<tr>
<td></td>
<td>Inadequate drugs and medical supplies</td>
<td>Without access to treatment,</td>
<td>Set standards of care</td>
</tr>
<tr>
<td></td>
<td></td>
<td>unable to deliver care</td>
<td>Set guidelines</td>
</tr>
<tr>
<td>Policy and strategic</td>
<td>Non-standardized drug policies</td>
<td>Appropriate medical care is</td>
<td>Identify what is basic requirements within each country and ensure that this is available</td>
</tr>
<tr>
<td>management in the health</td>
<td></td>
<td>unavailable</td>
<td></td>
</tr>
<tr>
<td>sector</td>
<td>Limited communication and transport</td>
<td>Unable to get the right</td>
<td>Satellite centres increase access for patients to seek specialized healthcare. Helps to mitigate problems of distance</td>
</tr>
<tr>
<td>Government policy</td>
<td>infrastructure</td>
<td>information and adequate care to people</td>
<td></td>
</tr>
<tr>
<td>Political and physical</td>
<td>Climatic and geographic predisposition to</td>
<td>Affects how care is delivered</td>
<td>Brings medical care closer to patients</td>
</tr>
<tr>
<td>environment</td>
<td>disease, physical environment unfavorable for service delivery</td>
<td>and accessed by patients</td>
<td></td>
</tr>
<tr>
<td>Global</td>
<td>Fragmented governance and management structures for global health</td>
<td>No treatment consensus guidelines, leading to unequal care</td>
<td>Standardized good quality treatment and sharing of best practices</td>
</tr>
</tbody>
</table>
large sarcoma MDT to support the care of a more limited number of patients. Thus development of regional sarcoma referral centres or centres of excellence with established networks and referrals may be very important to maintain expertise and justify resources needed (26).

Pathologist

An accurate pathological diagnosis is vitally important in sarcoma management. A sarcoma pathological diagnosis must confirm the presence of sarcoma, indicate the grade and histological subtype of disease (27). However, it is well-recognized that pathological diagnosis is challenging in STS. Given the rarity of the disease, inappropriate medical management has been reported in more than 70% of sarcoma patients (28). Articles recommending histopathologic peer reviews for sarcoma have continued to appear in recent years (29-34). But this service is frequently not available in many parts of Asia. Clearly this is an issue given the frequency of misdiagnosis with carcinoma, melanoma, benign tumor, or even between sarcoma histological subtypes (35,36). The reproducibility of STS diagnosis is also relatively poor across unfamiliar pathologists and the histopathological classification of this cancer in cancer registries is often inconsistent (31). With increasing sub-specialization, it is inevitable that histopathologists become less skilled outside their areas of practice. There is a consistent discrepancy between referring and expert diagnosis of approximately 25% for diagnosis and grade, and about 5% between a malignant and benign diagnosis (29,33,35-44). The rate of false-negative is underestimated as such reportedly benign cases will only be diagnosed when they recur or metastasize (29).

A concordance study performed in Rhone-Alpes comparing primary diagnosis and systematic review by expert showed that 46% of diagnosis were modified at second reading and up to 19% of cases showed a discordance in the histological subtype (31). It was also found that only 56% had full concordance between primary diagnosis and second opinion with about 19% having a complete discordance. A centralized pathological review providing rapid and efficient help with access to molecular biology analysis is of vital importance in these rare tumors (31).

According to figures from the Royal College of Pathologists (RCP), in 2010, there were almost 1,100 diagnostic practicing pathologists in the UK. Assuming that the number of sarcoma cases was evenly distributed, each pathologist would see fewer than 2 cases of STS annually (29).

And with so many different histological subtypes, it may be nearly impossible for a single general pathologist to encounter all subtypes in adequate numbers in one lifetime (45). This argues strongly for the training of an expert sarcoma pathologist in each MDT. The European Society for Medical Oncology (ESMO) acknowledges the need for accurate diagnosis and states that a pathological expert second opinion is recommended in all cases where the original diagnosis was made outside of reference sarcoma centres (46). In the US, the National Comprehensive Cancer Network (NCCN) guidelines recommend that the pathologic assessment of biopsies should be carried out by an experienced sarcoma pathologist. This amount of experience is, however, undefined (29,47). Similar recommendations are in preparation by the RCP of Australasia. In the UK, the National Institute for Health and Clinical Excellence (NICE) issued guidelines on the management of STS (26), recommending that all STS should be either first reported or reviewed by a specialist STS pathologist. Central to this all is sub-speciality training, exposure to sarcoma cases, both in terms of numbers and difficulty, and building of pathology networks in the region to share challenging cases. Anchored by good basic sarcoma pathology training, ancillary investigations oftentimes help confirm the diagnosis (48). These tests include immunohistochemical (IHC) stains and specific tests for molecular alterations. However, these are not routinely available and their use require experienced pathologists with relevant expertise (31). For example, the diagnosis of an atypical lipomatous tumor can be supported by demonstrating MDM2 and CDK4 amplification by Fluorescence in situ hybridization (FISH) or IHC. These are usually available only in specialized centers (29).

There is an ever-expanding array of molecular tests ranging from conventional morphology, karyotyping to deep exome sequencing (3). Morphology is basic requirement, while IHC testing is a useful tool in diagnosis and molecular studies, including chromosomal and translocation studies, are aspirational. The policies and guidelines governing use of these tests in each country needs to be carefully deliberated, weighed against the access, availability and effectiveness of therapeutics to allow for cost-effective medicine across countries. To this end, the Asia-Pacific sarcoma group put together a set of pathological recommendations according to individual country’s resource availability (49). In the basic clinical setting, morphologic assessment with haematoxylin and eosin stains (H&E) is recommended. While in the limited setting, H&E stains...
assisted by IHC workup would be available. In the enhanced setting, sarcoma-related translocation studies would be performed to aid in diagnosis. And finally in the most ideal of settings, histological examination would be further assisted by genomic analysis to guide molecularly-targeted treatment and research (49).

**Surgeon**

Surgery is a key component of the curative multi-modal treatment of localized STS and in selected patients with metastatic disease. While the aim of oncological surgery is good negative margins, there is no universally agreed numerical cut-off, though 1 cm is oftentimes suggested (7,50). Challenging this mission is the rarity of tumor, ubiquitous primary location and potentially large tumor. Additionally, many STS are incidentally discovered following an excision of what was supposed to be a benign lump (“whoops surgery”). This initial excision is usually inadequate and a repeat proper oncological surgery is required. As such, delays in diagnosis is not uncommon (51). An American observational study of patients with STS noted that in non-specialist treatment centres, 59% of surgical resections reported as “wide” were found to contain residual disease on specialist pathological review (51).

To achieve best surgical outcomes for patients, key strategies include primarily, formal sarcoma surgical training, secondly, raising disease awareness amongst the public and fellow healthcare professionals to avoid “whoops surgery”. Taking into consideration of each country’s resources, there would be differences expected with types of surgical treatment availability. Basic surgical care may include amputation or wide resection by a general surgeon while maximal care would include surgeries being done by a dedicated multidisciplinary team (49).

**Medical oncologist**

The sarcoma medical oncologist has intimate knowledge of the disease particularly at the molecular level as well as the rapidly developing armamentarium of systemic treatment options personalized to the individual patient. While the role of the medical oncologist may be more limited in the setting of adjuvant treatment in STS (52–54), he/she occupies a key role in the palliative setting, directing care and managing systemic treatment.

Due to the complexity, geography and widely varying affluence of countries across Asia, no single uniform guideline exists to inform of chemotherapeutic options. First-line palliative chemotherapy option remains an anthracycline-based therapy and is accessible to most patients. Second-line and beyond is more heterogeneous and is subjected to healthcare reimbursements within each country. According to the set of resource-stratified guidelines drawn up by Asia-Pacific sarcoma investigators (49), the most basic of chemotherapy availability would include vincristine, cyclophosphamide, doxorubicin and dacarbazine (49). At the other end of the spectrum, would be agents such as pazopanib and trabectedin in addition to the more common drugs such as ifosfamide, gemcitabine, docetaxel and paclitaxel (49).

**Radiation oncologist**

(Neo)-adjuvant radiation is standard of care for resected high-risk STS of the extremities though its role in retroperitoneal sarcoma is less clear and the subject of a large international study (STRASS study) (55). Radiation treatment is accessible in different forms across Asia. Radiation therapy at the most basic level would include a simple field setup with a cobalt machine; in a limited setting delivered through linear accelerators. And in the maximal and enhanced settings the availability of intensity modulated radiotherapy (IMRT)/brachytherapy and proton beam/carbon ion capabilities respectively (49,56).

**Supportive and palliative care**

The diagnosis of sarcoma involves complicated treatment processes and decision-makings with potentially life-changing functional outcomes. Many patients find it helpful when they are able to speak to patients who have gone through similar treatments. Practical needs resources are important to enable patients to continue living with their functional capability maximized.

Patients and their carers often need a variety of support, from symptom-management, to assessing social care and benefits, as well as specific needs such as orthoses and prosthetics (51). Psychological support is also important and the development of sarcoma-specific self-help groups and support groups should be encouraged (51). The 2004 Sarcoma UK survey found that sarcoma patients were not routinely offered formal psychosocial support, but those who attended counselling found it useful (51). Evidence from three systematic reviews considered in the NICE guidance on “Improving supportive and palliative care for
adults with cancer” suggests that psychosocial interventions are useful for the reduction of anxiety in cancer patients (51,57). In two small observational studies, peer support programmes were also viewed positively by patients, with many having decreased anxiety and depression (51). However, only 15% of the respondents to the Sarcoma UK survey had attended a sarcoma-specific patient support group (51).

Improved integration of palliative care services throughout the course of the illness will enhance quality of life for both patients and their carers (51). Integrated care is particularly important at the end of life, and the contribution of palliative care specialists will help to create a more appropriate balance between efforts to preserve life and the need for comfort, peace and support for close family members when it becomes clear that death is inevitable (51).

Adolescent and young adult (AYA) sarcomas

There is limited data on the incidence and epidemiology on AYA-specific sarcomas, largely contributed by the difficulty in obtaining such data. According to the SEER Database 1975–1999, approximately 70% of sarcomas diagnosed were in the age group 16–39 years old (58). Sarcomas comprise 10–20% of cancers in the young and the overall mortality is about 50%, among the worst for any cancer type in AYAs (59,60). Preliminary data from our centre showed that in the period between 2002 and 2016, we accrued 344 AYA sarcoma patients into our prospective database. Twenty-two percent had bony sarcomas, most commonly in the 16–19 years old age group, and 78% had STS, most commonly in the 35–39 years old age group. Among AYA STS patients, the 3 most common sarcomas subtypes were synovial sarcomas, liposarcomas and dermatofibrosarcoma protuberans (61). Up to 40% was lost to follow-up.

Treatment outcomes in AYA patients have not improved significantly nor kept pace with the rest of oncology over the past 20 years (62). There is evidence to suggest that AYAs with sarcomas do substantially worse than children. 5-year overall survival for both osteosarcoma and EWS in AYAs between 1983–2003 is 45%, compared to 76% for children under 15 (60). There are specific concerns with this age group, with psychological distress being significantly greater compared to the other counterparts. They face serious problems related to loss of fertility, disruptions of education and careers, limitations in social interactions, psychosocial distresses in addition to the threat to their mortality. Some of the more straightforward issues such as adhering to clinic visits and treatment schedules can be made more complicated in an AYA patient as they balance between managing their young careers and staying on treatment (62). Additionally, AYA sarcomas arising from limbs may entail aggressive surgery including amputation leading to psychological distresses, body image issues as well as physical limitations.

In a recent UK survey of an AYA group, 42% of those with STS said they had visited their general practitioner more than 5 times before they were referred to a hospital (51). Similarly, in an American study, AYA sarcoma patients reported a mean of 4.85 physician visits before a referral to a specialist unit for a bone or STS was made (51). This clearly highlights the issue of delayed diagnosis in AYA sarcoma patients and the urgent need to raise awareness amongst the public and fellow healthcare workers about this disease. The field of AYA is in its infancy, even more so in Asia. Having an AYA service in the sarcoma MDT greatly enhances care to patients.

Genetics service

Various cancer syndromes are associated with sarcomas, including Li-Fraumeni, familial adenomatous polyposis and Gardner’s syndromes. The incidence rates of such genetic diseases in the Western population are well published but is less known in Asia. Germline sarcoma mutations in patients were found to be more common than previously thought. The largest study to-date, the International Sarcoma Kindred study (ISKs), performed in 1,162 probands of both Caucasian and Asian descent, interrogated germline mutations in sarcoma patients and found that 55% of sarcoma patients harbor putatively pathogenic monogenic and polygenic variation in known and novel cancer genes (63). While work from our group focused on a cohort of 66 Asian patients younger than 50 years of age and found that 13.6% of patients had at least one predicted pathogenic germline mutation (64). Collectively, these data suggest that germline mutation in sarcoma patients may be higher than anticipated and lends strong support to the inclusion of a cancer genetics service into the sarcoma MDT.

Accreditation process

Accreditation, at the level of the MDT as well as individual clinical services, is an essential step to ensure standards are maintained and guidelines are adhered to. For example, the NICE guidelines recommend that an STS pathologist is one
who regularly reports STS as a significant component of his/her workload. In addition, the pathologist should participate in external quality assessment (EQA) and be a member of a properly constituted sarcoma MDT. The MDT should also meet requirements to ensure that standards are achieved and maintained. This can be potentially audited at the basic level by the minimum number of patients discussed/ managed per year and complexity of cases discussed (51).

**Treatment guidelines**

Several international sarcoma guidelines are available and commonly used by practicing clinicians. They include the NCCN and ESMO guidelines (4,46). However, these international guidelines do not take into account resource limitations that exist in Asia, between countries, and sometimes within disparate geographical locations within a country. As alluded to earlier, investigators in the region published a set of resource-stratified guidelines for the management of sarcoma in the Asia-Pacific region, providing a framework of care at each resource level and serve as a roadmap for sites to aspire to in their developmental process. In Singapore, there also exists the existence of the SCAN (Singapore Cancer Network) guidelines to provide some guidance on the diagnosis and management of sarcomas (7).

**Sarcoma collaborations**

STSs are rare and heterogeneous tumors. To improve clinical care, education and research, one needs to exploit opportunities, both regionally and internationally for collaboration.

**Clinical care collaboration**

Given the complexity and heterogeneity of sarcoma, the management requires integrated care. The ability to strengthen international collaborations result in maximization of resources, thereby improving care for our patients (65). This allows better and greater access to drugs across Asia, and also allow a better understanding of how these drugs work for our patients. In addition, international collaboration also allows sharing of the most up-to-date diagnostic and treatment possibilities, ultimately benefitting the patient.

In the real world, the SCE complete with its MDT do not exist on its own. It is the hub which directs and strengthens all sarcoma-related care activities in the community. Hence building a strong collaborative network and expedited referral process (where necessary) are key factors in improving patient care. Collaboration allows integration pre-and-post operatively to allow precise coordination of chemotherapy, radiotherapy and surgery (66). This approach not only allows best care for patients (24,67), but at the same time allows for the concentration of these rare cancers into specific centres to continually hone the skills of sarcoma clinicians.

**Educational collaborations**

Education in sarcoma can be looked upon at three levels, namely general public, community health and professional levels.

For the public, efforts have been largely singular and country-specific. There needs to be proper public education to minimize patient-related delay in diagnosis. In a Belgian study, 47% of STS patients had a delay of more than 1 month before seeking medical advice, with a median delay of 4 months. In a Dutch study, 36% of patients with retroperitoneal STS waited for more than 6 months before seeing a doctor (51).

At the community health level, inaccurate and late diagnoses lead to mismanagement in 70% of patients with a number receiving wrong treatment (2). It is therefore important to raise awareness and knowledge of sarcomas within the community (2). Non-sarcoma specialists need a reliable source of information and network of care for their patients (51). Various efforts have been made to raise sarcoma awareness in healthcare and allied health professionals, but these have not been uniform. It is estimated that a GP is only likely to see one or two sarcomas in his/her career (24). In a study of referrals to a UK specialist STS unit, 20% had more than a 3-month delay, with the median being 14 months. The most frequent reason was a lack of clinical suspicion. A second UK study showed that a referral from the primary doctor took 7.5 months (51). Educating our fellow colleagues will hopefully decrease referral delays.

At the professional level, there exists an unmet need across Asia to have sarcoma-specific forums where sarcoma experts and specialists meet to share knowledge on the subject. There is also a lack of preceptorships where trainees can learn from. Such meetings are beneficial to create awareness and educate our fellow colleagues.

The biannual Asia-Pacific Musculoskeletal Tumor Society (APMSTS) meeting, with an orthopedic oncology
focus, is one such example. The annual Singapore Sarcoma Symposium, initiated in 2013, focuses on a broad multi-disciplinary range of sarcoma topics, for the local and regional sarcoma community. These efforts are invaluable in promoting the sharing of knowledge and education of our fellow healthcare professionals and allied health partners. Additionally, sarcoma-specific preceptorship programs catered for practicing clinicians are highly effective tools for sharing state-of-art medical knowledge with fellow colleagues. To this end our group hosted 2 programs in 2017. The first was a clinically focused interactive preceptorship built around an MDT framework on the various sarcoma subtypes. While the other was a sarcoma pathology focused event for regional and international sarcoma pathologists to share, learn and network, paving the way for future collaborations.

One commendable healthcare professionals-targeted education effort would be the Sarcoma Policy Checklist. This is a joint effort by a multi-stakeholder group of experts from the medical, patient advocacy and pharmaceutical industry fields. It aims to help policymakers close the gap in access to high quality information and care for sarcoma patients across Europe (2). In the UK, EQAs are as of the UK National Orthopedic Pathology EQA scheme. The scheme provides 2 circulations a year with relevant questions to encourage ongoing education (26). Training should be developed and provided for all members of both the core and extended sarcoma MDT (51).

**Research collaboration**

In research of rare tumors like sarcoma, collaboration is key. Management of sarcomas is difficult due to limited information on natural history. Continuous prospective long-term databases are important to obtain knowledge for rare tumors (68). This is consistent both in the Western and Asian populations. Currently there is a paucity of prospective data coming out of Asia (6). Across Asia, national registries are limited for rare tumors like sarcoma. Current Asian studies tend to be small, are usually single-centred and retrospective. This makes it difficult to draw firm conclusions.

Research is made even more challenging by the heterogeneity and reproducibility in histological diagnoses. To produce robust Asian data with large sample size, there exists a need to build regional and international collaborations to pool and share data. Of particular interest would be genetic differences in Asian patients with regards to disease incidence, response and tolerance to treatment.

In this respect the Asian Sarcoma Consortium (ASC), was formed to bring together the Asian sarcoma research community. In its inaugural efforts, the group reported on the epidemiology and real world treatment of a large cohort of 423 patients with angiosarcoma from 8 sites in 6 Asian countries (69). Additionally the STAR study (5) was an Asian multi-centred observational study initiated to describe epidemiology, treatment patterns and clinical outcomes in STS patients diagnosed between 2006–2010. More efforts must be placed into building formalized research collaborations within Asia to advance the science of sarcoma.

**Conclusions**

In summary, STS is an uncommon and heterogeneous group of malignancies. Due to the unique and disparate socio-economic statuses and healthcare systems of the individual countries in Asia, it has made understanding this disease in Asian context even more difficult. Most data present are largely country-specific and retrospective.

We propose the development of SCEs across Asia. This will hopefully allow high-quality sub-specialized sarcoma care to patients, especially to areas where access may particularly be difficult. SCEs should include:

- MDTs;
- Pathologist;
- Surgeon;
- Medical oncologist;
- Radiation oncologist;
- Supportive and palliative care;
- AYA care;
- Genetics services.

SCEs should be responsible for the development of region-specific treatment guidelines and the accreditation process. This will ensure standards are maintained and guidelines adhered to.

In order to advance sarcoma care in Asia, no individual country can do this alone. There are multiple avenues possible for collaboration, and these should be done regionally and internationally:

- Clinical care collaboration;
- Educational collaborations (general public, community health and professional level);
- Research collaboration.

This report serves to highlight the challenges and opportunities of sarcoma in Asia. Asia is huge and diverse.
While the inherent geographical and socio-economic factors may be outside the scope of this report, there are however many opportunities to improve clinical care, expertise, education and research in sarcoma within Asia. The sarcoma MDT within centres of excellence serves as the foundation from which we build upon. National guidelines to set standards of care and prospective databases to capture real world data are useful tools to understand and better manage this disease. While the ASC has paved the way in the past 5 years with fruitful collaborations, more will need to be done in order to ensure advancements in sarcoma care continue. Moving forwards, formalized national and Asia-wide research collaborative networks should be set up to provide the much-needed platforms for us to study this unique disease in the Asian setting.

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References
and-affordable/


47. NCCN. NCCN Practice Guidelines in Oncology, Soft Tissue Sarcoma v2. 2010.


55. EORTC. Surgery With or Without Radiation Therapy in Untreated Nonmetastatic Retroperitoneal Sarcoma (STRASS).


66. Sarcoma Care: Meeting of the Minds, 2016.


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