Economic evaluations were developed more than 50 years ago to assist those responsible for allocating resources to prioritize alternatives with the maximum benefit at the lowest cost, as health care resources were, and remain, scarce. Economic evaluations provide a decision-making framework in which outcomes (benefits) and costs are assessed for various alternative options. Although the interest in complete and partial economic evaluations has increased over the past 2 decades, the quality of studies has been marginal due to methodological challenges or incomplete cost determination. This paper provides an overview of the main types of complete and partial economic evaluations, reviews key methodological elements to be considered for any economic evaluation, and reviews concepts of cost determination. The goal is to provide the clinician neurosurgeon with the knowledge and tools needed to appraise published economic evaluations and to direct high-quality health economic evaluations.

Overview of Complete Economic Evaluations

Four main types of complete economic evaluations are currently used: cost minimization analysis (CMA), cost-effectiveness analysis (CEA), cost-utility analysis (CUA), and cost-benefit analysis (CBA). The most critical initial step is to formulate the study question; this will guide selection of the appropriate type of economic evaluation. Reviewing in depth each type of economic evaluation would be beyond the scope of this paper; instead, we will review key methodological elements to be included in any economic evaluation and concepts of cost determination.
evaluation is beyond the scope of this overview. A concise description of each type follows. Table 1 presents a summary of the 4 main economic evaluations.

**Cost Minimization Analysis**

In a CMA, the alternative options are known to have the same effectiveness (efficacy or clinical outcome). The costs of the 2 or more procedures or programs under consideration are compared to establish the lowest cost alternative. It is essential that the same or similar effectiveness has been demonstrated by a clinical study that is specifically designed to assess the equivalence of treatments. Studies have demonstrated that the drugs had equivalent effectiveness in controlling seizures, and therefore the authors performed a CMA to compare only the direct costs of similarly effective therapies. Published CMAs are rare in the neurosurgical literature as outcomes are most frequently different between two alternative options.

**Cost-Effectiveness Analysis**

In CEA, the clinical outcomes and costs of each procedure or program are assessed and compared. The costs are expressed in monetary units and the outcomes in nonmonetary natural units, common to all alternatives. Examples of clinical outcomes include symptom-free days, specific clinical parameters (e.g., remission rates in functional pituitary surgery), hospital days prevented, and life years gained. Identification of the most appropriate outcomes to serve as effectiveness end points is crucial. Selected outcomes often provide a narrow view, focusing on one condition or specific care pathway, and limiting the use of CEA to guide allocation of resources across different health programs.

Karinen and colleagues assessed the cost-effectiveness of nimodipine administration in patients who presented with a subarachnoid hemorrhage and underwent aneurysm clipping. The authors used direct costs for the hospitalization, and the nonmonetary clinical outcome was measured as life years gained. They found that patients receiving nimodipine had an average of 3.46 years of longer life expectancy compared with patients receiving placebo. The “cost-effectiveness” of one alternative option can be compared with that of the benchmark alternative by calculating the incremental cost-effectiveness ratio (ICER): (Costalternative − Costbenchmark) / [Outcomealternative − Outcomebenchmark]. In this study, the ICER was $223 per life years gained; nimodipine increases patient life years at a low incremental cost.

As another example of CEA, Vuong and colleagues assessed the cost-effectiveness of 2 brain metastatic treatment modalities, namely resection versus stereotactic radiosurgery from the perspective of Germany’s health system. For both treatment strategies, only costs of initial intervention and costs of retreatment (including repeat surgery, stereotactic radiosurgery, linear accelerator, or whole-brain radiotherapy) were calculated until death or 5.5 years of follow-up. The nonmonetary clinical outcome assessed was life year saved. In this study, use of stereotactic radiosurgery cost €1683 less than resection and increased the number of life years saved by 0.45 years. There are numerous examples of CEA in all neurosurgical subspecialties.

**Cost-Utility Analysis**

Cost-utility analysis follows the same framework as CEA. In fact, CUA is viewed by many economists as a subtype of CEA. The costs are expressed in monetary units. The outcomes are utility adjusted, where utility refers to a patient’s preference for a particular health outcome. The outcome is expressed in quality-adjusted life years (QALYs), where the length of time spent in one health state is multiplied by the utility value of that health state (ranging from 0 [death] to 1 [perfect health]). Generic preference-based measurement systems such as the EuroQol 5 Dimension (EQ-5D) and the Short Form 6D (SF-6D) have facilitated the determination of the utility value. Cost-utility analysis presents cost per QALY gained by selecting one procedure or program over another. This type of economic evaluation is especially useful when mortality difference is not expected between interventions and/or quality of life is an important aspect of one treatment alternative. In comparison with CEA, CUA enables comparison of different interventions across numerous diseases, as a QALY has equal value irrespective of the nature of the alternative that delivers it. In North American studies, the cost/QALY threshold typically used to make resource allocation decisions varies from $50,000/QALY to $100,000/QALY. Figures of £20,000 to £30,000 per QALY are also used in the United Kingdom.

Parker and colleagues questioned if minimally invasive versus open multilevel hemilaminectomy in patients with degenerative lumbar spinal stenosis was cost-effective. In this CUA, the authors described how they estimated the hospital costs, the physician costs, and indirect costs related to productivity losses. The authors used a

<table>
<thead>
<tr>
<th>Type of Economic Evaluation</th>
<th>Outcome Measure</th>
<th>Cost Measure</th>
<th>Pearl</th>
</tr>
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<tbody>
<tr>
<td>CMA</td>
<td>none (considered stable between alternatives)</td>
<td>monetary unit</td>
<td>need to be certain both alternatives provide the same outcome</td>
</tr>
<tr>
<td>CEA</td>
<td>any nonmonetary natural unit (disease specific)</td>
<td>monetary unit</td>
<td>inability to make interdisease comparison</td>
</tr>
<tr>
<td>CUA</td>
<td>utility score (nonmonetary)</td>
<td>monetary unit</td>
<td>capable to make interdisease comparison</td>
</tr>
<tr>
<td>CBA</td>
<td>monetary unit attributed to health effects</td>
<td>monetary unit</td>
<td>not frequently used in medicine</td>
</tr>
</tbody>
</table>
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standard utility assessment, the EQ-5D, to estimate the QALYs before and at 2 years postprocedure. Although minimally invasive and open multilevel hemilaminectomy were associated with similar mean cost over 2 years (minimally invasive surgery: US$23,109 vs open hemilaminectomy: US$25,420), both treatment strategies provided equivalent improvement in QALYs (0.72) 2 years after surgery.26 Reporting findings of cost equivalence are of significance, as they may inform preconceived impressions of physicians.

As another example, Kumar and Rizvi evaluated the utility of spinal cord stimulation (SCS) and conventional medical management (CMM) compared with CMM alone for patients with multiple conditions including failed back surgery syndrome and complex regional pain syndrome.21 Outcome (QALY) was assessed at baseline and at 6 months using the EQ-5D.28 For each base case, the various categories of costs were tabulated for each treatment option (SCS+CMM vs CMM alone), including the cost of the implantable devices. In this study, the ICER for SCS+CMM was CAD$9,293 (Canadian dollars) per QALY gained in failed back surgery syndrome and CAD$11,216 per QALY gained in complex regional pain syndrome.21 Importantly, the ICER may vary between study groups, depending on the difference in QALYs between SCS+CMM versus CMM alone and the difference in cost between both treatment strategies.

Last, Ho and colleagues assessed the cost-effectiveness of decompressive craniectomy performed in patients with refractory intracranial hypertension after severe traumatic brain injury.13 The authors detailed which costs were tabulated and how QALY gained at the 18-month follow-up was calculated. This study showed that the average total hospital costs per QALY gained increased significantly when the predicted risk of a poor outcome was greater than 80%: US$682,000 if predicted risk > 80% versus US$140,000 if predicted risk ≤ 80%, which is more than the usual accepted value of US$100,000 deemed cost-effective.15 In trauma and oncology, the utility of a procedure can be difficult to rationalize solely on the information provided by economic evaluations (which integrate benefits and costs).9,14

Cost-Benefit Analysis

In CBA, the outcomes and the costs are expressed in monetary units. The human capital approach and the willingness to pay approach are 2 methods used to quantify in a monetary unit how much patients value improvement of their health (outcome).11 A person’s willingness to pay may be influenced not only by personal preferences and values, but also by income and state of health.32 This type of economic evaluation is seldom used in the medical literature given the difficulty in placing a monetary value on various health states.29 To our knowledge, no study has reported a true CBA as described above regarding a neurosurgical treatment. Some papers include the key words “costs” and “benefit” in their titles but are in fact CEA.

True CBAs have been performed in the context of drug administration. Tang and colleagues conducted a cost-benefit analysis of administering ondansetron (Zofran) to prevent emesis in a high-risk patient population undergoing outpatient surgery.20 The overall weighted costs per patient for postoperative nausea and vomiting management were US$21 ± $7. The authors used the “willingness to pay” method to attribute a monetary value of the benefits of preventing emesis. On average, patients were willing to pay US$117 ± $82 to prevent postoperative nausea and vomiting if they underwent a similar surgery in the future.29 Dividing the assigned monetary value of “benefits” by the assigned cost results in the benefit-to-cost ratio, noted to be 5:1 in this study.

As a second example, Wasserfallen and colleagues assessed parents’ willingness to pay for local anesthetics during blood sampling in children.32 The unit cost of the local anesthetic used in this study was €5.6. Among the various scenarios proposed, parents were willing to pay approximately €4–€7, testifying that the use of this drug has a high societal value.32 This type of information, as noted by the authors, could be used to support billing of the drug to the parents through the health care facility or argue in favor of having the health care system absorb its cost as it delivers painless care.

Existing Guidelines for Economic Evaluation Studies

As neurosurgeons engage in the value paradigm, lead various economic evaluations, and share their findings with the scientific community, reviewing existing guidelines for economic submissions is of great value. Many societies and journals have described specific guidelines, most inspired by those put forth by Drummond and collaborators of the BMJ Economic Evaluation Working Party.5,17 They are useful for both authors and peer reviewers of economic submissions.8,17 A detailed review of the guidelines is beyond the scope of this overview. Table 2 summarizes key elements to be considered and reviewed when preparing either an economic evaluation or a cost analysis.8,17,28

Partial Economic Evaluation

When outcomes are not considered in the analysis along with the cost data, the economic evaluation is qualified as partial. Studies that include a cost analysis report the cost incurred for either one specific part of the treatment (e.g., surgery-related costs), one phase of the episode of care (e.g., costs of inpatient care), or attempt to cost the entire episode of care.

The cost analysis is performed only for one procedure or program at one specific time. A partial economic evaluation in the form of cost analysis is the first step in conducting some form of complete economic evaluation.

Mendez and colleagues analyzed the costs of treatment of glioblastomas in Nova Scotia, Canada.23 This study was timely for 2 reasons: 1) Health care costs of illnesses with high mortality rates were, and remain, of interest in times of constrained health care resources. 2) At the time of publication, no information regarding cost of care of these patients existed in Canada.23 The authors provided a detailed description of their cost determination methodology and presented a breakdown of costs.
In the health care accounting setting, costs refer to the monetary value of resources used in performing an activity. Costs can be qualified as either direct versus indirect and variable versus fixed. Direct costs are defined as costs charged specifically to a revenue-producing cost center. Examples include radiology, laboratory, and pharmacy costs. These costs are most closely related to providing patient care and are most directly impacted by care redesign initiatives. Indirect costs are costs allocated to revenue-producing centers from nonrevenue departments, which support patient care. Examples include nutrition, administration, and patient escort. Of importance, the term “indirect cost” has a significance when used in an economic setting that is different from that in an accounting setting; it then includes changes in productivity resulting from morbidity (wait time, loss of earnings) or mortality. To prevent confusion, the term “productivity cost” is preferred.

Overall, there are 4 basic unit costs: variable direct, fixed direct, variable indirect, and fixed indirect. Variable costs refer to costs that change when volume changes versus fixed costs, which refer to costs that do not change when the volume changes. As an example, the cost of incurred during the inpatient and outpatient care of glioblastoma patients. From the time of diagnosis to death, the mean cost of medical care was CAD$17,149 per patient. The cost analysis enabled observation that highest costs were related to hospitalization with ward costs alone accounting for 48% of all costs.25

Once a potential area is identified for intervention, a CMA could be conducted between the benchmark approach and the proposed intervention assuming that the outcome is the same with both alternatives. Cost analysis can also be used to follow cost of treatment trends over time, especially when coupled with care redesign initiatives.24

### Cost Determination for Economic Evaluations

Cost information can be obtained from financial departments within health systems. Hospital-related costs and physician service costs may be managed by two distinct departments. Outside of a clinical trial, quantity of resources used and cost per unit of resource used should be based as much as possible on real patients included in the study. Physicians involved in health economics should build and nourish collaborations with their institution’s finance department and learn to speak the same accounting language. They need to recognize the difference between costs and charges and understand the different subsidiary components of total costs.

The extent of cost determination is guided by the study question and whose perspective is being considered. Performing an economic evaluation from a patient’s perspective (narrowest) versus the society in general (broadest) will influence the extent of included costs.27 Although the societal perspective may be the gold standard, capturing the data on all costs incurred is nearly impossible. For this reason, and because provider cost data are more readily available, the provider’s perspective is usually used.

### Hospital-Related Costs

In the health care accounting setting, costs refer to the monetary value of resources used in performing an activity. Costs can be qualified as either direct versus indirect and variable versus fixed. Direct costs are defined as costs charged specifically to a revenue-producing cost center. Examples include radiology, laboratory, and pharmacy costs. These costs are most closely related to providing patient care and are most directly impacted by care redesign initiatives. Indirect costs are costs allocated to revenue-producing centers from nonrevenue departments, which support patient care. Examples include nutrition, administration, and patient escort. Of importance, the term “indirect cost” has a significance when used in an economic setting that is different from that in an accounting setting; it then includes changes in productivity resulting from morbidity (wait time, loss of earnings) or mortality. To prevent confusion, the term “productivity cost” is preferred.

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###TABLE 2: Summary of key elements to review when preparing an economic evaluation or cost analysis

<table>
<thead>
<tr>
<th>Study Design</th>
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<tbody>
<tr>
<td>1) study question</td>
</tr>
<tr>
<td>2) selection of alternative options</td>
</tr>
<tr>
<td>3) type of economic evaluation</td>
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<table>
<thead>
<tr>
<th>Data Collection</th>
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</thead>
<tbody>
<tr>
<td>4) effectiveness data (clinical outcome)</td>
</tr>
<tr>
<td>5) “benefit” measurement &amp; valuation</td>
</tr>
<tr>
<td>6) costing measurement</td>
</tr>
<tr>
<td>7) modeling</td>
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</tbody>
</table>

<table>
<thead>
<tr>
<th>Analysis &amp; Interpretation of Results</th>
</tr>
</thead>
<tbody>
<tr>
<td>8) adjustments for timing of outcomes &amp; costs</td>
</tr>
<tr>
<td>9) allowance for uncertainty</td>
</tr>
<tr>
<td>10) presentation of results</td>
</tr>
</tbody>
</table>
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contrast is variable, as the more head CTs with contrast are done, the more the cost of this resource. However, the cost of the CT scanner is fixed, because the cost does not change when the number of head CT studies changes. In general, economists agree that cost analysis should preferentially use direct costs.

In the absence of cost data, various strategies have been developed to provide approximation of costs. The most frequent is to use interchangeably cost and charges. However, charges and costs are not equivalent. Charges refer to the expenses posted to a patient account for hospital services. Another common approximation for cost is the use of cost-to-charge ratios, which modifies the charges using a ratio reflecting historical proportional differences. Its interpretation is challenging: a low cost-to-charge ratio can represent either lower costs or higher charges. In addition, the cost-to-charge ratio can vary tremendously within the same institutions for different services. There is no substitute for cost data for economic evaluations.

Physician Reimbursement (Not Cost)

The physician reimbursement from Center for Medicare and Medicaid Services relies on appropriate coding for the diagnosis and the service provided. The reimbursement fee is based on the resource-based relative value scale. The relative value units are multiplied by a geographic practice cost index and then converted to dollars by a conversion factor that changes annually. Similarly, private insurers have established various methodologies to calculate their own reimbursement fee schedule, with multiple influencing factors such as geographic location and regional demand for subspecialty.

Importantly, monetary values obtained for physician services are “reimbursement fees.” This is different from the charges and also from the costs. As a general rule, the charges are the most expensive, the costs are the least expensive, and the reimbursement is somewhat in between. Adding hospital cost (or charges as a suboptimal surrogate for cost) and physician reimbursement, although not optimal, might be the closest to assess total cost of care delivery using patient-specific data.

Costs From National Databases

Depending on the study question and on access to cost data within the institution, the Nationwide Inpatient Sample (NIS) may be a resource for economic data. The NIS is one of the databases developed for the Healthcare Cost and Utilization Project (HCUP), sponsored by the Agency for Healthcare Research and Quality. The NIS approximates a 20% sample of discharges from HCUP-participating community hospitals within the US. The NIS contains clinical information such as age, sex, primary diagnosis (ICD-9 diagnosis), comorbidity measures, and length of stay. It also holds nonclinical information such as hospital characteristics (location, teaching/nonteaching, rural/community, and low/high volume) and expected payment source. The NIS provides total charges (not costs) for inpatient care only; it does not include charges (or costs) for physician services. In addition, costs related to outpatient clinic, rehabilitation, or long-term acute care hospitals are not included. Longitudinal studies are possible using the NIS database as it holds more than 20 years of data. However, this may be biased as only a subset of hospitals have been NIS members for multiple continuous years. Careful selection of data elements and preanalysis with descriptive statistics are essential to highlight missing data and hospital-specific data anomalies, as these can compromise the quality of the analysis.

The University Health Consortium offers access to multi-center economic data to its members, using cost-to-charge ratios to generate their cost data. Crimson Continuum of Care, a business analytical tool, can also be used to analyze economic data. Each participating institution determines what subsidiary of cost is fed into the database (i.e., total cost, direct cost, or direct variable cost). Since different institutions may feed different types of cost, no comparison is feasible between participating institutions. Physicians using any type of database should understand well how it is built and how to extract meaningful and sound data.

Costs for a Phase of Care Versus the Entire Episode of Care

In studies reporting on complete economic evaluations or cost analysis, the cost determination is most often captured for a specific point of care or phase of care (e.g., preoperative outpatient, inpatient hospitalization, postoperative outpatient, and readmissions). However, there is a movement toward assessment of cost incurred to deliver care throughout the entire episode of care. It would be challenging to calculate the cost of the entire episode of care using current cost determination methods. The development of accountable care organizations will help facilitate this task by consolidating the care delivered to a patient population within one health system, using one constant costing method.

A promising accounting method called the time-driven activity-based costing method, developed by Kaplan et al., is being piloted in various specialties and institutions throughout the nation. It integrates the cost of each resource used in the process and the quantity of time the patient spends with each resource. This model has the potential to capture all costs (hospital- and physician-related costs) incurred throughout the episode of care using one consistent methodology. It is being piloted by some specialties to prepare bundle pricing estimates.

Impact of Accessible Clinical Data on Economic Evaluations

Health information technology has significantly spread over the last 2 decades; 59% of US hospitals now have at least a basic electronic health record. In addition to not yet being widely adopted, modern health information technology systems are most often not used to their maximal capacity regarding communication, coordination, measurement, clinical decision support, and hard-wiring of care pathway redesigns. Furthermore, current health care payment models, such as the fee-for-service model, might be counterproductive to health information technology’s potential favorable economic impact, resulting in increased hospital charges.
awaits all institutions as they transition from ICD-9 code sets to ICD-10 code sets, which requires more detailed documentation. From a national health care budget perspective, it will be key to mitigate the risk of increased charges due to changes in the documentation process (but not representing different care delivery). In the future, it may be pertinent to report in health economic analysis various payment models, electronic health record status, and coding system used, especially if 2 groups are compared. Overall, access to data, clinical or financial, is the first step. Physicians will need to have a comprehensive understanding of the data they are using to guide care redesign and obtain the greatest health benefits and cost savings.

Future Perspectives

Economic evaluations and cost analysis are essential for the future of health care delivery. High-quality and methodologically sound economic evaluations are needed to confidently steer decision-making processes and care redesign initiatives toward optimal value of care. A multidisciplinary team should be coordinated, including economists, accountants, finance administrators, business professors, performance excellence specialists, and care providers. Although this alignment has been lacking in many health care institutions, those leading the path on health economics have understood the priceless value of collaborative teamwork performed by this new multidisciplinary team. Just like basic science developed a bench to bedside approach, health economics needs to develop a calculator to bedside approach, seeking to translate economic evaluation findings into the care delivery setting.

Disclosure

The authors report no conflict of interest concerning the materials or methods used in this study or the findings specified in this paper.

Author contributions to the study and manuscript preparation include the following: Concept and design: McLaughlin, Martin. Acquisition of data: McLaughlin. Analysis and interpretation of data: all authors. Drafting the article: McLaughlin. Critically revising the article: all authors. Reviewed submitted version of manuscript: McLaughlin, Ong, Tabbush, Martin. Approved the final version of the manuscript on behalf of all authors: McLaughlin. Study supervision: McLaughlin.

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