Levels of Evidence for Primary Research Question.^[1,2]

| Study type | Question | Level I | Level II | Level III | Level IV | Level V |
|--|---|--|--|---|---|---|
| Diagnostic—Investigating a diagnostic test | Is this (early detection) test worthwhile? | • Randomized controlled trial | • Prospective ^[3] cohort ^[4] study | Retrospective ^[5] cohort ^[4] study | Case series | • Mechanism-based reasoning |
| | | | | Case-control ^[6] study | | |
| | Is this diagnostic or monitoring test | • Testing of previously developed diagnostic | • Development of diagnostic criteria | Nonconsecutive patients | • Poor or nonindependent reference standard | • Mechanism-based reasoning |
| | accurate? | criteria (consecutive patients with consistently applied reference standard and blinding) | (consecutive patients with consistently applied reference standard and blinding) | No consistently applied reference standard | | |
| Prognostic—Investigating the effect of a patient characteristic on the outcome of a disease | What is the natural history of the condition? | Inception^[3] cohort study (all patients enrolled at an early, uniform point in the course of their disease) | • Prospective ^[3] cohort ^[4] study (patients enrolled at different points in their disease) | Retrospective^[5] cohort^[4] study Case-control^[6] study | • Case series | • Mechanism-based reasoning |
| | | | Control arm of randomized trial | | | |
| Therapeutic—Investigating the results of a treatment | Does this treatment help? What are the | Randomized controlled trial | • Prospective ^[3] cohort ^[4] study | • Retrospective ^[5] cohort ^[4] study | • Case series | Mechanism-based reasoning |
| | harms? ^[7] | | Observational study with dramatic effect | • Case-control ^[6] study | Historically controlled study | |
| Economic | Does the intervention offer good value for dollars spent? | Computer simulation model (Monte Carlo simulation, Markov model) with inputs derived from Level-I studies, lifetime time duration, outcomes expressed in dollars per quality-adjusted life years | Computer simulation model (Monte Carlo simulation, Markov model) with inputs derived from Level-II studies, lifetime time duration, outcomes expressed in dollars per QALYs and uncertainty examined using | Computer simulation model (Markov model) with inputs derived from Level-II studies, relevant time horizon, less than lifetime, outcomes expressed in dollars per QALYs and stochastic multilevel sensitivity analyses | Decision tree over the short time horizon with input data from original Level-II and III studies and uncertainty is examined by univariate sensitivity analyses | Decision tree over the shor time horizon with input data informed by prior economic evaluation and uncertainty is examined by univariate sensitivity analyses |
| | | (QALYs) and uncertainty examined using probabilistic sensitivity analyses | probabilistic sensitivity analyse | | | |

[1]This chart was adapted from "OCEBM Levels of Evidence Working Group"* and "The Journal of Bone&Joint Surgery"**. A glossary of terms can be found here: http://www.cebm.net/glossary/.

*The Oxford 2011 Levels of Evidence," Oxford Centre for Evidence-Based Medicine, http://www.cebm.net/ocebm-levels-of-evidence/.

**http://jbjs.org/instructions-for-authors#LevelsofEvidence.

^[2] Level-I through IV studies may be graded downward on the basis of study quality, imprecision, indirectness, or inconsistency between studies or because the effect size is very small; these studies may be graded upward if there is a dramatic effect size. For example, a high-quality randomized controlled trial (RCT) should have ≥80% follow-up, blinding, and proper randomization. The Level of Evidence assigned to systematic reviews reflects the ranking of studies included in the review (i.e., a systematic review of Level-II studies is Level II). A complete assessment of the quality of individual studies requires critical appraisal of all aspects of study design.

^[3] Investigators formulated the study question before the first patient was enrolled.

^[4] In these studies, "cohort" refers to a nonrandomized comparative study. For therapeutic studies, patients treated one way (e.g., cemented hip prosthesis) are compared with those treated differently (e.g., cementless hip prosthesis). ^[5] Investigators formulated the study question after the first patient was enrolled.

^[6] Patients identified for the study on the basis of their outcome (e.g., failed total hip arthroplasty), called "controls."

^[7] Sufficient numbers are required to rule out a common harm (affects >20% of participants). For long-term harms, follow-up duration must be sufficient.