

O knee, where art thou? Patient-reported outcomes and physical performance in individuals with acute knee injury

marviduais with acute knee mjury.		

Flosadottir, Vala

2018

Document Version: Other version

Link to publication

Citation for published version (APA):

Flosadottir, V. (2018). O knee, where art thou? Patient-reported outcomes and physical performance in individuals with acute knee injury. [Doctoral Thesis (compilation), Sport Sciences]. Lund University: Faculty of Medicine.

Total number of authors:

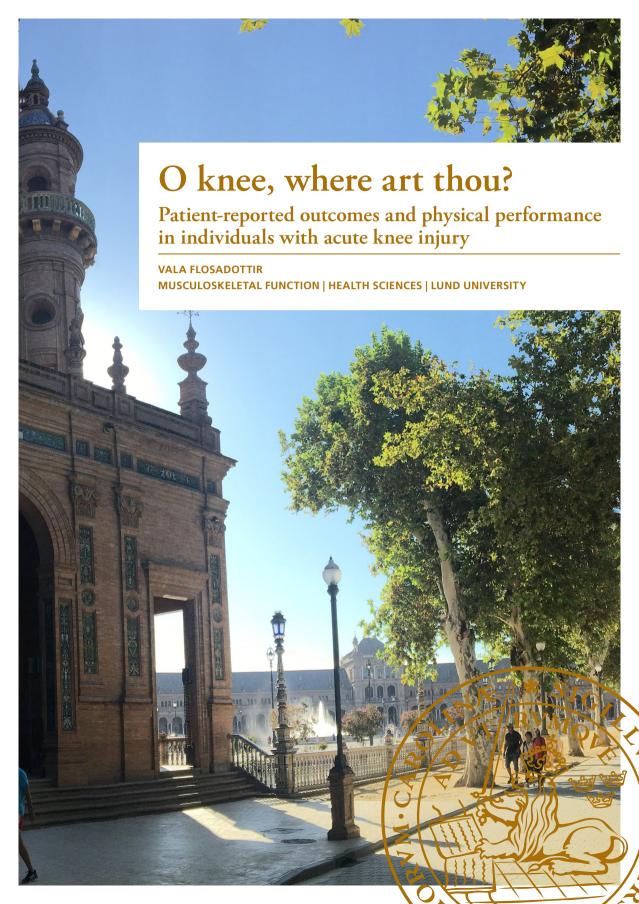
Unless other specific re-use rights are stated the following general rights apply:
Copyright and moral rights for the publications made accessible in the public portal are retained by the authors and/or other copyright owners and it is a condition of accessing publications that users recognise and abide by the legal requirements associated with these rights.

• Users may download and print one copy of any publication from the public portal for the purpose of private study

- or research.
- You may not further distribute the material or use it for any profit-making activity or commercial gain
 You may freely distribute the URL identifying the publication in the public portal

Read more about Creative commons licenses: https://creativecommons.org/licenses/

If you believe that this document breaches copyright please contact us providing details, and we will remove access to the work immediately and investigate your claim.



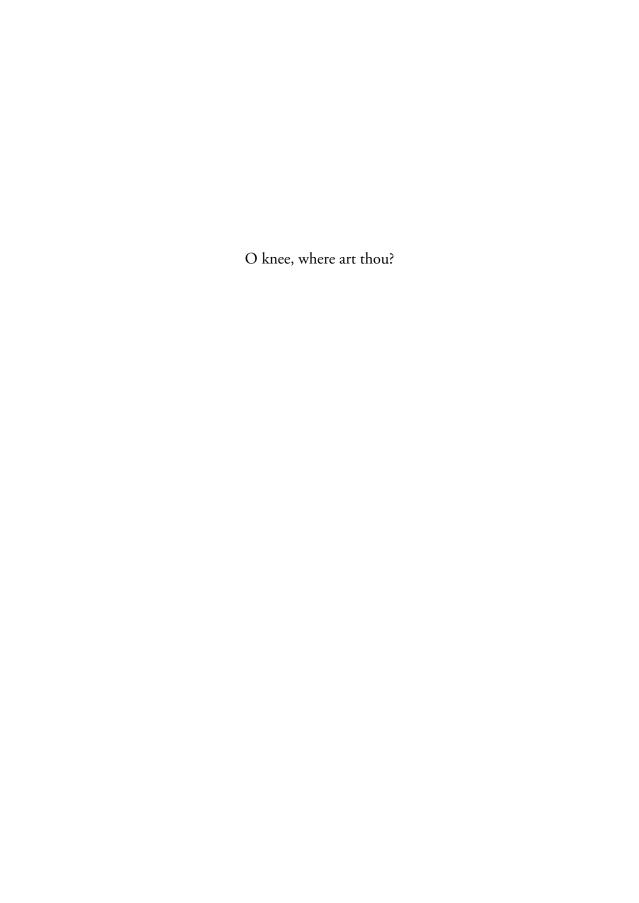




The Musculoskeletal function research group
Department of Health Sciences

Lund University, Faculty of Medicine Doctoral Dissertation Series 2018:35 ISBN 978-91-7619-602-1 ISSN 1652-8220





O knee, where art thou?

Patient-reported outcomes and physical performance in individuals with acute knee injury

Vala Flosadottir



DOCTORAL DISSERTATION

by due permission of the Faculty of Medicine, Lund University, Sweden. To be defended at the Health Sciences Centre, Lund on April 20^{th} 2018 at 09.00.

Faculty opponent
Assistant Professor Karin Grävare Silbernagel
Department of Physical Therapy, University of Delaware
Department of Orthopaedics, Institute of Clinical Sciences at Sahlgrenska Academy,
University of Gothenburg

Organization:

LUND UNIVERSITY, Sweden
Department of Health Sciences
Research group: Musculoskeletal function
Author: Vala Flosadottir

Document name:
DOCTORAL DISSERTATION

Date of issue:
April 20, 2018

O knee, where art thou?

Patient-reported outcomes and physical performance in individuals with acute knee injury.

Knee injuries are severe and common injuries in young and physically active individuals. Knee injury treatment, non-surgical and surgical, is aimed at improving patient-reported outcomes (PROs), reducing symptoms and restoring physical function. Physical performance tests and patient-reported outcome measures (PROMs) are often used in the evaluation of outcome after knee injury treatment. For this purpose, it is critical that the PROMs that are used demonstrate adequate measurement properties. Furthermore, it is important to identify modifiable factors that may be associated with improved outcome after knee injury. The first aim of this thesis is to investigate and present the measurement properties, specifically the responsiveness, of knee-specific and generic PROMs, in individuals undergoing knee injury rehabilitation. The second aim of this thesis is to investigate the associations between lower extremity physical performance and self-reported activity participation, knee function and knee self-efficacy in individuals after anterior cruciate ligament (ACL) injury or ACL reconstruction (ACLR).

Two methodological studies (I and II) were performed to assess the cross-cultural validity of the Swedish version of the Activity Rating Scale (ARS) for disorders of the knee (n=100) and to assess the responsiveness of the Knee injury and Osteoarthritis Outcome Score (KOOS), the Tegner Activity Scale (TAS), the Activity Rating Scale (ARS) for disorders of the knee, and the Knee Self-Efficacy Scale (K-SES) (n=76), respectively. Two cohort studies (III and IV) were conducted to assess the associations between single-leg hop performance, lower extremity muscle strength and postural orientation, respectively, and the KOOS, the TAS, the ARS (n=54) and the K-SES (n=89).

The results of this thesis show that the Swedish version of the ARS has good reliability and validity in the settings of and after knee injury rehabilitation. The results also indicate that the ARS, the K-SES, the KOOS, and the Physical Component Summary of the 36-Item Short-Form Health Survey have good responsiveness in young adults undergoing rehabilitation for a knee injury. Furthermore, the results of this thesis demonstrate that worse single-leg physical performance, in particular performance in the vertical hop tests and the side hop tests and postural orientation, correlated moderately with lower future self-reported activity participation and worse future knee function after ACL injury/ACLR. Also, worse symmetry between legs in the single-leg hop test and in knee flexion muscle strength correlated moderately with lower future knee self-efficacy after ACL injury /ACLR. Treatment strategy, in terms of exercise therapy only, or in combination with early or delayed ACLR, after ACL injury did not appear to have an impact on future knee self-efficacy.

Key words: Knee injury, patient outcome assessment, performance-based measures, validation studies

Supplementary bibliographical information

Language: English

ISSN 1652-8220 Lund University, Faculty of Medicine Doctoral
Dissertation Series 2018:35

Recipient's notes

Number of pages 94

Price

Security classification

Signature Date 2018-03-13

I, the undersigned, being the copyright owner of the abstract of the above-mentioned dissertation, hereby grant to all reference sources permission to publish and disseminate the abstract of the above-mentioned dissertation.

O knee, where art thou?

Patient-reported outcomes and physical performance in individuals with acute knee injury

Vala Flosadottir



Cover photo by Lára Flosadottir, my sister. Plaza de España, Sevilla, Spanien. Copyright Vala Flosadottir

Faculty of Medicine Department of Health Sciences

Lund University, Faculty of Medicine Doctoral Dissertation Series 2018:35 ISBN 978-91-7619-602-1 ISSN 1652-8220

Printed in Sweden by Media-Tryck, Lund University Lund 2018



Media-Tryck is an environmentally certified and ISO 14001 certified provider of printed material. Read more about our environmental work at www.mediatryck.lu.se



Content

Adstract	11
Populärvetenskaplig sammanfattning.	13
Definitions and abbreviations	15
Definitions	
Abbreviations	
List of studies	19
Introduction	21
Knee injury	21
Epidemiology	
Anatomy and biomechanics	
Treatment	22
Outcomes after knee injury	23
Patient-reported outcomes measures	
Measures of physical performance	
Patient-reported outcomes or physical performance?	
Measurement properties of outcome measures	27
Rationale of the thesis	30
Aims of the thesis	33
Overall aims	33
Specific aims	33
Methods	35
Study designs and cohorts	35
The Knee Injury data	
The KANON data	
Participants in studies I-IV	40
Outcome measures	41
Patient-reported outcome measures, PROMs (I-IV)	
Knee-specific PROMs (I-IV)	
Generic PROM (II)	
Comparative PROMs (I and II)	
Performance-based outcome measures (III and IV)	
Translation and cross-cultural adaptation of the ARS	47
Data analyses	48

Studies I and II	49
Study III	
Study IV	50
Ethics	51
Results	53
Study I	
Translation, cross-cultural adaptation and validation of the ARS	
Studies I and II	55
Responsiveness of knee-specific and generic PROMs	55
Studies III and IV	
Knee-specific PROs.	
Associations between physical performance and knee-specific PR	.Os57
Discussion	59
Evaluation of measurement properties	59
Cross-cultural adaptation and validation of the Activity Rating Sc	ale
for disorders of the knee (ARS)	
Responsiveness of knee-specific and generic PROMs	
Associations between physical performance and knee-specific PROs	
Associations at 3 years after ACL injury	
Associations over time	
Associations between treatment strategy and knee-related self-efficacy	
ACL injury	
Methodological considerations and limitations	67
Clinical implications	71
Conclusions	73
Acknowledgements	75
References	77
Appendices	
Studies I-IV	

Abstract

Knee injuries are severe and common injuries in young and physically active individuals. Knee injury treatment, non-surgical and surgical, is aimed at improving patient-reported outcomes (PROs), reducing symptoms and restoring physical function. Physical performance tests and patient-reported outcome measures (PROMs) are often used in the evaluation of outcome after knee injury treatment. For this purpose, it is critical that the PROMs that are used demonstrate adequate measurement properties. Furthermore, it is important to identify modifiable factors that may be associated with improved outcome after knee injury. The first aim of this thesis is to investigate and present the measurement properties, specifically the responsiveness, of knee-specific and generic PROMs, in individuals undergoing knee injury rehabilitation. The second aim of this thesis is to investigate the associations between lower extremity physical performance and self-reported activity participation, knee function and knee self-efficacy in individuals after anterior cruciate ligament (ACL) injury or ACL reconstruction (ACLR).

Two methodological studies (I and II) were performed to assess the cross-cultural validity of the Swedish version of the Activity Rating Scale (ARS) for disorders of the knee (n=100) and to assess the responsiveness of the Knee injury and Osteoarthritis Outcome Score (KOOS), the Tegner Activity Scale (TAS), the Activity Rating Scale (ARS) for disorders of the knee, and the Knee Self-Efficacy Scale (K-SES) (n=76), respectively. Two cohort studies (III and IV) were conducted to assess the associations between single-leg hop performance, lower extremity muscle strength and postural orientation, respectively, and the KOOS, the TAS, the ARS (n=54) and the K-SES (n=89).

The results of this thesis show that the Swedish version of the ARS has good reliability and validity in the settings of and after knee injury rehabilitation. The results also indicate that the ARS, the K-SES, the KOOS, and the Physical Component Summary of the 36-Item Short-Form Health Survey have good responsiveness in young adults undergoing rehabilitation for a knee injury.

Furthermore, the results of this thesis demonstrate that worse single-leg physical performance, in particular performance in the vertical hop tests and the side hop tests and postural orientation, correlated moderately with lower future self-reported activity participation and worse future knee function after ACL injury/ACLR. Also, worse

symmetry between legs in the single-leg hop test and in knee flexion muscle strength correlated moderately with lower future knee self-efficacy after ACL injury /ACLR. In addition, treatment strategy, in terms of exercise therapy only, or in combination with early or delayed ACLR, after ACL injury did not appear to have an impact on future knee self-efficacy.

Populärvetenskaplig sammanfattning

Knäskador är vanligt förekommande bland unga och fysiskt aktiva individer. En skada på ligament, menisker eller brosk i knäet kan leda till att leden blir instabil samt att rörelsemönstret och belastningen i själva leden förändras. Dessutom ökar risken för en tidig artrosutveckling i knäleden. Behandlingen av en knäskada går ut på att optimera självskattade variabler (t.ex. livskvalitet, upplevd knäfunktion och aktivitetsnivå), samt att återställa den fysiska funktionen och minimera knä-relaterade symptom. Därför utvärderas behandling efter knäskada ofta med hjälp av självskattningsinstrument och funktionella test, t.ex. hopp- och styrketest, som mäter den fysiska prestationsförmågan.

På senare tid har användning av självskattningsinstrument ökat och antalet instrument som används i bedömningen av behandling av knäskada har efter hand blivit större. Instrument som ofta används för självskattning i dessa sammanhang är Knee Injury and Osteoarthritis Outcome Score (KOOS) för upplevd knäfunktion, Activity Rating Scale for disorders of the knee (ARS) och Tegners aktivitetsskala (TAS) för fysisk aktivitetsnivå samt Knee Self-Efficacy Scale (K-SES) för upplevd tilltro till knäet. För självskattning av allmän fysisk och mental hälsa används ofta Medical Outcomes Study 36-Item Short-Form Health Survey (SF-36). Individer som råkar ut för en knäskada har oftast höga förväntningar på att de ska återhämta sig till fullo efter behandling samt att de ska kunna återgå till tidigare aktivitetsnivå eller idrott. Trots rehabilitering efter en knäskada, med eller utan kirurgisk behandling, kan dock inte alltid ett lyckat resultat uppnås.

Det är viktigt att de instrument som används för utvärdering av behandling är relevanta för de individer som drabbas av en knäskada och att dessa instrument uppvisar goda mätegenskaper, dvs. god validitet, tillförlitlighet samt känslighet för förändring inom det område de är avsedda för att mäta. Ökad kunskap om mätegenskaperna kan underlätta för fysioterapeuter, läkare och forskare att välja lämpliga instrument för utvärdering av behandling efter knäskada. God fysisk prestationsförmåga efter främre korsbandsskada har visats sig vara prediktiv för bättre upplevd knäfunktion, återgång till idrott och minskad risk för artros i knäleden. Syftet med studierna i den här avhandlingen var att undersöka och jämföra mätegenskaperna, särskilt känsligheten för förändring, hos ARS, KOOS, TAS, K-SES och SF-36. Dessutom, var syftet att undersöka vilka faktorer av fysisk prestationsförmåga som kan ha samband med bättre självskattad knäfunktion, aktivitetsnivå och tilltro till knäet efter främre korsbandsskada.

Vi genomförde två metodstudier, där vi i den ena översatte och utvärderade mätegenskaperna hos den svenska versionen av the Activity Rating Scale for the disorders of the knee (ARS). I den andra metodstudien undersökte vi känsligheten för förändring i fem självskattningsinstrument, som används för bedömning av upplevd knäfunktion, aktivitetsnivå, tilltro till knäet och allmän hälsa. I dessa studier ingick individer som var i rehabilitering för en knäskada. I två longitudinella studier undersökte vi betydelsen av fysisk prestationsförmåga för framtida upplevd knäfunktion, aktivitetsnivå och tilltro till knäet efter främre korsbandsskada. Dessutom undersökte vi betydelsen av olika behandlingsstrategier av främre korsbandsskada för framtida tilltro till knäet.

Resultaten i avhandlingen visar att den svenska versionen av ARS är valid och tillförlitlig, samt att den kan användas för att upptäcka förändring i aktivitetsnivå under och efter rehabilitering av en knäskada. Vidare visar resultaten att KOOS, K-SES och SF-36 är lämpliga instrument som kan användas för att upptäcka förändring i självskattad funktion i idrotts- och fritidsaktiviteter, knä-relaterad livskvalitet, tilltro till nuvarande knäfunktion respektive allmän fysisk hälsa hos individer som har genomgått rehabilitering för en knäskada.

Nedsatt fysisk prestationsförmåga, speciellt i vertikalhopp och sidohopp utförda på ett ben, samt förmågan att stabilisera de olika kroppsdelarna i relation till varandra och omgivningen, verkar ha ett tydligt samband med lägre aktivitetsnivå och självskattad knäfunktion efter en främre korsbandsskada. Dessutom verkar en större skillnad i resultat mellan skadat och oskadat ben i enbenslängdhopp samt fram- och baklårsstyrka ha ett klart samband med tilltro till knäet efter främre korsbandsskada. När det gäller främre korsbandsskada verkar behandlingsstrategin, dvs. rehabilitering bestående av strukturerad träning, eller rehabilitering i kombination med tidig eller senarelagd kirurgisk behandling, inte påverka den upplevda tilltron till knäet. Dessa resultat understryker betydelsen av målinriktade övningar i rehabiliteringen av en främre korsbandsskada för att förbättra den fysiska prestationsförmågan, framför allt i hoppoch styrketest.

Definitions and abbreviations

Definitions

Exercise therapy A regimen or plan of physical activities designed and

prescribed for specific therapeutic goals. Its purpose is to restore normal musculoskeletal function or to reduce

pain caused by diseases or injuries.

Measurement properties Features of a measurement instrument that reflect the

quality of the measurement instrument

Patient-reported outcome A measurement or a response that comes directly from

the patient without any interpretation by anyone else.

Rehabilitation A treatment by a physical therapist that includes exercise

therapy and other physical therapy modalities.

Abbreviations

ACL Anterior cruciate ligament

ACLR Anterior cruciate ligament reconstruction

ACL-D Treated with exercise therapy only

ACL-R Treated with exercise therapy and early reconstruction

ACL-X Treated with exercise therapy and optional delayed reconstruction

ADL Activities of daily living

ARS Activity Rating Scale for disorders of the knee

BMI Body Mass Index

COSMIN COnsensus-based Standards for the selection of health Measurement

INstruments

ES Effect size

GRC Global Rating of Change

ICC Intraclass Correlation Coefficient

IQR Inter Quartile Range

KOOS Knee injury and Osteoarthritis Outcome Score

- Pain Symptoms of pain, subscale in KOOS

- Symptoms Other symptoms than pain, subscale in KOOS

- ADL Activities of daily life, subscale in KOOS

- Sport/Rec Function in sport and recreation, subscale in KOOS

- QOL Knee-related quality of life, subscale in KOOS

KOOS₄ Mean score of KOOS subscales Pain, Symptoms, Sport/Rec and QoL

K-SES Knee Self-Efficacy Scale

- Present Perception present knee-related capability, subscore in K-SES

Future Perception about RTS and fear of reinjury, subscore in K-SES

LCL Lateral collateral ligament

LOA Limits of agreement

MCL Medial collateral ligament

OA Osteoarthritis

PCL Posterior cruciate ligament
PROs Patient-reported outcomes

PROMs Patient-reported Outcome Measures

RCT Randomized Controlled Trial

SD Standard deviation

SDC Smallest detectable change

SEM Standard error of measurement

SF-36 Medical Outcomes Study 36-Item Short-Form Health Survey

- PF Physical functioning, subscale in SF-36

- RP Role physical, subscale in SF-36

- BP Bodily pain, subscale in SF-36

- GH General health, subscale in SF-36

- VT Vitality, subscale in SF-36

- SF Social functioning, subscale in SF-36

- RE Role emotional, subscale in SF-36

- MH Mental health, subscale in SF-36

- PCS Physical component summary, subscore in SF-36

- MCS Mental component summary, subscore in SF-36

RTS Return to sports

SGPALS Modernized Saltin-Grimby Physical Activity Level Scale

SRM Standardized response mean

TAS Tegner Activity Score

VAS Pain Visual Analogue Scale for Pain

List of studies

This thesis is based on the following studies, which will be referred to by their Roman numerals in the text.

- I. Flosadottir V, Roos EM, Ageberg E. Cross-cultural translation and measurement properties of the Swedish version of the Activity Rating Scale (ARS) for disorders of the knee. Orthop J Sports Med. 2017 Sep 25;5(9):2325967117729361. doi: 10.1177/2325967117729361.
- II. Flosadottir V, Roos EM, Ageberg E. Responsiveness of knee-specific and generic patient-reported outcome measures in people undergoing rehabilitation for a knee injury. In manuscript.
- III. Flosadottir V, Roos EM, Ageberg E. Muscle function is associated with future patient-reported outcomes in young adults with ACL injury. BMJ Open Sport & Exercise Medicine, 2016 Oct 10;2(1):e000154
- IV. Flosadottir V, Frobell RB, Roos EM, Ageberg E. Impact of treatment strategy and physical performance on future knee-related self-efficacy in individuals with ACL injury. BMC Musculoskelet Disord. 2018 Feb 13;19(1):50. doi: 10.1186/s12891-018-1973-2.

Introduction

Knee injury

Epidemiology

Acute injury affecting the knee joint, including injuries to ligaments, menisci and/or cartilage, isolated or in combination, are most frequent among physically active young adults and middle-aged individuals (1). These injuries are a major risk factor for the development of early onset knee osteoarthritis (OA) (2-4). The injury mechanism often includes pivoting and twisting movements and sudden changes of directions, typically as seen in sports such as soccer, handball, floorball, down-hill skiing and basketball (5, 6). The anatomical location of the knee results in enormous weight bearing forces acting on the joint during exercise and physical activities. Therefore, it may not be surprising that knee is one of the most common injured joint (7), and that knee injuries account for up to 48% of all injuries in soccer, handball, basketball and floorboll (8).

Injury to the anterior cruciate ligament (ACL) is the most common knee ligament injury among athletes and the estimated incidence is around 80 injuries per 100 000 person-years (9, 10). In Sweden, this corresponds to approximately 6000 ACL injuries per year, and around 250 000 injuries per year in the United States (5). Injury to the ACL is commonly associated with injuries to the menisci, joint cartilage or other ligaments (11, 12). Gender wise, female athletes have 1.5 times increased risk of ACL compared to male athletes (13). Meniscal injury, another common knee injury, has an estimated incidence of 60-70 injuries per 100 000 person-years (6, 14). Articular cartilage lesions are relatively frequent in the knee, and based on studies of knee arthroscopies, the prevalence is estimated at around 60% in young to middle-aged individuals (6, 15). These injuries; ligament, meniscus and cartilage, often occur in combination (6, 16). At least 50% of all ACL injuries are associated with meniscus injuries (11) and around 70% of articular cartilage injuries are associated with either ACL or meniscus injuries (6). The incidence of dislocation of the patella, most commonly seen in adolescents, is estimated at 43 injuries per 100 000 person-years (17).

Anatomy and biomechanics

The different structures in the knee joint each have a specific purpose for joint function and health. The menisci act as load transmitters, shock absorbents and stabilizers in the knee joint (18-22). The joint cartilage has a fundamental role in varying load bearing, which reduces the pressure on subchondral bone, and in minimizing friction through range of motion and functional activities (16). The ACL plays an important role in the knee joint by providing stability in the anterior-posterior translational and internal-external rotational direction (23, 24). An injury to these structures causes mechanical joint instability (25), changed kinematics and joint loading (26, 27), reduced muscle strength (28, 29) and worse balance (30). These physical impairments increase the risk of secondary injuries (31), and in the long-term, may lead to early onset knee osteoarthritis (11, 14, 27, 32, 33) and increased risk of total knee replacement (34). Subsequently, all these consequences and unsuccessful outcomes after ACL injury can result in direct and indirect costs for the individual, as for the society (35).

The knee joint is dependent on both dynamic and passive stability, which is provided by muscles and ligaments, respectively (25, 36). After a ligament injury, patients often experience reduced control of the knee joint in weight bearing activities (i.e. giving way events), referred to as functional instability (37). The functional joint instability caused by a ligament injury may be replaced with dynamic stability through adequate rehabilitation, including structured exercise therapy (38-42). It is suggested that optimal knee and muscle function after knee injury is first regained at approximately 2 years after injury (43-47).

Treatment

The treatment after a knee injury is aimed at improving patient-reported outcomes (PROs), reducing symptoms and restoring physical function. Individualized exercise therapy, with or without the addition of a surgery, is considered the main choice of treatment for knee ligament, meniscal and cartilage injury, and, typically includes exercises aimed at increasing functional stability and lower extremity strength, restoring range of knee joint motion, sensorimotor function (balance, coordination and proprioception), and minimizing joint effusion and pain (14, 45, 48-52). Surgery may be considered in cases of frequent activity participation and significant knee instability (10, 53-56), or mechanical interference of joint movements (14) despite completed exercise therapy.

The treatment of an ACL injury, surgical or non-surgical, should include individualized exercise therapy and all progress should be based on clinical milestones rather than a specific timeline (57). It is still debated whether a surgical or a non-surgical treatment is the best approach in the long term (35, 58-60). A surgical reconstruction of the ACL

may offer the prospect of better anatomical reconstruction, improved mechanical stability during recovery and over time, as well as return to preinjury level sports participation (61-65). However, not all patients experience normal knee function or return to preinjury sports after ACL reconstruction (ACLR) (66, 67). Furthermore, the outcomes up to 5 years after ACL injury, in terms of PROs and physical function, do not appear to differ between surgical or non-surgical treatment (40, 61, 68-71). Therefore, a non-surgical approach may be advocated as the primary choice of treatment for ACL injury (40, 41, 71-73). In addition, recent findings suggest that it is favorable to start exercise therapy before considering ACL reconstruction, especially for young and physically active individuals with acute ACL injury and associated meniscus injury, and for individuals that report severe knee-related pain, symptoms and impaired knee function early after injury (74). Therefore, it could be speculated that the best approach for treating an ALC injury lies in optimizing exercise therapy treatment. However, if lower perceived knee function, joint instability, lower extremity strength deficits, and movement asymmetries still persists after structured exercise therapy treatment, and if the intent is to return to high-demanding work or physical activity including pivoting and cutting movements, an ACLR may be considered (56). This period of structured exercise therapy before ACLR has been shown to be associated with better outcomes, in terms of improved self-reported knee function and increased lower extremity muscle strength (75, 76).

There have been great advances in the treatment of ACL injuries during the last decades (77-79), and recently an evidence statement for the rehabilitation after ACLR was published (80). In addition, an international consensus statement for the prevention, diagnosis and treatment of paediatric ACL injury has been established (81). Yet, there are no evidence-based guidelines available for the optimal selection of non-surgical or surgical treatment strategies for patients with acute ACL injury. In a global perspective, there is an apparent difference in the management of ACL injuries, specifically in the trends and incidence of ACLR. This may be illustrated by the majority of the patients in the United States undergo surgical reconstruction after ACL injury (12, 63), compared to only one in every three patients in Sweden (10). Consequently, patients need uniform guidance based on high-quality evidence in their decision-making regarding treatment options following a knee injury.

Outcomes after knee injury

Assessment of treatment outcome is critical in the clinical practice. The information regarding the outcome during or after treatment can be used to make decisions about the choice or progress of treatment, and to evaluate the treatment effect. Evaluation of outcome after knee injury treatment includes generic and knee-specific PROMs and

tests of clinical measures, such as physical performance, joint range of motion and joint stability (50, 82-84). It is important to evaluate the effect and process of the treatment using measures that are meaningful and relevant to the patient (85, 86). For individuals with a knee injury, measures of knee function, symptoms, quality of life, physical activity level and return to sports and psychological factors are often used in treatment evaluation (68, 84, 85, 87-89).

Patient-reported outcomes measures

PROMs constitute the recommended main outcome in clinical trials (90) and according to international consensus, PROMs should be the primary outcome in the evaluation of outcome after knee injury (84). Generic PROMs, such as Medical Outcomes Study 36-Item Short-Form Health Survey (SF-36), allow for comparisons across different medical conditions, yet, important aspects may not be noticed for specific populations (91). Therefore, the development of PROMs should include condition-specific patient involvement and be validated in the population of interest (92). There are several knee-specific PROMs commonly used to evaluate outcome after knee injury treatment, including the Knee Injury and Osteoarthritis Outcome Score (KOOS), the Tegner Activity Scale (TAS), the Activity Rating Scale for disorders of the knee (ARS), and the Anterior Cruciate Ligament Return to Sport after Injury scale (ACL-RSI) (84, 85). These knee-specific PROMs measure the patient's perception on how the knee injury affects their daily life, body structure and function, quality of life, activity, participation as well as contextual factors.

In the short- and the long-term, knee injuries can result in serious concerns and consequences for the individual (11). Despite knee injury treatment, non-surgical or surgical, impairments often persist in terms of worse patient-reported outcomes (3, 17, 40, 41). Impairments in knee-specific PROs compared to pre-injury levels and population norms have been reported for quality of life (68), knee function (93), knee-related pain (3, 6), and physical activity participation (94). In individuals with ACL injury, generic PROs have also been shown to be impaired, in terms of worse general health (86) and increased depression (95) compared to normative values.

Pre-injury, present, and desired physical activity levels are often assessed both for clinical and research purposes in individuals with knee injury. In addition, return to physical activity participation at a pre-injury, or at a modified activity level, is commonly used to evaluate success in knee injury treatment (84, 85, 87, 96, 97). It has been shown that individuals with ACL injury have high expectations of good recovery of knee function and return to sports (RTS) (65, 98). However, these expectations may never be accomplished, as many patients do not return to their pre-injury level sports participation despite completed rehabilitation and good self-reported knee function. Around 82% return to some type of physical activity participation after ACL injury or

reconstruction, and only 39-55% return to competitive sports despite completed rehabilitation (67, 89). Physical activity participation in terms of intensity and type is often assessed with the Tegner Activity Scale (TAS) and the Activity Rating Scale for disorders of the knee (ARS) is used to assess the frequency of activity participation (85). The ARS has been recommended to evaluate progression and outcome after ACLR rehabilitation (80). Until now, a validated version of the ARS has not been published in Swedish.

It has been shown that sport injuries can be accompanied with negative psychological responses, such as sadness, fear of reinjury, and loss of confidence and identity (99) and that these responses can affect the outcome after injury (100). In recent years, there has been increased focus on psychological factors and their association to recovery and RTS after ACL injury (101-104). Knee-related self-efficacy, defined as the perception of one's capability to perform knee-related activities, is one psychological factor that has shown to be important for outcome after ACL injury (105). Knee-related self-efficacy has been reported to be low soon after ACL injury or ACLR but appears to increase during the process of rehabilitation (106, 107). However, factors, such as physical performance or treatment strategies (surgical or non-surgical), that may affect future knee-related self-efficacy after ACL injury need further investigation. The Knee Self-Efficacy Scale (K-SES) has been recommended to be included in the evaluation of psychological changes during and after ACLR rehabilitation (80).

Measures of physical performance

Knee function is reliant on numerous biomechanical factors and conditions (25). Therefore, it may be speculated that no single factor, in terms of physical performance, can be used to determine knee function after knee injury. Tests of single leg hop performance and lower extremity muscle strength are often used to assess physical outcome and knee function after knee injury (108). The advantage of these tests is that they may be used to determine progress during rehabilitation, discharge from rehabilitation and a save return to sports (109). Often, the performance of the non-injured leg is used as control for the performance of the injured leg, by calculating the Limb Symmetry Index (LSI = injured leg/non-injured x 100) (110). However, this method may raise some concerns as altered movement patterns and strength deficits have been reported not only for the injured leg but also for the non-injured leg after ACL injury (111-113). Therefore, deficits in performance may be underestimated when using the LSI. Despite this, LSI is frequently used and a 90% symmetry between legs (LSI \geq 90%) in several physical performance tests has been suggested as criteria for a successful outcome and return to sports (84, 114, 115).

Single-leg hop tests, assessing maximum hop performance or endurance, are commonly used and are designed to reflect the demands of a high level of physical activity (109,

116). The single leg hop for distance is the most commonly used hop performance test after ACL injury (110). In addition, tests of lower extremity muscle strength, assessing knee extension and flexion strength, are recommended in outcome evaluation after knee injury (31, 110, 117, 118). Assessment of postural orientation, defined as the ability to stabilize joints in relation to each other and to the environment (119), may also be used to complement strength and hop assessment after knee injury (120). Extensive test batteries, which include hop tests, strength test and assessment of postural orientation, can preferably be used to quantify knee-related performance in patients after ACL injury or ACLR (80).

Patient-reported outcomes or physical performance?

The use of PROMs and physical performance tests to evaluate recovery and progress during and after rehabilitation after ACL injury has shown that these measures also appear to be predictive of several outcomes after ACL injury. The associations between physical performance and knee-specific PROs in individuals with ACL injury have been investigated to a certain extent.

PROMs assessing psychological factors, in terms of fear of reinjury, reduced self-efficacy and reduced motivation, have been shown to be associated with lower rate of return to sports after ACL injury (88, 102, 121, 122). High pre-operative knee-related self-efficacy seems to be predictive of better single-leg hop performance at 1 year after ACLR (105). Furthermore, worse score in the SF-36 Bodily pain prior to ACLR seems to be a predictor for worse self-reported knee function at 2 years after ACLR (76). Low physical activity participation at 2 years after ACLR has been reported to be associated with an increased risk of painful knee at 6 years (123).

Physical performance tests can provide valuable information about self-reported knee function, knee-related quality of life, treatment strategies, reinjury and knee OA development after ACL injury. The LSI for the single leg hop test for distance at 2.5 months and the LSI for the triple-leg crossover and the 6-meter timed hop at 6 months, respectively, have been shown to be predictive of self-reported knee function at 1 year after ACL injury or ACLR (124, 125). Lower LSI in the single leg hop and the triple hop for distance at 1 year after ACLR was associated with lower rates of return to preinjury level sports at 2 years (103). Also, a LSI of < 90% in the single leg hop for distance at 1 year was found to be associated with knee OA development at 10 after ACLR (126). Worse performance in the one leg rise test after completed rehabilitation following ACL injury or ACLR has been reported to be associated with worse self-reported knee function at 2 and 5 years (70). A failure to achieve LSI \geq 90% in any single-leg hop or lower extremity muscle strength test, included in a battery of discharge tests, has been shown to be associated with higher risk of reinjury up to 2 years after ACLR (31, 127).

The moderate correlations previously reported between measures of physical performance and PROMs after ACL injury suggest that while these measures share some similarities, the information they gather is not the same (128-130) (70, 103, 124, 125, 129). Therefore, PROMs and measures of physical performance are obvious in complementing, rather than replacing each other.

Measurement properties of outcome measures

Over the past 20 years, there has been a considerable growth in the number of PROMs designed to evaluate outcome after knee injury. The process of choosing the appropriate PROM to accurately evaluate changes after treatment needs to include information about the construct to be measured, the target population and the measurement properties of the specific PROM (131). This is fundamental to enable correct investigation and comparison of different elements and options of treatment.

In recent years, the effectiveness of different treatment strategies for knee injuries, such as ACL and meniscus tears, have been investigated and compared in a number of randomized clinical trials (29, 40, 41, 132-134). PROMs are extremely important in the evaluation and interpretation of these trials. There is a need for a consistency in the PROMs that are used to enable comparisons between trials and to find the most appropriate PROM for each population. In this context, the measurement properties of the PROMs are critical.

The concepts of measurement properties, i.e. what they represent and how they should be assessed, have been defined in different and inconsistent manner (135, 136). These varieties of definitions and terms have led to different methods for assessing measurement properties of PROMs. The results of a study of measurement properties is dependent on the methods that are used. Therefore, it is important to use one definition in addition to an accompanying method in the assessment of measurement properties. In the Consensus-based Standards for the selection of health Measurement Instruments (COSMIN) study, an international consensus was reached on the taxonomy, the terminology and the measurement properties of health-related PROMs (Figure 1) (92). These consensus-based recommendations can be used as guidelines to evaluate and develop the measurement properties of PROMs.

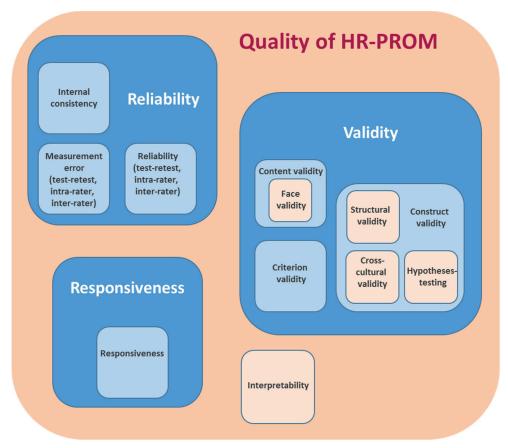


Figure 1.

The COSMIN taxonomy of relationships between measurement properties. COSMIN, COnsensus-based Standards for the selection of health Measurement INstruments; HR-PROM, health-related patient-reported outcome measure. Adapted from Mokkink et al 2010.

According to the COSMIN guidelines, there are three basic domains of measurement properties, including validity, reliability and responsiveness (Table 1). These domains are commonly assessed to provide evidence for the measurement properties of a PROM. Validation studies investigate whether the underlying construct of interest is properly assessed. Reliability studies probe into the stability of repeated measurements and the error or the "noise" of the measurements. Responsiveness studies investigate the longitudinal validity of the instrument and confirm that the instrument can detect changes over time in the construct of interest. (92) The three domains of measurement properties each include subcategories, which are further defined in Table 1.

Table 1. COSMIN definitions of domains, measurement properties, and aspects of measurement properties

Term			
Domain	Measurement property	Aspect of a measurement property	Definition
Reliability			The degree to wich the measurement is free from measurement error
	Internal consistency		The degree of the interrelatedness among the items
	Reliability		The proportion of the total variance in the measurements which is due to true differences between patients
	Measurement error		The systematic and random error of a patient's score that is not attributed to true changes in the construct to be measured
Validity			The degree to which a PROM measures the construct(s) it is intended to measure
	Content validity		The degree to which the content of a PROM is an adequate reflection of the construct to be measured
		Face validity	The degree to which (the items of) a PROM really looks as though they are an adequate reflection of the construct to be measured
	Construct validity		The degree to which the scores of a PROM are consistent with hypotheses based on the assumptions that the PROM validly measures the constructs to be measured
		Structural validity	The degree to which the scores of a PROM are an adequate reflection of the dimensionality of the construct to be measured
		Hypotheses testing	Same definition as construct validity
		Cross- cultural validity	The degree to which the the performance of the items on a translated or culturally adapted PROM are an adequate reflection of the performance of the items of the original version of the PROM
	Criterion validity		The degree to which the scores of a PROM are an adequate reflection of a "gold standard"
Responsiveness			The ability of a PROM to detect change over time in the construct to be measured
	Responsiveness		Same definition as responsiveness
Interpretability*			The degree to which one can assign clinical or commonly understood meaning to a PROM's quantitative scores or change in scores

PROM, patient-reported outcome measure. *Interpretability is not considered a measurement property, but an important characteristic of a PROM. Adapted from Mokkink et al 2010.

Cross-cultural validity

Cross-cultural validity is an aspect of validity, which includes the process of translation and re-evaluation of the measurement properties of a PROM in another language or cultural setting (92). The recommended method to linguistically and conceptually validate a PROM in a new language and culture consists of six steps (137, 138). These steps include a 3-stage translation, a review of an expert committee and a pre-testing of

the new version of the PROM (137). Lastly, the measurement properties of the PROM should be evaluated in an adequate sample ($n \ge 50$) of the target population (138).

Responsiveness

The definition of responsiveness is the ability of a PROM to detect change over time in the construct to be measured (92). In the evaluation of treatment outcome after an injury or disease, PROMs are often used to assess outcome over time. In that aspect, responsiveness is an important measurement property. According to the COSMIN guidelines, responsiveness should be assessed by testing hypotheses regarding the strength of correlations between change scores in a PROM and change scores in other PROMs that measure related constructs (139). Another approach for responsiveness assessment is the calculation of effect sizes. Cohen's effect size and the Standardized Response Mean (SRM) have been suggested to be appropriate measures to assess responsiveness (140). In addition, it has been suggested that a standardized effect size known as Cliff's delta should be calculated when data deviates from the normal distribution (141). These two approaches, hypotheses testing and different calculations of effect sizes, are suggested to evaluate the validity of change (longitudinal validity) and the magnitude of change, respectively (142).

The responsiveness of a PROM can vary depending on the specific populations, treatments and follow-up periods that are used (143). Hence, it is important to test and assess the responsiveness in the context the specific PROM is to be used. Furthermore, relevant PROMs should be assessed and compared alongside within the same study to facilitate the selection of appropriate PROMs in the evaluation of change in treatment outcome.

Rationale of the thesis

Knee injuries are severe injuries that are common in the young and physically active population. At present, there are numerous patient-reported outcome measures available to assess outcome after knee injury treatment. Highly physically active individuals, that have high demands on their knee function, need to be evaluated accordingly during and after knee injury treatment. The ARS, which assesses the frequency of knee-specific activity participation, may be an appropriate outcome measure for this purpose. Until now, a Swedish version of the ARS has not been validated.

It is recommended to establish the responsiveness of a PROM in the target population and in the relevant setting (92). The responsiveness of PROMs commonly used in the evaluation of knee injury treatment, including the KOOS, the TAS, the ARS, the K-SES and the SF-36, have previously, yet, separately been investigated in populations

with knee injury (85, 86, 144-147). The responsiveness, in terms of both magnitude and validity of change, of these knee-specific and generic PROMs has not yet been assessed and compared head-to-head within the same knee injury cohort. Increased knowledge about the responsiveness of these PROMs may facilitate the selection and the use of appropriate PROMs to evaluate change in outcome after knee injury.

The associations between physical performance and knee-specific PROMs in individuals with ACL injury have been investigated to a certain extent. Yet, the evidence is limited and there are differences in the methods used in previous studies, in terms of post-injury and post-surgical time intervals, the physical performance tests and the PROMs that are assessed (124, 125, 128, 129, 148, 149). Furthermore, the associations between physical performance after ACL injury/ACLR and future knee-related self-efficacy have not been investigated before. Understanding which components of physical performance are associated with improved patient-reported outcomes after ACL injury or ACLR may contribute to optimization of exercise therapy programs, improve adherence to rehabilitation and assist patients and clinicians in the decision and advisement of different treatment strategies.

High self-efficacy can facilitate one's initiative for action, level of effort and resilience to setbacks (150). After ACL injury, knee-related self-efficacy may be low but has been shown to improve during the course of rehabilitation (146). However, the levels of future knee-relate self-efficacy in patients treated with exercise therapy alone or in combination with either early or the option of delayed ACLR have not previously been reported. Therefore, the impact of surgical or non-surgical treatment strategies on knee-related self-efficacy is largely unknown.

Consequently, the studies in this thesis aim to investigate and present the measurement properties, specifically the responsiveness, of several knee-specific and generic PROMs, in individuals undergoing knee injury rehabilitation. In addition, the included studies are intended to complement and increase the knowledge about which physical performance factors and treatment strategies may be associated with improved PROMs after ACL injury or ACLR.

Aims of the thesis

Overall aims

The overall aims of this thesis were to 1) assess the measurement properties, in particular the responsiveness, of knee-specific and generic PROMs and to 2) investigate the association between lower extremity physical performance and knee-specific PROs in individuals with knee injury.

Specific aims

Study I: To 1) translate and cross-culturally adapt the ARS into Swedish and 2) assess measurement properties of the Swedish version of the ARS.

Study II: To evaluate and compare the responsiveness, in terms of magnitude and validity of change, of commonly used knee-specific and generic PROMs in patients undergoing rehabilitation for a knee injury.

Study III: To investigate the cross-sectional and longitudinal associations between lower extremity physical performance at 3 years and knee-specific PRO scores at 5 years after ACL injury/ACLR.

Study IV: To 1) report knee-related self-efficacy 6 years after acute ACL injury in patients treated with exercise therapy alone or in combination with either early or the option of delayed ACLR, and to 2) investigate associations between objectively measured single-leg physical performance at various time points after ACL injury/ACLR and self-reported knee self-efficacy at 6 years after injury.

Methods

Study designs and cohorts

The research for this thesis was performed using data from two separate study cohorts, the Knee Injury cohort and the Knee Anterior Cruciate Ligament, Nonsurgical versus Surgical Treatment (KANON) cohort (Figure 2). The Knee Injury data, including individuals in rehabilitation for a knee injury, was specifically collected for studies I and II in this thesis. The KANON data originates from the KANON study (40), a randomized controlled trial, which included 121 individuals with acute ACL injury. The measures used in the studies include quantitative variables from patient-reported questionnaires and physical performance tests.

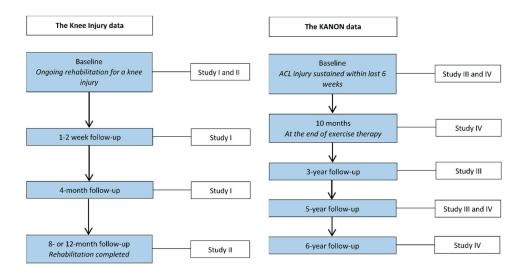


Figure 2. Settings and follow-ups in study I-IV

The Knee Injury data

Study I

The data for study I was collected between December 2014 and May 2016. The participants were recruited at seven physical therapy clinics in southern Sweden. The inclusion criteria for the participants were: ongoing rehabilitation for a knee injury (ligament, meniscus or cartilage) or post-injury knee osteoarthritis, surgically or non-surgically treated, and age 15-49 years. Participants were excluded if they had completed their knee injury rehabilitation between recruitment and baseline, were not limited in their activities due to their knee injury, had other disease or disorders overriding their knee injury, if they had an overuse knee injury (i.e. jumper's or runner's knee), or if they were physically inactive (Tegner Activity Scale score < 3). In May 2016, 182 individuals had completed the baseline assessment (Figure 3).

In study I, the measurement properties of the Swedish version of the ARS were assessed according to the COSMIN guidelines (138). Participants (n=100) that had completed the test-retest assessment were included in the reliability and the validity analyses. Participants (n=70) that had completed the assessment of the patient-reported questionnaires at the 4-month follow-up were included in the responsiveness analysis.

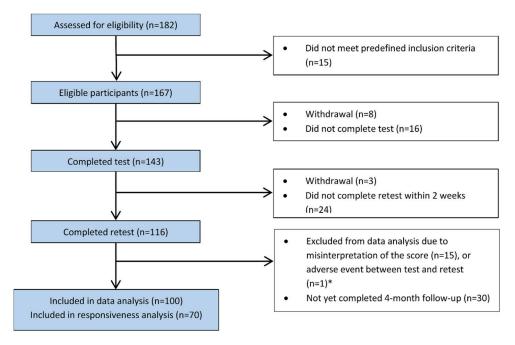


Figure 3.
Flow diagram study I. Recruitment and eligibility assessment performed until May 2016.

Study II

The data collection for study II continued through June 2017, using the same inclusion and exclusion criteria as were used in study I. At the end of the data collection, 225 individuals had completed the baseline assessment (Figure 4).

In study II, the responsiveness of the KOOS, the ARS, the TAS and the SF-36 was assessed according to the COSMIN guidelines (138). Participants (n=76) that had completed the patient-reported questionnaires at follow-up (8- or 12-months) and at that time reported that they had completed their knee injury rehabilitation were included in the responsiveness analysis.

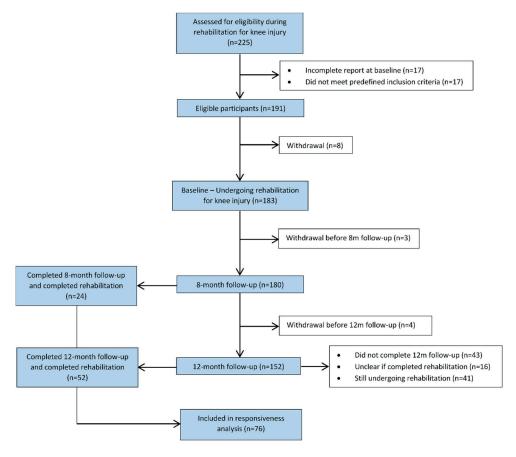


Figure 4.
Flow diagram study II. Recruitment and eligibility assessment performed until July 2017.

The KANON data

Study III

In study III, we used data from the original KANON (40) study, a treatment RCT, and data from an ancillary study of the RCT by Ageberg et al. (69). The inclusion criteria for the KANON study were: an acute ACL injury sustained within the last 4 weeks, a moderate to high physical activity level (Tegner Activity Scale score 5-9), and age 18-35 years. The main exclusion criteria were: professional athlete (TAS score 10), less than moderate physical activity level (TAS score < 5), a complete collateral ligament rupture, or full thickness cartilage lesion. All participants underwent a structured exercise therapy program, supervised by a physical therapist, for at least 4 months. The participants were randomized to a treatment consisting of exercise therapy only, or exercise therapy in addition to an early or an optional delayed ACL reconstruction. Inclusion criteria for the ancillary study (69) were: 2-5 years since injury.

In study III, the associations between physical performance and knee-specific PROMs scores were investigated. Participants (n=54) that had completed an extensive physical performance testing at mean 3 years after injury were included in the analyses (Figure 5). These participants also completed knee-specific PROMs, including the KOOS, the TAS and the ARS, at mean 3 and 5 years after ACL injury or ACLR.

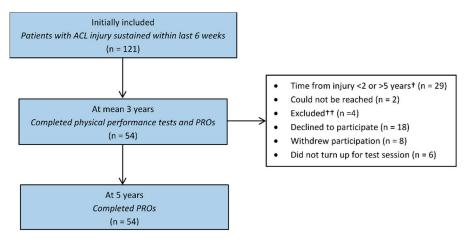


Figure 5. Flow diagram study III. † Inclusion criterion for physical performance tests, †† Pregnancy (n=1), still using cruthes (n=3).

Study IV

In study IV, we used data from the original KANON study (40) and data from an ancillary study of the KANON study by Ericsson et al. (70). In the ancillary study (70), 87 participants, of the original 121, underwent physical performance testing at the end of the exercise therapy (at mean 10 months).

In study IV, the K-SES scores for the groups-as-treated at 6 years were reported, and the associations between physical performance and K-SES scores were investigated. Participants (n=89) that had completed the physical performance tests at the end of the exercise therapy and at 5 years, and, in addition to having completed K-SES at 6 years were included in the analyses (Figure 6).

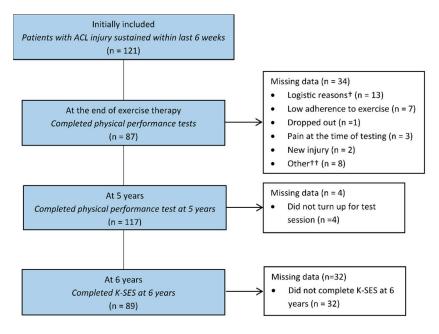


Figure 6.Flow diagram study IV. †Long-distance relocation or transferal to a physical therapist (PT) not involved in the study. ††Pregnancy (n=1), disc herniation (n=1), advised against performing test by PT (n=3), missing test protocols (n=3).

Participants in studies I-IV

Table 2. Settings and baseline characteristics of the participants included in studies I-IV

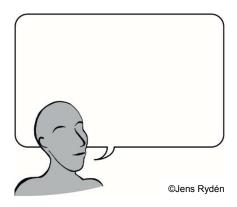
	Study I	Study II	Study III	Study IV
Data origin	Knee injury cohort	Knee injury cohort	KANON-trial	KANON-trial
Setting	Ongoing rehabilitation for a knee injury	Completed rehabilitation for a knee injury	Exercise therapy only or in addition to early or option of delayed ACLR after ACL injury	Exercise therapy only or in addition to early or option of delayed ACLR after ACL injury
Number of participants	100	76	54	89
Age, mean ± SD, y	27.0 ± 10.5	28.3 ± 8.8	29.7 ± 5.3	26.2 ± 5.1
Female, n (%)	55 (55)	40 (53)	15 (28)	25 (28)
BMI, mean ± SD, kg/m2	24.7 ± 3.8	24.3 ± 3.3	24.6 ± 3.0	23.8 ± 2.8*
Type of knee injury, n (%)				
ACL	74 (74)	54 (71)	54 (100)	89 (100)
Meniscal	12 (12)	11 (14)	NA	NA
Cartilage	4 (4)	2 (3)	NA	NA
MCL or LCL	4 (4)	5 (7)	NA	NA
Patellar dislocation	5 (5)	3 (4)	NA	NA
PCL	1 (1)	1 (1)	NA	NA
Surgical treatment, n (%)	58 (58)	48 (63)	36 (67)	69 (78)

^{*}n=87. NA, not applicable in these studies

Outcome measures

Patient-reported outcome measures, PROMs (I-IV)

The patient-reported outcome measures that are assessed in the studies in this thesis include four knee-specific and one generic PROMs. Four additional PROMs were exclusively used for comparative purposes in the validity assessment in studies I and II.



Knee-specific PROMs (I-IV)

In study I, the Swedish version of the Activity Rating Scale (ARS) for disorders of the knee was cross-culturally adapted and validated. In study II, the ARS, the Tegner Activity Scale (TAS), the Knee Self-Efficacy Scale (K-SES), and the Knee injury and Osteoarthritis Outcome Scores (KOOS) were assessed for responsiveness. In studies III and IV, the knee-specific PROMs were used to investigate cross-sectional and longitudinal associations with physical performance in individuals after ACL injury/ACLR.

Activity Rating Scale for disorders of the knee, ARS

The ARS (151) is a scale used to assess the frequency of participation in four separate activities with high demands on knee function; running, cutting, decelerating and pivoting. The frequency of activity participation is scored on a five-level scale from 0 (no participation) to 4 (participation four or more times a week), resulting in a total score between 0 and 16. The ARS has demonstrated good face, content and construct validity, and good test-retest reliability for the evaluation of physical activity participation in individuals with knee injury (151).

Tegner Activity Scale, TAS

The TAS (152) is a 11-level scale used to assess physical activity level based on recreational/sporting or occupational activities. The TAS ranges from 0 (sick leave or disability due to knee problems) to 10 (participation in competitive sports at national or international level). The TAS has demonstrated good construct validity and adequate test-retest reliability for evaluation of activity level in groups with knee injury (ACL and meniscal injury, patellar dislocation and knee OA) (144, 152, 153). Moderate to large effect sizes have been reported for the TAS at 9, 12 and 24 months after ACLR (153) and at 2 years after meniscal surgery (154).

Knee Self-Efficacy Scale, K-SES

The K-SES (145) holds four sections; A) daily activities, B) sports and leisure activities, C) knee function tasks, and D) future knee function. The certainty about the capability of performing an activity is rated on an 11-point Likert scale, with a range from 0 (not at all certain) to 10 (very certain). The full K-SES includes 22 items, which are summarized and divided by the number of items, producing a total K-SES score (A-D) ranging from 0 to 10. In sections A through C, the perception about one's present capability of performing knee-related tasks is reported, resulting in the subscore K-SES Present. In section D, the perception about one's future knee function capability is reported, i.e. perception about return to preinjury level sport and fear of reinjury (and future knee function if undergoing ACLR), resulting in the subscore K-SES Future. The subscores are calculated separately, with a range of 0 to 10. The K-SES has shown good internal consistency, test-retest reliability, face validity, content validity and construct validity in individuals after ACL injury or ACLR (145, 146).

Knee injury and Osteoarthritis Outcome Score, KOOS

The KOOS (155) is a 42-item score that includes five subscales with separate normalized subscale score with a range from 0 (worst) to 100 (best). The KOOS subscales cover knee-related pain, symptoms, activities of daily living (ADL), function in sports and recreational activities (Sport/Rec), and quality of life (QOL). In the subscale QOL, item 3 (Q3) covers perceived knee confidence. The KOOS has demonstrated good face validity, content validity, construct validity and good internal consistency and test-retest reliability in individuals undergoing surgery due to knee injury (ACL injury, meniscus or chondral injury) (144). Large effect sizes have been reported for the KOOS in individuals at 6 months after ACL (155) and moderate to large effect sizes have been reported at 3 months after meniscectomy (156, 157).

Generic PROM (II)

In study II, the magnitude of change was assessed for the SF-36 and compared with the magnitude of change for the knee-specific PROMs. The SF-36 was also used as a comparative instrument in the hypotheses testing for the responsiveness assessment of the knee-specific PROMs.

The Medical Outcomes Study 36-Item Short-Form Health Survey, SF-36

The SF-36 (158) holds 36 items scored within eight components of generic health: Physical Functioning (PF), Role Limitations due to Physical Health (Role-Physical, RP), Bodily Pain (BP), General Health Perceptions (GH), Vitality (VT), Social Functioning (SF), Role Limitations due to Emotional Problems (Role-Emotional, RE), and General Mental Health (MH). The score of the physical components (PF, RP, BP, GH) is summarized in the Physical Component Summary (PCS) and the mental components (VT, SF, RE, MH) are summarized in the Mental Component Summary (MCS). The eight components and the two component summaries each produce separate scores between 0 (worst health) and 100 (best health). The SF-36 has shown good criterion and construct validity and good internal consistency in a normative population (159, 160).

Comparative PROMs (I and II)

We used the following PROMs as comparative instruments in the hypotheses testing for the construct validity assessment of the Swedish version of the ARS and for the responsiveness assessment of the ARS, the KOOS, the TAS and the K-SES.

Modernized Saltin-Grimby Physical Activity Level Scale, SGPALS

The SGPALS (161) is a 4-level scale used to evaluate the level of general physical activity with a score ranging from 1 (physical inactivity) to 4 (regular, hard training for competitive sports). The SGPALS has demonstrated good construct validity and reliability in a general population (162, 163).

Visual Analogue Scale for Pain, VAS Pain

The VAS Pain (164) is an analogue scale used to assess pain intensity. In our study (II), the scale was marked on a horizontal line, generating a score from 0 (no pain) to 100 (worst imaginable pain). We asked the participants to grade their pain based on the question "How much knee pain did you experience during the past week?". The VAS Pain has demonstrated good construct validity and good test-retest reliability to be used in individuals with acute and chronic joint pain (165).

Pain manikin for measurement of musculoskeletal pain, Pain manikin

The Pain manikin (166) is a manikin displaying 44 body parts, 21 anterior and 23 posterior parts. The participants marked painful sites on the manikin to indicate whether they had pain from any muscles, joints or bones for at least one week during the last month. The pain should not be related to menstruation or influenza. In our study (II), the prevalence of pain was calculated for the number of marked sites. Pain assessed with the Pain manikin has demonstrated high agreement with pain assessed using written questions (166). Pain manikins have demonstrated good test-retest reliability in individuals with low back pain (167).

Global Rating of Change, GRC

We used GRC scales to assess the perceived change in the specific constructs of the PROMs (ARS, TAS, K-SES and KOOS) being evaluated in studies I and II. At follow up, the participants answered the eight following questions based on a 7-point Likert scale. 1) GRC reflecting changes in the ARS: "How often do you perform kneedemanding activities (running, cutting, decelerating to a quick stop, twisting/pivoting with your foot planted while playing sports) now compared to 8/12 months ago?" The response options: much more often, more often, somewhat more often, about the same, somewhat less often, less often or much less often. 2) GRC reflecting changes in the TAS: "How is your current activity level compared with 8/12 months ago?" The response options: much higher, higher, slightly higher, about the same, slightly lower, lower or much lower. 3) GRC reflecting changes in the K-SES: "How certain are you that you can perform physical activities, such as running after a bus, go on bike trip, jump on one leg or make quick turns, compared to 8/12 months ago?" The response options: much more certain, more certain, slightly more certain, about the same, slightly less certain, uncertain or very uncertain. GRC scales reflecting changes in the KOOS subscales Pain, Symptoms, ADL, Sport/rec, and QOL, respectively: 4) "How is your knee pain now compared to 8/12 months ago?" 5) "How are the other symptoms in your knee (swelling, stiffness, decreased range of motion) now compared to 8/12 months ago?" 6) "How is your ability to perform daily activities (sitting, standing, walking, stairs, dressing and household work) now compared to 8/12 months ago?" 7) "How is your ability to perform sport and recreational activities (running, jumping, squatting, kneeling, twisting on loaded knee) now compared to 8/12 months ago?" 8) "How is your quality of life that relates to your knee (trust in knee, life style, how often you think of your knee, etc.) now compared to 8/12 months ago?" The response options for questions 4-8 were: better, somewhat better, very small change (but not enough to be an important improvement), about the same, very small change (but not enough to be an important worsening), worse, or much worse.

Performance-based outcome measures (III and IV)

The Limb Symmetry Index (LSI) was calculated as the performance of the injured leg divided by that of the uninjured leg and multiplied by 100. For the analyses in study III, we used both absolute values (W, cm) and the LSI. In study IV, the LSI was used for all analyses of physical performance.

Muscle power tests

In study III, the muscle power tests were performed in weight training machines intended for knee extension, knee flexion and leg press, respectively, where the average power output (W) was measured using an electronic system (Muscle Laboratory; Ergotest Technology) (69). The knee extension power was recorded from 110° knee flexion to full extension, the knee flexion power was recorded from full knee extension to 110° knee flexion, and the leg press test was used to assess lower extremity power from 90° knee and hip flexion to fully extended knee and 30° hip flexion.

In study IV, the muscle power tests for hamstring and quadriceps strength were performed in an isokinetic device (BIODEX(168)) or in weight training machines intended for knee extension and knee flexion, respectively, using the principle of one maximal repetition (1 RM) (169).

Hop performance tests

In all of the four hop performance tests, both legs were assessed separately.

The single-leg hop for distance test (116) was performed from a standing starting position on one leg, with both hands placed on the lower back. From this position, the participants were instructed to jump as far as possible and land on the same foot (Figure 7). For a qualified jump, balance had to be remained at landing for 2-3 seconds and hands had to remain on the back during jump and landing. The distance (cm) was measured from the toes at the starting position to the heel at the landing position.

The vertical hop test (116) included an upright starting position on one leg and instructions to jump as high as possible (Figure 7). The participants were allowed to bend their knee as much as desired to initiate the jump. In study III both arms were held behind the back during the jump, whereas, in study IV the arms were free to help during the jump.

The side hop test (116) was performed standing on the test leg with both hands placed on the back. The participants jumped on one leg from side to side between two parallel lines on the floor, 40 cm apart (Figure 7). The instructions were to jump as many times as possible during 30 seconds, without touching the lines. Only the number of successful jumps was counted.

The square hop test (170) started with the participants standing outside a square (35×35 cm) marked with tape on the floor. The instructions were to jump in and out of the square in a clockwise rotation, as many times as possible, during 30 seconds. The number of landings inside the square, without touching the taped frame of the square, was recorded.

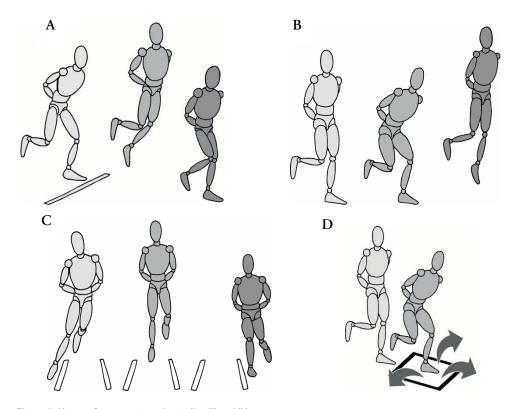


Figure 7. Hop performance tests in studies III and IV
A: Single-leg hop test for distance, B: Vertical hop test, C: Side hop, D: Square hop. ©Jens Rydén

Additional physical performance tests

In study III, the Test for Substitution Patterns (TSP) was used to visually assess postural orientation during the performance of five lower extremity functional tasks: 'Body weight-altering', 'Tip-toe standing knee flexion', 'Knee flexion-extension standing on one leg', 'Forward lunge from stairs' and 'Mini-squat' (120). The TSP contains a total score for all five tasks based on the orientation of several body segments. The total TSP scores has a range of 0 to 54 points where lower scores indicate better outcomes. The absolute values for the injured leg (TSP total score) and the difference between the absolute values of the injured and the uninjured leg (TSP diff) were used in the analyses of postural orientation.

In study IV, the single-leg rise and the single-leg balance test were used to assess hip-knee extensor strength and single-leg balance (70). In the single-leg rise test (170), the participants were asked to rise from a sitting to a standing position using the test leg with the other leg and both arms lifted in front of the body. The sitting height was adjustable and was lowered for each successful standing. The participants got three trials on each new height. The test continued until a failure to stand or until a height of 0 cm was recorded. The lowest height for a successful standing was recorded in centimeters. In the single-leg balance test (70) the participants were asked to stand as long as possible on the test leg, inside a marked square (35 x 35 cm), with the other leg fixed in maximal hip and knee flexion by both hands. Time (in seconds) was recorded from the closing of eyes until failure to stand on one leg (touching of the borders of the square with the test leg, touching the floor with the other leg or opening eyes).

Translation and cross-cultural adaptation of the ARS

In study I, the ARS was translated and cross-culturally adapted to Swedish in five steps according to recommended guidelines (Figure 8) (137, 138). This process included initial translation, synthesis of the translations and back translation by four translators, an extensive review by an expert committee and a pre-testing (n=12) of the preliminary Swedish version of the ARS. In the sixth and final step, the measurement properties of the Swedish version of the ARS were assessed in terms of reliability, validity and responsiveness.

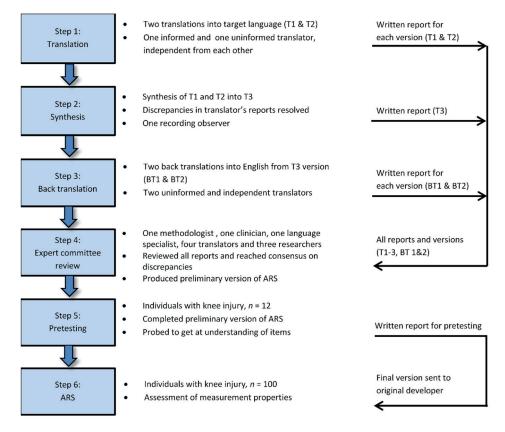


Figure 8.The process of translation and cross-cultural adaptation of the Acitivy Rating Scale in study I.

Data analyses

Statistical analyses were performed using IBM SPSS for Windows, V.22.0 (I and III) and V.23.0 (II and IV) (IBM Corp., Armonk, New York, USA). Descriptive statistical methods were used to describe baseline characteristics. The data in all studies was tested for normality by visual interpretation of histograms, Q-Q plots and the Kolmogorov-Smirnov test. The assumptions for normal distribution were not met for any of the PROMs, and therefore, non-parametric tests were used in the analyses. Depending on data type, the Chi-square test (categorical) or the Mann-Whitney *U* test (continuous), were used for between group comparisons. The Wilcoxon signed-rank test (continuous and ordinal) was used for within-group comparison. The one-way ANOVA test (categorical) and the Kruskal-Wallis test (continuous and ordinal) were used for comparisons between three groups. In all correlation analyses, correlation coefficients

thresholds suggested by Cohen (171) were used as follows; ≥ 0.10 to 0.29 denote low association, ≥ 0.30 to 0.49 moderate association and coefficients ≥ 0.50 large association. P values ≤ 0.05 were considered statistically significant. Further analyses were performed as follows.

Studies I and II

Floor and ceiling effects (I and II)

Floor and ceiling effects were considered present if more than 15% of the participants achieved the lowest or the highest score (143) in the ARS (I) or in any of the PROMs included in the analyses of longitudinal validity (II).

Face validity (I)

The face validity of the ARS was assessed by the expert committee during the cross-cultural adaptation process and through qualitative analysis of the pretest interviews.

Reliability (I)

Cronbach alpha was used to measure the scale and item internal consistency, were a Cronbach alpha of 0.70 to 0.95 was considered as good internal consistency (172). In the assessment of test-retest reliability; to prevent recall bias and a change in activity level among the participants, the questionnaires were completed on two different occasions 1-2 weeks a part (median, 7 days; interquartile range 7-9 days). The intraclass correlation coefficient (ICC), derived from a 2-way random –effects model (absolute agreement definition), was used to evaluate relative test-retest reliability. An ICC of \geq 0.70 was regarded as the minimum standard for reliability testing (143). Bland Altman plots with 95% limits of agreement (LOA), i.e. the mean difference \pm 1.96 * SD_{diff} (173), were used to determine the absolute reliability. The standard error of measurement (SEM = $SD_1 \times \sqrt{[1-r]}$, with SD_1 representing the standard deviation at baseline and r representing the ICC), was calculated including systematic differences to investigate to what extent the scores were the same for the repeated measurements. (92) The smallest detectable change (SDC) was determined both at an individual level ($SDC_{Ind} = 1.96 \times \sqrt{2} \times SEM$) and at a group level ($SDC_{Group} = SDC_{Ind}/\sqrt{n}$) (172).

Construct validity (I)

Construct validity was assessed by comparing the scores of the ARS with other knee-specific PROM scores by testing predefined hypotheses (Appendix B). Spearman's rank-order correlation analyses were used to test the associations between the scores. For the ARS to be accepted as a measure of good construct validity at least 75% of the predefined hypotheses had to be confirmed (172).

Responsiveness – Longitudinal validity (I and II)

The responsiveness of the ARS, the TAS, the KOOS, and the K-SES was assessed by comparing the change scores in these PROMs to the change scores in other construct-related PROMs and Global Rating of Change, by testing predefined hypotheses (Appendices B and C). To test the associations between the change scores Spearman's rank-order correlation analyses were used. To be accepted as a measure of good responsiveness at least 75% of the hypotheses had to be confirmed (172).

Responsiveness - Magnitude of change (II)

To evaluate the magnitude of change, the effect sizes (ESs) were calculated using Cohen's d, standardized response mean (SRM), and Cliff's d. Cohen's d (171) was calculated as the difference between the baseline mean scores and the post-rehabilitation mean scores divided by the SD at baseline. The SRM was defined as the mean change scores, divided by the SD of the change scores (140). Cliff's d (141) was used to compare all post-rehabilitation scores (A) to all baseline scores (B). When an A score was higher than a B score a value of +1 was noted, and when reversed, a value of -1 was noted. A tie yielded a value of zero. The sum of all scores was divided through the total number of counts ($n_A n_B$) resulting in Cliff's d (141). Cliff's d was classified as small (≥ 0.11), medium (≥ 0.28), and large (≥ 0.43) (174).

Study III

Spearman's rank-order correlation analysis was used to test the associations between the physical performance tests, where correlation coefficients above 0.8 between two physical performance tests resulted in the exclusion of one of the tests. The knee flexion (W), the leg press (W) and the one leg hop tests were excluded based on prior findings showing higher sensitivity for the knee extension and the side hop tests (69). The correlation coefficient between TSP difference and TSP total exceeded 0.90, thus, only the TSP total for the injured leg was included in further analyses. Spearman's and partial Spearman's rank-order correlation analyses were used to assess cross-sectional (3-year) and longitudinal (5-year) associations between physical performance and PROs, controlling for gender and treatment (surgical/non-surgical). Based on the exploratory character of the study, no adjustments were made for multiple comparisons.

Study IV

The data in this study were analyzed on a post-hoc as treated basis. Spearman's, and partial Spearman's, rank-order correlation analyses were used to assess associations between physical performance and K-SES, controlling for age, gender and treatment (exercise therapy only or with the addition of early or optional delayed ACLR).

Ethics

Studies I and II

The Research Ethics Committee at Lund University, Sweden, approved the studies (Dnr. 2014/672). The participants received information (verbal and written) about the data collection, its purpose, and its use. Since the data collection included sensitive personal data regarding health, we thoroughly explained that all data would be deidentified, i.e. the results could not be individually distinguished. Furthermore, all personal data was handled according to the Data Protection Act (personuppgiftslagen, PuL). All participants gave their written consent before entering the study. Data was collected through RedCap, a web based application for electronic data collection. The data was stored in RedCap, via the Medical Faculty of Lund University, according to current regulations. The ethical principles for medical research proposed in the Declaration of Helsinki were followed.

Studies III and IV

All participants received information about their participation and gave their written consent. The Research Ethics Committee at Lund University, Sweden, approved the studies (LU 535-01). The ethical principles for medical research proposed in the Declaration of Helsinki were followed.

Results

The general results of the studies are presented in this part of the thesis. The full details are available in the separate studies (I-IV).

Study I

Translation, cross-cultural adaptation and validation of the ARS

The translation and the cross-cultural adaptation of the ARS into Swedish included minor modifications of the original English version. We found the measurement properties of the Swedish version of the ARS (appendix A) to be adequate.

Steps 1-4: Translation and Cross-Cultural Adaptation

The English version of the ARS was translated into Swedish with minor adaptations. The expert committee questioned and discussed the wording of the second item and the recited sports in the fourth items. The word "cutting" was changed to "fast change of direction" in the Swedish version. In the cultural adaptation, three changes were made to the fourth item: "pivoting", where the wording "kicking, throwing, hitting a ball" was changed to "ball sports/games" and the sample sports were changed to "soccer, team handball, floorball, basketball, and racquet sports" were added. In addition, the a flexible recall period, i.e., 1 month or 1 week, was proposed as an option to 1 year in the English version to facilitate treatment follow-up. The review by the expert committee resulted in a preliminary Swedish version of the ARS.

Steps 5: Face Validity and Pretesting

The Swedish version of the ARS was considered a comprehensive and an appropriate outcome measure for knee-specific activity levels by the expert committee and the pretesting participants. No modifications of the Swedish version of the ARS were made after the pretesting.

Step 6: Assessment of Measurement Properties

Floor and Ceiling Effects

More than half of the participants (56%) reported the lowest score (0) in ARS_{Month} at baseline. We observed a floor effect of 22% and a ceiling effect of 20% for ARS_{Year}. Participants who had sustained their injury during sports activity reported significantly higher ARS_{Year} scores (mean, 8.6 ± 5.7), than those who had sustained an injury during other activity (mean, 2.8 ± 3.9) (P < .001). When we analyzed these groups separately, we observed no floor effect for ARS_{Year} (14.5%) in the group of participants with an injury sustained during sports activity, whereas, the floor effect (59%) remained for the group of participants with an injury sustained during other activity.

Reliability

The results from the assessment of test-retest reliability, internal consistency, SEM and SDC of the ARS are presented in table 5.

Table 5
Mean ARS scores at first and second assessment, test-retest reliability, internal constincency, SEM, and SDC of the ARS (n = 100)

	ARS Month	ARS Year
Mean score (SD) 1 st assessment	3.6 (5.3)	7.6 (5.9)
Mean score (SD) 2 nd assessment	3.6 (5.2)	8.1 (5.7)
Mean diff (95% CI)	-0.06 (0.65 to 0.53)	-0.42 (-1.08 to 0.24)
ICC (95% CI)	0.91 (0.87 to 0.94)	0.91 (0.87 to 0.94)
95% LOA	-5.86 to 5.74	-6.97 to 6.13
Cronbach's alpha	0.957	0.958
SEM	1.55	1.78
SDC ind	4.29	4.93
SDC group	0.43	0.49

Range of ARS scores = 0 (no participation) to 16 (very frequent participation). ARS, Activity Rating Scale; ICC, intraclass correlation coefficient; LOA, limits of agreement; SDCGroup, smallest detectable change at group level; SDCInd, smallest detectable change at individual level; SEM, standard error of measurement.

Construct Validity

Seven out of the eight (88%) predefined hypotheses were confirmed. The only rejection was the hypothesis regarding the correlation between higher ARS_{Month} scores and longer time since injury ($r_s = 0.134$). The correlations observed between ARS scores and TAS, SGPALS, and single KOOS item scores were at least moderate ($r_s \ge 0.310$). Higher ARS scores correlated with longer time since surgery and injury sustained during sports activity ($r_s \ge 0.342$).

Studies I and II

Responsiveness of knee-specific and generic PROMs

Longitudinal validity of the ARS in young adults undergoing knee injury rehabilitation (I) The ARS reached the criterion of 75% with six out of the seven (86%) predefined hypotheses being confirmed.

The single hypothesis that was rejected was the one regarding changes in ARS_{Month} scores and KOOS item P2 scores ($r_s = -0.027$). As hypothesized, the change in the ARS scores correlated higher with a change in the TAS scores ($r_s = 0.229$) than with the SGPALS scores ($r_s = 0.197$). The correlations between the change in the ARS scores and the change in single-item subscores of the KOOS Sport/Rec subscale and the perceived change in the ability to perform sports and recreational activities were $r_s \ge -0.270$. There was a low correlation between the change in the ARS scores and the change in the KOOS ADL subscores and the perceived change in function in ADL, respectively ($r_s \ge 0.011$).

Longitudinal validity of knee-specific PROMs in young adults after completed knee injury rehabilitation (II)

The ARS, the K-SES Present and all the subscales of the KOOS achieved the criterion of confirming at least 75% of the predefined hypotheses. For the TAS and the K-SES Future only one out of the seven (14%) predefined hypotheses could be confirmed, respectively.

The correlations between the specific GRCs and change scores in the ARS, K-SES Present and Future were at least moderate ($r_s \ge 0.332$). As hypothesized, the change scores in the KOOS Sport/Rec and QOL, and in the SF-36 Physical functioning and Role physical correlated at least moderately with the changes in ARS and K-SES Present scores ($r_s \ge 0.341$), respectively. The change scores in the ARS, the K-SES Present and the KOOS subscales correlated higher ($r_s \ge 0.173$), with the change scores in the SF-36 Physical component summary than with the change scores in the SF-36 Mental component summary. Change scores in TAS and the K-SES Present correlated higher with change scores in knee-specific activity than in generic activity, respectively ($r_s \ge$ 0.180). The change scores in all the subscales of the KOOS correlated at least moderately with the specific GRC ($r_s \ge 0.321$), except the change scores in the KOOS QOL ($r_s = 0.263$). As we hypothesized, the change scores in the VAS knee pain, the Pain manikin and the SF-36 subscores Bodily pain, Physical functioning and Role physical correlated at least moderately with the change scores of each of the KOOS subscales ($r_s \ge 0.315$). Only the change scores in the KOOS Sport/Rec and QOL correlated as hypothesized with the change in ARS scores ($r_s = 0.357$).

Magnitude of change in knee-specific and generic PROMs in young adults after completed knee injury rehabilitation (II)

The Cohen's *d* for the KOOS Sport/Rec, QOL, K-SES, K-SES Present, SF-36 Physical component summary and SF-36 Physical functioning, Role Physical and Bodily Pain ranged between 0.81 and 1.36, and were considered large. The Cohen's *d* for the ARS, KOOS Symptoms, Pain, ADL, SF-36 Mental component summary and SF-36 Vitality ranged from 0.50 to 0.79, and were considered moderate. The Cohen's *d* for the TAS and K-SES Future was 0.26 and 0.05, respectively, and were considered small.

The majority of the effect sizes calculated using the SRM and the Cliff's *d* were classified at levels corresponding to the classification levels when using Cohen's *d*. The SRM for the SF-36 Physical functioning and Role physical were 0.78 and 0.73, respectively, and were considered moderate. The SRM for the ARS was 0.43 and was considered small. For three of the PROMs, the classification increased one level when using Cliff's *d* compared to Cohen's *d*. For the KOOS Pain and ADL the Cliff's *d* were 0.49 and 0.48, respectively, and were considered large. The Cliff's *d* for the SF-36 Social Functioning was 0.31 and was considered moderate.

Studies III and IV

Knee-specific PROs

Changes in KOOS, TAS and ARS scores between 3 and 5 years (III)

We observed no statistically significant changes in the KOOS scores between 3 and 5 years (KOOS Pain -1 (95% CI -3 to 2), KOOS Sport/rec 2 (95% CI -3 to 7) and KOOS QoL -6 (95% CI -6 to 3)). Out of 54 participants, ten (19%) reported worse scores in the KOOS Q3, 15 (28%) in the TAS and 22 (41%) in the ARS at 5 years in comparison to 3 years.

K-SES scores at 6 years (IV)

At 6 years after injury, the median K-SES scores for all participants (n=89) were K-SES 7.8 (IQR 5.9-9.0), K-SES present 8.7 (IQR 6.8-9.6) and K-SES future 4.8 (IQR 2.5-7.5). We observed no differences between the three treatment groups in K-SES scores 6 years after ($p \ge 0.501$).

Associations between physical performance and knee-specific PROs

We found several associations between physical performance and PROs above 0.1 but we observed no correlation coefficients equal to or above 0.5. The correlation coefficients were quite similar when using Spearman's (r_s) and partial Spearman's rank-order correlation (r_{sp}) analyses. Therefore, only the results from the partial correlation analyses are presented.

Associations at 3 years (III)

At 3 years, worse performance in the vertical hop (cm, LSI) and the TSP total score were associated with worse scores on the KOOS Pain ($r_{sp} \ge 0.302$, $p \le 0.031$). Worse performance in the vertical hop (cm), the side hop (n, LSI) and the TSP total score, at 3 years, were associated with worse scores on KOOS Sport/rec ($r_{sp} \ge 0.320$, $p \le 0.021$). Worse knee extension power (LSI), worse performance in the vertical hop (cm, LSI), the side hop (LSI) and the TSP total score were associated with worse KOOS QoL ($r_{sp} \ge 0.314$, $p \le 0.023$), at 3 years.

Worse knee extension power (W), worse performance in the vertical hop (cm), the side hop (n, LSI) and the TSP total score were associated with lower TAS score ($r_{sp} \ge 0.330$, $p \le 0.017$) at 3 years. Worse performance in the vertical hop (cm, LSI) and the TSP total score were associated with lower ARS score ($r_{sp} \ge 0.302$, p = 0.031) at 3 years.

The correlations observed between the remaining physical performance tests and the PROs at 3 years ranged from $r_{sp} = 0.003$ (p = 0.984) to $r_{sp} = 0.272$ (p = 0.051).

Associations over time (III and IV)

Worse performance in the vertical hop (cm), at 3 years, was associated with worse scores on KOOS Pain ($r_{sp} = 0.308$, p = 0.026), at 5 years. Worse performance in the side hop (LSI), at 3 years, was associated with worse KOOS Sport/rec ($r_{sp} = 0.280$, p = 0.045) at 5 years. Worse performance in the vertical hop (LSI, cm), at 3 years, was associated with worse KOOS QoL ($r_{sp} = 0.281$, p = 0.044 and $r_{sp} = 0.284$, p = 0.041, respectively), at 5 years. Worse TSP total score, at 3 years, was associated with worse KOOS QoL ($r_{sp} = 0.334$, p = 0.017), at 5 years. Worse performance in the vertical hop (LSI) and worse TSP total score, at 3 years, were associated with worse KOOS Q3 scores ($r_{sp} = -0.324$, p = 0.019 and $r_{sp} = 0.372$, p = 0.007, respectively) at 5 years. Worse knee extension strength (W) at 3 years was associated with lower ARS score ($r_{sp} = 0.281$, p = 0.044) at 5 years. The correlations observed between the remaining physical performance tests at 3 years and the PROs at 5 years ranged between $r_{sp} = 0.008$ (p = 0.955) and $r_{sp} = 0.274$ (p = 0.052).

Worse knee flexion power LSI ($r_{sp} = 0.341$, p = 0.042) at the end of the exercise therapy, as well as worse single-leg hop LSI at 5 years ($r_{sp} = 0.310$, p = 0.005), correlated with worse scores on K-SES at 6 years after injury (Table 3). The associations observed

between the remaining physical performance tests at the end of the exercise therapy and the sores on the K-SES, ranged from r_{sp} = 0.148 (p = 0.264) for the single-leg hop LSI to r_{sp} = 0.265 (p = 0.045) for the square hop LSI.

Discussion

The first aim of this thesis was to assess the measurement properties, in particular the responsiveness, of knee-specific and generic PROMs, in young adults undergoing rehabilitation for a knee injury. For that reason, we conducted two methodological studies in accordance with the COSMIN guidelines. Our results showed that the Swedish version of the ARS has good reliability and validity in the settings of and after knee injury rehabilitation. Furthermore, based on the results we can recommend the use of the ARS, the K-SES, the KOOS, and the SF-36 Physical Component Summary to evaluate changes in activity participation, knee-related self-efficacy, knee-related pain, symptoms, function in daily living, function in sports and recreation and knee-related quality of life, and general physical health, respectively, in young adults undergoing rehabilitation for a knee injury.

The second aim of this thesis was to investigate the association between lower extremity physical performance and self-reported activity participation, knee function and knee self-efficacy in individuals after ACL injury/ACLR. The results demonstrated that worse single-leg physical performance correlated moderately with lower future self-reported activity participation, worse future knee function and lower future knee self-efficacy after ACL injury/ACLR. In addition, treatment strategy, in terms of exercise therapy only, or in combination with early or delayed ACLR, after ACL injury did not have an impact on future knee self-efficacy.

Evaluation of measurement properties

Cross-cultural adaptation and validation of the Activity Rating Scale for disorders of the knee (ARS)

The ARS was successfully translated and cross-culturally adapted into Swedish. Our results show that the Swedish version of the ARS demonstrates good reliability and validity to evaluate the frequency of knee-demanding activities during the past year, or the past month, in young to middle-aged adults with knee injury.

The floor and the ceiling effects that we observed supported that the ARS_{Month} may be better suited for use later in the treatment phase, i.e. when knee-demanding exercises have been introduced, and that the ARS_{Year} is appropriate for use in individuals that are physically active in sports.

The reliability of the Swedish version was considered good based on the high internal consistency and high test-retest reliability (172). Our results are in line with previous studies assessing the measurement properties of the original (151) and the Persian version (175) of the ARS. As many knee-specific PROMs (176-178), the ARS appears to be useful for group comparisons, supported by the low SDC_{group} values. In contrast, the high values observed for the SDC_{ind}, SEM, and LOA, indicate a need for quite a large change in ARS scores (> 6 points) to detect a real difference for an individual.

We concluded the construct validity of the Swedish version of the ARS to be good since 88% of the hypotheses were confirmed (172). As we hypothesized, frequent participation in knee-demanding activities correlated with higher physical activity level, less trouble during running and pivoting activities, longer time since surgery and injury sustained during sports activity. However, longer time since injury did not correlate with a more frequent participation in knee-demanding activities. A possible explanation for this may be activity modifications among the participants resulting in reduced participation in knee-demanding (i.e. pivoting) activities based on clinical advice or personal reasons (96, 179)

The ARS was originally developed as a baseline measure of activity participation (151). However, our results show that when using a recall period of one month, instead of one year, the ARS can be used to evaluate change in activity participation in young adults in rehabilitation for a knee injury. This may facilitate treatment follow-up of activity participation in individuals that are active in pivoting and cutting sports. Further assessment of the responsiveness of the ARS is discussed in the following section of this thesis.

Responsiveness of knee-specific and generic PROMs

In the assessment of the magnitude and validity of change, several instruments and subscales stood out. For the knee-specific PROMs, we observed large effect sizes for the KOOS subscales Sport/Rec, and QOL, the K-SES, K-SES subscore Present. As for the generic PROM, the SF-36 Physical Component Summary, and SF-36 subscales Physical Functioning, Role Physical and Bodily Pain demonstrated large effect sizes. Our results also demonstrated good longitudinal validity for the ARS, the K-SES subscore Present and all the subscales of the KOOS, with more than 75% confirmation of the predefined hypotheses (172). We observed small effect sizes for the TAS and the K-SES Future, and, in addition, only 14% of the predefined hypotheses could be confirmed for these instruments. Taken together, our results of the evaluation of

responsiveness of commonly used knee-specific PROMs indicate that the ARS, the K-SES and the KOOS are valid instruments to assess change in activity participation, knee self-efficacy and knee-related pain, symptoms, function in daily living, function in sports and recreation and knee-related quality of life, respectively, in individuals undergoing rehabilitation for a knee injury.

TAS and ARS – Change in knee-specific activity participation

In a recent systematic review (180), it was concluded that among the four most common activity scales (including the ARS) used for knee-specific activity participation, the TAS was the only scale showing adequate reliability, validity and responsiveness in an ACL injured population. However, the authors (180) called for a comparative analysis of the measurement properties of these four scales. In this thesis, we compared two of these activity scales. In our cohort, we observed low magnitude and validity of change for the TAS, and, therefore, we did not consider it responsive to rehabilitation after knee injury, surgically or non-surgically treated. However, in previous studies, effect sizes between 0.6-1.1 have been reported after ACLR (153) and meniscus surgery (154). The low effect sizes we observed for the TAS may be explained by the difference in starting points between the studies. Our cohort reported TAS scores when they were in rehabilitation for a knee injury, whereas in previous studies (153, 154) the TAS scores were reported before the participants received any intervention. It is possible that our participants reported higher TAS scores at the starting point, in comparison to previous studies (153, 154), which consequently may have resulted in lower change scores, and, thereby, lower effect sizes. Another potential explanation is that changes in the intensity of activity participation may have occurred within each activity level of the TAS, and simply were not detected. This may be supported by the moderate magnitude and good longitudinal validity that we noted for the ARS. The TAS is based on the intensity and type of activity participation, whereas the ARS is based on the frequency of participation. Our results suggest that the construct measured by the ARS better reflected the changes in activity participation than the TAS. In addition, our results indicate that the ARS is a responsive measure of changes in kneespecific activity participation in young adults during and after completed rehabilitation for a knee injury.

K-SES – Change in knee-related self-efficacy

There was a marked difference between the responsiveness demonstrated by the K-SES Present and the K-SES Future. These subscores measure separate constructs, the perception of present functional knee capability (K-SES Present) versus the fear of reinjury and uncertainty of future knee function (K-SES Future). The large effect sizes we noted are in line with previous findings that have shown significant changes in K-SES Present scores, and no changes in K-SES scores, during rehabilitation at 4-, 6-, and 12-month follow-up after ACL injury (106). The confirmation of all the seven

predefined hypotheses, where associations between change scores in K-SES Present showed high correlations with changes in perceived knee-related quality of life, knee-specific and generic physical function, support the good longitudinal validity of K-SES Present. However, for the K-SES Future the observed effect sizes were close to none, and with only one out of the seven predefined hypotheses being confirmed, the standards for longitudinal validity were not fulfilled. Based on our results, in addition to previous results (106), this may suggest that the perception of present functional knee capability is more modifiable than the perception of one's ability to return to sport and future knee function as well as fear of reinjury. On the other hand, this may indicate that rehabilitation after knee injury require additional interventions aimed at reducing fear of reinjury and increasing the certainty of future knee function. We found the K-SES subscore Present may be used to evaluate change in knee-related self-efficacy in individuals undergoing rehabilitation for a knee injury.

KOOS – Change in knee-related pain, symptoms, function in daily living, function in sports and recreation and knee-related quality of life

All the KOOS subscales met the requirements for good longitudinal validity, through change scores that correlated as hypothesized at a moderate to large level with changes in related PROs, in terms of pain, knee-specific activity participation, and specifically with general physical functioning. We noted the highest effect sizes for the KOOS subscales Sport/Rec and QOL, which is in line with results presented in a recent systematic review and a meta-analysis evaluating the measurement properties of the KOOS (176). Our results, based on individuals with a range of acute knee injuries (ligament, meniscal and cartilage injuries), surgically or non-surgically treated, add to these previous results (176), primarily based on individuals undergoing ACLR. The KOOS subscale ADL was the only subscale that demonstrated ceiling effects at baseline (in rehabilitation), which increased even more after completed rehabilitation. Despite this, we observed moderate to large effect sizes and a significant change in scores for the KOOS ADL, as for the KOOS Pain and Symptoms. This indicates that all the KOOS subscales, yet, specifically the KOOS subscales Sport/Rec and QOL, can be used to evaluate change in self-reported knee-related pain, symptoms, function in daily living, function in sports and recreation and knee-related quality of life, respectively, after rehabilitation for a knee injury.

SF-36 – Change in general physical and mental health

Our results showing higher effect sizes for the SF-36 physical components compared to mental components are consistent with previous findings (86) in in individuals undergoing meniscectomy or ACLR. Since the goals of knee injury rehabilitation and treatment include improved knee function and reduced knee symptoms it is reasonable that the treatment effect is higher for components that measure physical health in comparison to mental health.

In the evaluation of outcome after treatment, it is recommended to use at least one generic health-related PROM, in addition to disease-specific PROMs (181). The SF-36 is the most widely used health-related PROM (158, 182). On the basis of our results, we can endorse the use of the SF-36, in particular the SF-36 Physical Component Summary, as a generic PROM to complement the knee-specific PROMs in the assessment of change in general physical health after knee injury rehabilitation.

Associations between physical performance and kneespecific PROs

We observed several moderate correlations ($r_{sp} \approx 0.3$ to 0.5) in the cross-sectional and the longitudinal analyses between physical performance and knee-specific PROs. In the longitudinal analyses, worse single-leg hop performance was associated with worse future knee-related function and self-efficacy, worse postural orientation was associated with worse future knee-related function, worse knee extension muscle power was associated with lower future knee-specific activity participation, and worse knee flexion muscle power was associated with worse future knee-specific activity participation, and worse knee flexion muscle power was associated with worse future knee-self-efficacy.

Associations at 3 years after ACL injury

In the cross-sectional analyses at 3 years after ACL injury/ACLR, four of the physical performance tests showed moderate associations with knee-related pain, symptoms, quality of life, knee confidence and activity participation (study III). These levels of correlations are in line with previous results (129, 183), demonstrating at most moderate correlations between physical performance and patient-reported measures at 2-3 years after ACLR. Among the performance tests included in our analyses (study III), we found that absolute performance in the vertical hop test and postural orientation showed the strongest correlation with KOOS Pain, Sport/Rec, QOL, TAS and ARS scores at 3 years. Our results support previous findings showing that measures that evaluate performance versus perception, indeed measure similar, yet, different constructs, and therefore, complement one another as outcome measures after ACL injury/ACLR (82, 183, 184).

Associations over time

In studies III and IV, we observed that physical performance assessed at the end of rehabilitation period, at 3 years (time point of optimal muscle function) and at 5 years

correlated moderately with future knee-related pain and quality of life, knee-related activity participation and self-efficacy. In particular, we found that performance in the vertical hop and the single-leg hop tests was associated with future knee-related pain, knee confidence and knee self-efficacy. Furthermore, knee extension and flexion strength demonstrated moderate correlations with future knee-related activity participation and self-efficacy, respectively. Finally, postural orientation was moderately associated with future knee-related function in sports and recreational activities, quality of life and knee confidence.

To our knowledge, this is the first prospective study to investigate the associations between physical performance and the ARS in individuals with ACL injury. In a previous study (148), where the TAS was used as a measure of activity participation, physical performance (knee flexion and extension strength and hop performance) at 6 months after ACLR was shown to correlate with the TAS at 4 years. We observed only one significant association in our analyses (study III) between worse knee extension muscle strength at 3 years and lower future knee-related activity participation, measured with the ARS, at 5 years. Furthermore, no physical performance tests correlated with future physical activity level measured with the TAS. The main difference between our study and the study by Sousa et al (148) was the time point for the assessment of physical performance and the difference in sample sizes, 54 versus 223 participants (148), which may explain the difference in results. Yet, we expected physical performance to be more associated with future activity participation since the ability to generate high forces during high movement velocities is considered an important factor in physical and athletic performance (185). Furthermore, recent results have shown that worse hop performance, at 1 year after ACLR, was associated with lower rates of return to pre-injury sport level at 2 years (103). We noticed a large difference in the numbers of associations in the cross-sectional (at 3 yrs) compared to the longitudinal analyses (from 3 to 5 yrs). In our cohort, near half of the participants modified their activity participation from 3 to 5 years. It is possible that the participants had a change of perspectives due to adaptation or acceptance (186) or that contextual factors, such as social or life-style changes may have led to activity modifications (88, 102, 187). Yet, our results showing that physical performance was more associated with the frequency of activity participation (ARS) than with the level of activity participation suggest that individuals that are active in knee-demanding activities (cutting and pivoting sports) could benefit from optimized physical performance for present and future frequency of activity participation.

We observed that worse vertical hop performance (cm, LSI) and worse postural orientation (TSP total score) at 3 years was associated with worse scores on the subscales KOOS subscales Sport/rec, QOL and item Q3 at 5 years after ACL injury or ACLR (study III). In line with our findings, longitudinal associations have previously been reported (124, 125) between single-leg hop performance (single-leg, triple and crossover hop), tested within 6 months after ACL injury or ACLR, and worse future knee

function (International Knee Documentation Committee (IKDC) Subjective Knee Evaluation Form) at 1 year. In addition, at 10 months after ACL injury or ACLR, performance in the single-leg rise has been shown to correlate with self-reported knee function (KOOS₄) at 2 to 5 years (70). However, in one study (188), with a considerably longer follow-up, no associations were reported between performance in single-leg hop or muscle strength tests at 2 years and KOOS scores at 9.5 years after ACLR. It is reasonable that other factors may have affected the results (188), given the long time between assessment of physical performance and PROs.

Several previous studies have focused on the influence of psychological responses on physical function and performance (121). To our knowledge, in study IV, we are the first to investigate the impact of physical performance on knee-related self-efficacy after ACL injury or ACLR. Thomee et al (105) showed that higher knee-related self-efficacy, measured with K-SES prior to ACLR, was associated with higher (better) LSI for single-leg hop performance (single-leg hop, side hop, counter movement jump) at 1-year follow up after ACLR. Our results demonstrated that higher LSI for knee flexion strength at the end of exercise therapy, and higher LSI in single-leg hop performance at 5 years, correlated moderately with knee-related self-efficacy at 6 years after ACL injury or ACLR. This suggests that symmetry between legs in single-leg physical performance is one factor that contributes to good future knee-related self-efficacy.

Although the associations between physical performance and knee-specific PROs, presented in this thesis, are only moderate, our results are supported by findings from previous longitudinal studies (70, 103, 124, 125). Altogether, despite differences in physical performance tests, PROMs and time points for baseline and follow-up assessments, the results indicate that single-leg physical performance after exercise therapy and up to 3 years after ACL injury or ACLR, is one factor that contributes to good future self-reported outcomes, in terms of physical activity participation, self-reported knee function and knee-related self-efficacy (103, 124, 125). This stresses the possibility that targeted training to optimize physical performance after ACL injury or ACLR may improve future self-reported outcomes. Still, further studies are required to investigate the relative contribution of single-leg physical performance, in addition to other factors, which may have an impact on self-reported outcomes after ACL injury or ACLR.

Associations between treatment strategy and knee-related self-efficacy after ACL injury

We found that knee-related self-efficacy at 6 years after ACL injury was not associated with treatment strategy (ACL-D, ACL-R and ACL-X), in terms of exercise therapy alone or in combination with early or optional delayed ACLR. These findings are in line with previous observations in the KANON-trial, where there was no difference across treatment groups in various outcomes, i.e. self-reported outcomes (40) and osteoarthritis (40, 41) at 2 and 5 years, and muscle function (69) assessed at 3 years after ACL injury. One possible explanation for our findings of similar K-SES scores for the as treated groups could be the extensive exercise therapy program that all participants followed regardless of treatment strategy. The progression of the exercise program was based on the achievement of different goals for muscle function, range of motion and functional performance, in contrast to a pre-set number of exercise sessions (40). Participants that received exercise therapy only progressed faster and required fewer exercise sessions than the participants who had early or delayed ACLR in addition to exercise therapy. This approach, of extensive exercise therapy, was chosen to make sure that all participants, regardless of treatment strategy, would achieve a similar and good level of physical function (40). As we presented in study IV, there were no differences in physical performance between the as treated groups. Consequently, the exercise program may be the common factor explaining the similarity in K-SES scores across all treatment groups, rather than the early or the optional delayed reconstruction.

In our cohort (study IV), at 6 years after ACL injury or ACLR, the K-SES scores (median 7.4-8.2) were similar or better than K-SES previously observed in other Swedish cohorts (145, 146, 189). Thomee et al. reported K-SES scores of mean 6.8 and 7.6 at 12 months after ACL injury and ACLR, respectively (146), mean 7.3 at 12 months after ACL injury or ACLR (189), and median 6.7 at 1-12 months after ACL injury and 3-12 months after ACLR (145). There are no normative or cut-off values available to grade the K-SES scores as high or low. Yet, previously, similar levels of K-SES scores (mean 6.9-8.3) have been shown to correlate with greater likelihood of satisfaction with outcome, in terms of self-reported knee function, at 3 years after ALCR (97). Furthermore, considering the subscores in our cohort, the K-SES Present scores (perception about one's present capability of performing knee-related tasks) can be regarded high in comparison to the K-SES Future scores (perception about one's return to preinjury level sports and fear of reinjury plus decreasing knee function if undergoing ACLR), for all three treatment groups. In previous studies (106, 145, 146, 189), the subscore K-SES Future has been reported to be comparable or higher than the K-SES Present scores. The main difference between our study (IV) and the previous studies (106, 145, 146, 189) is the time point for the assessment of knee-related selfefficacy. Our participants reported K-SES at 6 years after ACL injury or ACLR,

whereas, in previous studies K-SES scores were reported within one month after ACL injury or prior to ACLR and only up to 1 year after injury or ACLR (106, 145, 146, 189). Consequently, time as a factor may be the explanation for the difference in K-SES Present and Future scores in our cohort. As previously reported in the KANONtrial (41) at 5 years after ACL injury or ACLR, only about 20% of the participants were active at their preinjury activity level or higher. This suggests similar levels of return to preinjury activity levels for our participants (approx. 20 %) at 6 years. This is in line with previous reports showing that at 2-7 years after ACLR, no more than 50% return to sports, at preinjury or competitive level (94). The K-SES Future scores are based on the perception about return to preinjury level sports, fear of reinjury and decreasing knee function. It is reasonable to assume that over time the expectations and motivation for return to preinjury level sports and the perception of acceptable knee function may change. Also, results from previous studies have shown that individuals report high, and possibly overoptimistic, expectations before ACLR or meniscus surgery (65, 98, 190). As for our participants, lower, and perhaps more realistic, expectations may be reported some years later. Therefore, the low K-SES scores among our participants may be a reflection of natural changes that occur over time, such as increasing age or changes in social commitments. An additional explanation for the discrepancy between the K-SES Present (perception of good knee function capability) and the K-SES Future (uncertainty about return to preinjury level sports) scores may be that despite perceptions of good knee function return to sports is hindered due to fear of reinjury (191, 192). Furthermore, in spite of athletes returning to their pre-injury level of sports after ACLR, there are indications that these individuals end their sports participation earlier compared to individuals without ACL injury (4, 193). More longitudinal research is needed to further investigate these potential explanations for change in future knee self-efficacy in individuals after ACL injury.

Methodological considerations and limitations

There are some limitations to this thesis. The heterogeneity of the cohort in study I and II, in terms of age, gender and time spent in rehabilitation, may constitute a limitation. This may have influenced the results of the assessment of the measurement properties of the ARS. The floor effects that we observed may have been present due to participants that had spent a shorter time in rehabilitation, those at an early rehabilitation phase without engagement in knee-demanding activities. Measurement error and SDC are dependent on the standard deviation at baseline, where a higher standard deviation results in a higher SEM and an increased SDC. Therefore, a more homogenous sample could have resulted in lower SEM and SDC for the ARS. The majority of the participants (74% and 71%) in the studies had ACL injury. Based on this, it may be argued that we should have only included participants with ACL injury

in the analyses, since there may be differences associated with the various injuries that may have influenced the results. However, we performed subgroup analyses, were we only included ACL injured participants, which showed similar results for the main analyses in studies I and II. Still, we considered the knee injury rehabilitation to be the main treatment factor that would influence change in PROs when rehabilitation was completed. Therefore, the heterogeneity among the participant, including the variety of surgically and non-surgically treated knee injuries, may make the results of these studies more generalizable.

In the assessment of responsiveness in studies I and II, we chose to use both the traditional method by calculating effect sizes and standardized response mean, in addition to using the hypotheses testing according to the COSMIN guidelines. In line with the COSMIN guidelines, we consider effect sizes a measure of the magnitude of change and the hypotheses testing a measure of validity of change. It may be argued that the use of effect sizes facilitates cumulative research, as effects can be examined across studies. In additions, effect sizes can be used to calculate sample size for longitudinal studies. The COSMIN approach may be considered a qualitative approach in comparison to the quantitative calculations of ESs and SRM. It may be speculated that the COSMIN approach leaves more room for interpretation, and that it is dependent on the comparative PROMs that are chosen as external criterions. In addition, the content and the number of predefined hypotheses may be considered arbitrary, which may influence the results. However, in study II, we chose to combine these two methods of responsiveness assessment aiming for more complete analyses and valid results. Furthermore, the sample sizes were good (172), the response rates (≥64%) were higher than the minimum recommended (194) and the results can be generalized at least to patients with ACL injury.

The comparative PROMs that were used as external criterions in the hypotheses testing had not all been validated according to the COSMIN principles, which may question their use as comparative PROMs. Our results may have been different if we would have used other PROMs for comparison in the hypotheses testing. However, the majority of the PROMs that are included in our studies have been validated using other methods and are widely used for evaluation of treatment after knee injuries (85).

It may be considered a limitation that we did not calculate the minimal important change (MIC) for the PROMs included in study II. Recently, it has been suggested that no single value of MIC can be assigned to a PROM across different settings (195, 196). MIC is commonly used in RCTs to assess clinically relevant differences between groups; yet, the MIC for a PROM can vary greatly depending on the definitions and the methods that are used. Therefore, despite the great importance of clinical relevance, it may not be established through MIC.

A limitation to studies III and IV is the limited and conflicting evidence of the measurement properties of physical performance tests, despite their frequent use (197).

In addition, the naming of the tests and the methods used to conduct the tests differ across studies (198). Nonetheless, many physical performance tests are easy to conduct, require no equipment and are based on functional movements. In addition, physical performance tests often provide instant feedback and can be used to increase motivation and compliance during exercise therapy. The single-leg hop test has shown moderate responsiveness and can be used to assess progress during knee injury rehabilitation (198). This performance test has demonstrated moderate responsiveness and can be used to assess progress during knee injury rehabilitation (198). The single-leg hop test appears to be able to differentiate between injured and non-injured leg in individuals with ACL injury, in addition to be able to detect differences in the ACL-injured knee and the non-injured knee in age-matched controls (198).

The use of correlation analyses in study III and IV did not allow us to determine causality between physical performance and patient-reported outcomes. Also, since K-SES scores were only reported at 6 years after injury, we could not adjust for baseline scores or analyze change in K-SES scores. Although it may be simplest to speculate that improved physical performance causes better PROs, it is possible that individuals reporting better PROs engage in activities that in turn improve physical performance. Therefore, future prospective studies are required to fully understand the association between physical performance and PROs after ACL injury.

In study IV, we chose only to use the LSI to represent the values of the physical performance tests. However, studies have shown that bilateral deficits are present after ACL injury (111, 199, 200), which may result in misleading LSI values. Furthermore, results from previous studies have shown that the use of LSI may overestimate physical performance in comparison to the use of absolute values (201) or the use of LSI adjusted for BMI (202). However, the use of LSI is common in clinical studies and is included in the criteria for successful outcome (84) and return to sports (115) after ALC injury and ACLR.

Clinical implications

The ARS can be used to assess the frequency of activity participation in individuals that are physically active in cutting and pivoting sports. When the frequency of activity participation before and after a knee injury is compared, a recall period of one year can be used. During and after knee injury treatment, a recall period of one month or a week can be used when knee-demanding exercises or activities have been introduced to the exercise program. As many PROMs, the ARS is appropriate for comparisons on a group level. However, on an individual level, the change in ARS score must be relatively high (> 5 points, on the 0-16 point scale), to exclude measurement error.

Individuals that are highly physically active and participate in high-demand sports and activities need to be assessed accordingly in their physical activity participation. In clinical practice and in research of knee injuries, it is useful and relevant to evaluate the frequency of activity participation in activities that are challenging for the knee, in addition to the level or type of activity participation. However, the frequency and the level or type of activity do not necessarily converge. Individuals can be frequently physically active and place high demands on their knees, despite being physically active at a low recreational or competitive level. For that reason, the ARS can be used adjacent with self-reported measures of knee-specific activity level and knee function, e.g. the TAS and the KOOS, to assess treatment outcome after knee injury.

In the evaluation of outcome after treatment, it is important to recognize the responsiveness of the PROMS that are used to enable a valid evaluation. In this thesis, when comparing several knee-specific and generic PROMs, we found that the KOOS subscales Sport/Rec and QOL, the K-SES Present and the SF-36 Physical Component Summary appear to have the highest ability to detect change in outcomes that are relevant to individuals after completing rehabilitation for a knee injury. Furthermore, the ARS appears to be more appropriate than the TAS for evaluation of change in knee-specific activity participation in individuals after undergoing rehabilitation for a knee injury.

The results of this thesis support previous findings suggesting that rehabilitation of an ACL injury should include exercises aimed at optimizing and maintaining good single-leg physical performance and symmetry between legs in hop and strength performance. In particular, single-leg hop performance, knee flexion and extension strength in addition to postural orientation appear to be important factors for future self-reported

knee-related function, knee-specific activity participation and knee-related self-efficacy. In contrast, ACL treatment strategy, in terms of exercise therapy only, or exercise therapy in addition to an early or an optional delayed ACL reconstruction, does not appear to have an impact on knee-related self-efficacy at 6 years after ACL injury or ACLR.

Conclusions

- The Swedish version of the ARS is a valid, reliable and responsive instrument to evaluate activity participation in terms of frequency of participation in kneedemanding sporting activities in young adults with a knee injury.
- The KOOS subscales Sport/Rec and QOL, the K-SES Present and the SF-36
 Physical Component Summary showed the greatest potential for detecting
 change in outcomes relevant to patients after completing knee injury
 rehabilitation. The ARS may be preferred over TAS to optimize the evaluation
 of change in knee-specific activity participation following knee injury
 rehabilitation.
- The performance in the vertical hop and the side hop tests and postural orientation at 3 years were moderately associated with self-reported knee function and knee-specific activity participation, at 3 and 5 years after ACL injury/ACLR. Knee extension strength at 3 years was more associated with self-reported frequency rather than type and intensity of knee-specific activity participation at 5 years after ACL injury/ACLR.
- Knee-related self-efficacy at 6 years after an acute ACL injury/ACLR did not differ between patients treated with ACL reconstruction, performed early or as a delayed procedure, or those treated with exercise therapy alone. The self-efficacy for present knee function capability was higher than the self-efficacy for return to preinjury level sport and not being reinjured. Worse knee flexion strength LSI at the end of the exercise therapy, and worse LSI for the single-leg hop test at 5 years, correlated moderately with worse knee-related self-efficacy at 6 years. Targeted training to improve the symmetry between legs in knee flexion muscle strength and in the single-leg hop for distance may have a positive, yet small, impact on knee-related self-efficacy after ACL injury/ACLR.

Acknowledgements

In particular, I want to thank all the people that have participated in the studies that are included in this thesis. Your contribution made my thesis possible.

I am grateful to Eva Ageberg, my main supervisor, for introducing me to the world of research. Thank you for generously sharing your vast knowledge and vision. I truly value the time, dedication and ambition you have put into our journey.

Ewa M. Roos, my co-supervisor, thank you for sharing your knowledge in the research field, your strategic thinking and professional input.

I want to thank all my colleagues, Anders, Anna, Jenny, August, Niklas, Sofia RA, Ioannis, Anja and Sofia B, for creating the greatest working environment ever and making me feel a part of a fantastic team. Thank you for the discussions, all your input and last but not least all the laughs. You have upgraded my days over, and, over again. I genuinely thank you for your part in this journey.



The Musculoskeletal Function Research Group:
Anders Pålsson, Niklas Cedeström, Vala Flosadottir, Anna Cronström, Eva Ageberg, Sofia Ryman Augustsson,
August Estberger and Sofia Bunke. Absent: Jenny Nae, Ioannis Kostogiannis and Anja Eskilsson

I would also like to thank the following:

Richard Frobell, co-author, for your enthusiasm, ideas and rapid response.

The KANON steering group for kindly agreeing to let me to use your massive data.

Björn Schlaug for patiently providing numerous spreadsheets from the KANON database.

The staff at the physical therapy clinics; Kulan, Idrottsskademottagningen in Höör, Arena Fysio and Arena City in Helsingborg, Gerdahallens sjukgymnastik in Lund, iKlinik in Lund and Malmö, for facilitating the recruitment of participants to the studies.

Our statisticians, Vibeke, Christel and Tommy, for your valuable advice and guidance in the hills and winding paths of statistics.

I want to thank Clare Ardern and Markus Waldén for taking the time to read my work at the half-time review. Your input and comments certainly improved my work.

All my fellow PhD students, researchers and teachers, which I have come across in courses and committees during my research education. You have sparked my motivation and curiosity. You have also elevated my ability to reflect and be creative.

Björg and Hjalti, for leading with such a great example in everyday life and academic work.

All my family; in particular my mom and dad, my sister Lára, Jovan, Viggó, Göran, and all friends for giving me a healthy perspective and for your endless and loving support. You are my best!

My wonderful husband, Magnus and our amazing daughters, Emma and Klara, for reminding me of the most important things in life. I love you, always.

References

- 1. Majewski M, Susanne H, Klaus S. Epidemiology of athletic knee injuries: A 10-year study. The Knee. 2006;13(3):184-8.
- 2. Muthuri SG, McWilliams DF, Doherty M, Zhang W. History of knee injuries and knee osteoarthritis: a meta-analysis of observational studies. Osteoarthritis and cartilage / OARS, Osteoarthritis Research Society. 2011;19(11):1286-93.
- 3. Lohmander LS, Ostenberg A, Englund M, Roos H. High prevalence of knee osteoarthritis, pain, and functional limitations in female soccer players twelve years after anterior cruciate ligament injury. Arthritis and rheumatism. 2004;50(10):3145-52.
- 4. von Porat A, Roos EM, Roos H. High prevalence of osteoarthritis 14 years after an anterior cruciate ligament tear in male soccer players: a study of radiographic and patient relevant outcomes. Annals of the rheumatic diseases. 2004;63(3):269-73.
- 5. Griffin LY, Albohm MJ, Arendt EA, Bahr R, Beynnon BD, Demaio M, et al. Understanding and preventing noncontact anterior cruciate ligament injuries: a review of the Hunt Valley II meeting, January 2005. The American journal of sports medicine. 2006;34(9):1512-32.
- 6. Logerstedt DS, Snyder-Mackler L, Ritter RC, Axe MJ. Knee pain and mobility impairments: meniscal and articular cartilage lesions. The Journal of orthopaedic and sports physical therapy. 2010;40(6):A1-a35.
- Gage BE, McIlvain NM, Collins CL, Fields SK, Comstock RD. Epidemiology of 6.6
 million knee injuries presenting to United States emergency departments from 1999
 through 2008. Academic emergency medicine: official journal of the Society for
 Academic Emergency Medicine. 2012;19(4):378-85.
- 8. Åman M. Acute sports injuries in Sweden and their possible prevention: an epidemiological study using insurance data. Stockholm: The Swedish School of Sport and Health Sciences, GIH; 2018.
- 9. Frobell RB, Lohmander LS, Roos HP. Acute rotational trauma to the knee: poor agreement between clinical assessment and magnetic resonance imaging findings. Scandinavian journal of medicine & science in sports. 2007;17(2):109-14.
- Nordenvall R, Bahmanyar S, Adami J, Stenros C, Wredmark T, Fellander-Tsai L. A
 population-based nationwide study of cruciate ligament injury in Sweden, 2001-2009:
 incidence, treatment, and sex differences. The American journal of sports medicine.
 2012;40(8):1808-13.

- 11. Lohmander LS, Englund PM, Dahl LL, Roos EM. The long-term consequence of anterior cruciate ligament and meniscus injuries: osteoarthritis. The American journal of sports medicine. 2007;35(10):1756-69.
- 12. Mall NA, Chalmers PN, Moric M, Tanaka MJ, Cole BJ, Bach BR, Jr., et al. Incidence and trends of anterior cruciate ligament reconstruction in the United States. The American journal of sports medicine. 2014;42(10):2363-70.
- 13. Montalvo AM, Schneider DK, Yut L, Webster KE, Beynnon B, Kocher MS, et al. "What's my risk of sustaining an ACL injury while playing sports?" A systematic review with meta-analysis. British journal of sports medicine. 2018.
- 14. Englund M, Roemer FW, Hayashi D, Crema MD, Guermazi A. Meniscus pathology, osteoarthritis and the treatment controversy. Nature reviews Rheumatology. 2012;8(7):412-9.
- 15. Solheim E, Krokeide AM, Melteig P, Larsen A, Strand T, Brittberg M. Symptoms and function in patients with articular cartilage lesions in 1,000 knee arthroscopies. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2016;24(5):1610-6.
- 16. Mandelbaum BR, Browne JE, Fu F, Micheli L, Mosely JB, Jr., Erggelet C, et al. Articular cartilage lesions of the knee. The American journal of sports medicine. 1998;26(6):853-61.
- Schneider DK, Grawe B, Magnussen RA, Ceasar A, Parikh SN, Wall EJ, et al.
 Outcomes After Isolated Medial Patellofemoral Ligament Reconstruction for the
 Treatment of Recurrent Lateral Patellar Dislocations: A Systematic Review and Metaanalysis. The American journal of sports medicine. 2016;44(11):2993-3005.
- 18. Seedhom BB, Dowson D, Wright V. Proceedings: Functions of the menisci. A preliminary study. Annals of the rheumatic diseases. 1974;33(1):111.
- 19. Shrive NG, O'Connor JJ, Goodfellow JW. Load-bearing in the knee joint. Clinical orthopaedics and related research. 1978(131):279-87.
- 20. Walker PS, Erkman MJ. The role of the menisci in force transmission across the knee. Clinical orthopaedics and related research. 1975(109):184-92.
- 21. Fukubayashi T, Kurosawa H. The contact area and pressure distribution pattern of the knee. A study of normal and osteoarthrotic knee joints. Acta orthopaedica Scandinavica. 1980;51(6):871-9.
- 22. Kurosawa H, Fukubayashi T, Nakajima H. Load-bearing mode of the knee joint: physical behavior of the knee joint with or without menisci. Clinical orthopaedics and related research. 1980(149):283-90.
- 23. Bicer EK, Lustig S, Servien E, Selmi TA, Neyret P. Current knowledge in the anatomy of the human anterior cruciate ligament. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2010;18(8):1075-84.
- 24. Schillhammer CK, Reid JB, 3rd, Rister J, Jani SS, Marvil SC, Chen AW, et al. Arthroscopy Up to Date: Anterior Cruciate Ligament Anatomy. Arthroscopy: the

- journal of arthroscopic & related surgery: official publication of the Arthroscopy Association of North America and the International Arthroscopy Association. 2016;32(1):209-12.
- 25. Levangie PK NC. Joint structure and function. A comprehensive analysis. Philadelphia: F.A. Davis Company; 2005.
- Hart HF, Culvenor AG, Collins NJ, Ackland DC, Cowan SM, Machotka Z, et al. Knee kinematics and joint moments during gait following anterior cruciate ligament reconstruction: a systematic review and meta-analysis. British journal of sports medicine. 2015.
- 27. Chaudhari AM, Briant PL, Bevill SL, Koo S, Andriacchi TP. Knee kinematics, cartilage morphology, and osteoarthritis after ACL injury. Medicine and science in sports and exercise. 2008;40(2):215-22.
- 28. Petersen W, Taheri P, Forkel P, Zantop T. Return to play following ACL reconstruction: a systematic review about strength deficits. Archives of orthopaedic and trauma surgery. 2014;134(10):1417-28.
- Thorlund JB, Ostengaard L, Cardy N, Wilson F, Jorgensen C, Juhl CB. Trajectory of self-reported pain and function and knee extensor muscle strength in young patients undergoing arthroscopic surgery for meniscal tears: A systematic review and metaanalysis. Journal of science and medicine in sport / Sports Medicine Australia. 2017;20(8):712-7.
- 30. Hatton AL, Crossley KM, Clark RA, Whitehead TS, Morris HG, Culvenor AG. Between-leg differences in challenging single-limb balance performance one year following anterior cruciate ligament reconstruction. Gait & posture. 2017;52:22-5.
- Grindem H, Snyder-Mackler L, Moksnes H, Engebretsen L, Risberg MA. Simple decision rules can reduce reinjury risk by 84% after ACL reconstruction: the Delaware-Oslo ACL cohort study. British journal of sports medicine. 2016;50(13):804-8.
- 32. Oiestad BE, Engebretsen L, Storheim K, Risberg MA. Knee osteoarthritis after anterior cruciate ligament injury: a systematic review. The American journal of sports medicine. 2009;37(7):1434-43.
- 33. Ajuied A, Wong F, Smith C, Norris M, Earnshaw P, Back D, et al. Anterior cruciate ligament injury and radiologic progression of knee osteoarthritis: a systematic review and meta-analysis. The American journal of sports medicine. 2014;42(9):2242-52.
- 34. Khan T, Alvand A, Prieto-Alhambra D, Culliford DJ, Judge A, Jackson WF, et al. ACL and meniscal injuries increase the risk of primary total knee replacement for osteoarthritis: a matched case-control study using the Clinical Practice Research Datalink (CPRD). British journal of sports medicine. 2018.
- 35. Mather RC, 3rd, Koenig L, Kocher MS, Dall TM, Gallo P, Scott DJ, et al. Societal and economic impact of anterior cruciate ligament tears. The Journal of bone and joint surgery American volume. 2013;95(19):1751-9.

- 36. Williams GN, Chmielewski T, Rudolph K, Buchanan TS, Snyder-Mackler L. Dynamic knee stability: current theory and implications for clinicians and scientists. The Journal of orthopaedic and sports physical therapy. 2001;31(10):546-66.
- 37. Houck J, Yack HJ. Giving way event during a combined stepping and crossover cutting task in an individual with anterior cruciate ligament deficiency. The Journal of orthopaedic and sports physical therapy. 2001;31(9):481-9; discussion 90-5.
- 38. Chmielewski TL, Rudolph KS, Snyder-Mackler L. Development of dynamic knee stability after acute ACL injury. Journal of electromyography and kinesiology: official journal of the International Society of Electrophysiological Kinesiology. 2002;12(4):267-74.
- 39. Weiler R, Monte-Colombo M, Mitchell A, Haddad F. Non-operative management of a complete anterior cruciate ligament injury in an English Premier League football player with return to play in less than 8 weeks: applying common sense in the absence of evidence. BMJ case reports. 2015;2015.
- 40. Frobell RB, Roos EM, Roos HP, Ranstam J, Lohmander LS. A randomized trial of treatment for acute anterior cruciate ligament tears. The New England journal of medicine. 2010;363(4):331-42.
- 41. Frobell RB, Roos HP, Roos EM, Roemer FW, Ranstam J, Lohmander LS. Treatment for acute anterior cruciate ligament tear: five year outcome of randomised trial. BMJ (Clinical research ed). 2013;346:f232.
- 42. Grindem H, Risberg MA, Eitzen I. Two factors that may underpin outstanding outcomes after ACL rehabilitation. British journal of sports medicine. 2015.
- 43. Nagelli CV, Hewett TE. Should Return to Sport be Delayed Until 2 Years After Anterior Cruciate Ligament Reconstruction? Biological and Functional Considerations. Sports medicine (Auckland, NZ). 2017;47(2):221-32.
- 44. Hopper DM, Strauss GR, Boyle JJ, Bell J. Functional recovery after anterior cruciate ligament reconstruction: a longitudinal perspective. Archives of physical medicine and rehabilitation. 2008;89(8):1535-41.
- 45. Failla MJ, Arundale AJH, Logerstedt DS, Snyder-Mackler L. Controversies in Knee Rehabilitation Anterior Cruciate Ligament Injury. Clinics in sports medicine. 2015;34(2):301-+.
- 46. Palmieri-Smith RM, Thomas AC, Wojtys EM. Maximizing quadriceps strength after ACL reconstruction. Clinics in sports medicine. 2008;27(3):405-24, vii-ix.
- 47. Lautamies R, Harilainen A, Kettunen J, Sandelin J, Kujala UM. Isokinetic quadriceps and hamstring muscle strength and knee function 5 years after anterior cruciate ligament reconstruction: comparison between bone-patellar tendon-bone and hamstring tendon autografts. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2008;16(11):1009-16.
- 48. Ageberg E. Consequences of a ligament injury on neuromuscular function and relevance to rehabilitation using the anterior cruciate ligament-injured knee as model. Journal of

- electromyography and kinesiology: official journal of the International Society of Electrophysiological Kinesiology. 2002;12(3):205-12.
- 49. O'Donnell K, Freedman KB, Tjoumakaris FP. Rehabilitation Protocols After Isolated Meniscal Repair: A Systematic Review. The American journal of sports medicine. 2017;45(7):1687-97.
- Logerstedt DS, Scalzitti DA, Bennell KL, Hinman RS, Silvers-Granelli H, Ebert J, et al. Knee Pain and Mobility Impairments: Meniscal and Articular Cartilage Lesions Revision 2018. The Journal of orthopaedic and sports physical therapy. 2018;48(2):A1-a50.
- 51. Irrgang JJ, Fitzgerald GK. Rehabilitation of the multiple-ligament-injured knee. Clinics in sports medicine. 2000;19(3):545-71.
- 52. Mithoefer K, Hambly K, Logerstedt D, Ricci M, Silvers H, Della Villa S. Current concepts for rehabilitation and return to sport after knee articular cartilage repair in the athlete. The Journal of orthopaedic and sports physical therapy. 2012;42(3):254-73.
- 53. Moksnes H, Snyder-Mackler L, Risberg MA. Individuals with an anterior cruciate ligament-deficient knee classified as noncopers may be candidates for nonsurgical rehabilitation. The Journal of orthopaedic and sports physical therapy. 2008;38(10):586-95.
- 54. Ahlden M, Kvist J, Samuelsson K, Eriksson KO, Karlsson J. [Individualized therapy is important in anterior cruciate ligament injuries]. Lakartidningen. 2014;111(36):1440-3.
- 55. Stefancin JJ, Parker RD. First-time traumatic patellar dislocation: a systematic review. Clinical orthopaedics and related research. 2007;455:93-101.
- 56. Grevnerts HT, Faltstrom A, Sonesson S, Gauffin H, Carlfjord S, Kvist J. Activity demands and instability are the most important factors for recommending to treat ACL injuries with reconstruction. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2018.
- 57. Adams D, Logerstedt DS, Hunter-Giordano A, Axe MJ, Snyder-Mackler L. Current concepts for anterior cruciate ligament reconstruction: a criterion-based rehabilitation progression. The Journal of orthopaedic and sports physical therapy. 2012;42(7):601-14.
- 58. Monk AP, Davies LJ, Hopewell S, Harris K, Beard DJ, Price AJ. Surgical versus conservative interventions for treating anterior cruciate ligament injuries. The Cochrane database of systematic reviews. 2016;4:CD011166.
- 59. Smith TO, Postle K, Penny F, McNamara I, Mann CJ. Is reconstruction the best management strategy for anterior cruciate ligament rupture? A systematic review and meta-analysis comparing anterior cruciate ligament reconstruction versus non-operative treatment. The Knee. 2014;21(2):462-70.
- 60. Delince P, Ghafil D. Anterior cruciate ligament tears: conservative or surgical treatment? A critical review of the literature. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2012;20(1):48-61.

- 61. Meunier A, Odensten M, Good L. Long-term results after primary repair or nonsurgical treatment of anterior cruciate ligament rupture: a randomized study with a 15year follow-up. Scandinavian journal of medicine & science in sports. 2007;17(3):230-7
- 62. Fink C, Hoser C, Hackl W, Navarro RA, Benedetto KP. Long-term outcome of operative or nonoperative treatment of anterior cruciate ligament rupture--is sports activity a determining variable? International journal of sports medicine. 2001;22(4):304-9.
- 63. Marx RG, Jones EC, Angel M, Wickiewicz TL, Warren RF. Beliefs and attitudes of members of the American Academy of Orthopaedic Surgeons regarding the treatment of anterior cruciate ligament injury. Arthroscopy: the journal of arthroscopic & related surgery: official publication of the Arthroscopy Association of North America and the International Arthroscopy Association. 2003;19(7):762-70.
- 64. Heijne A, Axelsson K, Werner S, Biguet G. Rehabilitation and recovery after anterior cruciate ligament reconstruction: patients' experiences. Scandinavian journal of medicine & science in sports. 2008;18(3):325-35.
- 65. Feucht MJ, Cotic M, Saier T, Minzlaff P, Plath JE, Imhoff AB, et al. Patient expectations of primary and revision anterior cruciate ligament reconstruction. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2016;24(1):201-7.
- 66. Gobbi A, Francisco R. Factors affecting return to sports after anterior cruciate ligament reconstruction with patellar tendon and hamstring graft: a prospective clinical investigation. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2006;14(10):1021-8.
- 67. Ardern CL, Taylor NF, Feller JA, Webster KE. Fifty-five per cent return to competitive sport following anterior cruciate ligament reconstruction surgery: an updated systematic review and meta-analysis including aspects of physical functioning and contextual factors. British journal of sports medicine. 2014;48(21):1543-52.
- 68. Filbay SR, Culvenor AG, Ackerman IN, Russell TG, Crossley KM. Quality of life in anterior cruciate ligament-deficient individuals: a systematic review and meta-analysis. British journal of sports medicine. 2015;49(16):1033-41.
- 69. Ageberg E, Thomee R, Neeter C, Silbernagel KG, Roos EM. Muscle strength and functional performance in patients with anterior cruciate ligament injury treated with training and surgical reconstruction or training only: a two to five-year followup. Arthritis and rheumatism. 2008;59(12):1773-9.
- 70. Ericsson YB, Roos EM, Frobell RB. Lower extremity performance following ACL rehabilitation in the KANON-trial: impact of reconstruction and predictive value at 2 and 5 years. British journal of sports medicine. 2013;47(15):980-5.

- 71. Streich NA, Zimmermann D, Bode G, Schmitt H. Reconstructive versus non-reconstructive treatment of anterior cruciate ligament insufficiency. A retrospective matched-pair long-term follow-up. International orthopaedics. 2011;35(4):607-13.
- 72. Kessler MA, Behrend H, Henz S, Stutz G, Rukavina A, Kuster MS. Function, osteoarthritis and activity after ACL-rupture: 11 years follow-up results of conservative versus reconstructive treatment. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2008;16(5):442-8.
- 73. Meuffels DE, Favejee MM, Vissers MM, Heijboer MP, Reijman M, Verhaar JA. Ten year follow-up study comparing conservative versus operative treatment of anterior cruciate ligament ruptures. A matched-pair analysis of high level athletes. British journal of sports medicine. 2009;43(5):347-51.
- 74. Filbay SR, Roos EM, Frobell RB, Roemer F, Ranstam J, Lohmander LS. Delaying ACL reconstruction and treating with exercise therapy alone may alter prognostic factors for 5-year outcome: an exploratory analysis of the KANON trial. British journal of sports medicine. 2017.
- 75. Grindem H, Granan LP, Risberg MA, Engebretsen L, Snyder-Mackler L, Eitzen I. How does a combined preoperative and postoperative rehabilitation programme influence the outcome of ACL reconstruction 2 years after surgery? A comparison between patients in the Delaware-Oslo ACL Cohort and the Norwegian National Knee Ligament Registry. British journal of sports medicine. 2014.
- 76. Eitzen I, Holm I, Risberg MA. Preoperative quadriceps strength is a significant predictor of knee function two years after anterior cruciate ligament reconstruction. British journal of sports medicine. 2009;43(5):371-6.
- 77. Clancy WG, Jr. Knee ligamentous injury in sports: the past, present, and future. Medicine and science in sports and exercise. 1983;15(1):9-14.
- 78. Beynnon BD, Johnson RJ, Abate JA, Fleming BC, Nichols CE. Treatment of anterior cruciate ligament injuries, part I. The American journal of sports medicine. 2005;33(10):1579-602.
- 79. Beynnon BD, Johnson RJ, Abate JA, Fleming BC, Nichols CE. Treatment of anterior cruciate ligament injuries, part 2. The American journal of sports medicine. 2005;33(11):1751-67.
- 80. van Melick N, van Cingel RE, Brooijmans F, Neeter C, van Tienen T, Hullegie W, et al. Evidence-based clinical practice update: practice guidelines for anterior cruciate ligament rehabilitation based on a systematic review and multidisciplinary consensus. British journal of sports medicine. 2016;50(24):1506-15.
- 81. Ardern CL, Ekas G, Grindem H, Moksnes H, Anderson A, Chotel F, et al. 2018 International Olympic Committee consensus statement on prevention, diagnosis and management of paediatric anterior cruciate ligament (ACL) injuries. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2018.

- 82. Bent NP, Wright CC, Rushton AB, Batt ME. Selecting outcome measures in sports medicine: a guide for practitioners using the example of anterior cruciate ligament rehabilitation. British journal of sports medicine. 2009;43(13):1006-12.
- 83. Haines S, Baker T, Donaldson M. Development of a physical performance assessment checklist for athletes who sustained a lower extremity injury in preparation for return to sport: a delphi study. International journal of sports physical therapy. 2013;8(1):44-53.
- 84. Lynch AD, Logerstedt DS, Grindem H, Eitzen I, Hicks GE, Axe MJ, et al. Consensus criteria for defining 'successful outcome' after ACL injury and reconstruction: a Delaware-Oslo ACL cohort investigation. British journal of sports medicine. 2015;49(5):335-42.
- 85. Collins NJ, Misra D, Felson DT, Crossley KM, Roos EM. Measures of knee function: International Knee Documentation Committee (IKDC) Subjective Knee Evaluation Form, Knee Injury and Osteoarthritis Outcome Score (KOOS), Knee Injury and Osteoarthritis Outcome Score Physical Function Short Form (KOOS-PS), Knee Outcome Survey Activities of Daily Living Scale (KOS-ADL), Lysholm Knee Scoring Scale, Oxford Knee Score (OKS), Western Ontario and McMaster Universities Osteoarthritis Index (WOMAC), Activity Rating Scale (ARS), and Tegner Activity Score (TAS). Arthritis care & research. 2011;63 Suppl 11:S208-28.
- 86. Busija L, Osborne RH, Nilsdotter A, Buchbinder R, Roos EM. Magnitude and meaningfulness of change in SF-36 scores in four types of orthopedic surgery. Health and quality of life outcomes. 2008;6:55.
- 87. Harston A, Nyland J, Brand E, McGinnis M, Caborn DN. Collagen meniscus implantation: a systematic review including rehabilitation and return to sports activity. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2012;20(1):135-46.
- 88. Ardern CL, Taylor NF, Feller JA, Webster KE. A systematic review of the psychological factors associated with returning to sport following injury. British journal of sports medicine. 2013;47(17):1120-6.
- 89. Filbay SR, Ackerman IN, Russell TG, Crossley KM. Return to sport matters-longerterm quality of life after ACL reconstruction in people with knee difficulties. Scandinavian journal of medicine & science in sports. 2017;27(5):514-24.
- 90. U.S. Department of Health and Human Services FDA Center for Drug Evaluation and Research USDoHaHSFCfBEaR. Guidance for industry: patient-reported outcome measures: use in medical product development to support labeling claims: draft guidance. Health and quality of life outcomes. 2006;4:79.
- 91. Brazier JE, Harper R, Munro J, Walters SJ, Snaith ML. Generic and condition-specific outcome measures for people with osteoarthritis of the knee. Rheumatology (Oxford, England). 1999;38(9):870-7.
- 92. Mokkink LB, Terwee CB, Patrick DL, Alonso J, Stratford PW, Knol DL, et al. The COSMIN study reached international consensus on taxonomy, terminology, and

- definitions of measurement properties for health-related patient-reported outcomes. Journal of clinical epidemiology. 2010;63(7):737-45.
- 93. Ageberg E, Forssblad M, Herbertsson P, Roos EM. Sex differences in patient-reported outcomes after anterior cruciate ligament reconstruction: data from the Swedish knee ligament register. The American journal of sports medicine. 2010;38(7):1334-42.
- 94. Ardern CL, Taylor NF, Feller JA, Webster KE. Return-to-sport outcomes at 2 to 7 years after anterior cruciate ligament reconstruction surgery. The American journal of sports medicine. 2012;40(1):41-8.
- 95. Garcia GH, Wu HH, Park MJ, Tjoumakaris FP, Tucker BS, Kelly JDt, et al. Depression Symptomatology and Anterior Cruciate Ligament Injury: Incidence and Effect on Functional Outcome-A Prospective Cohort Study. The American journal of sports medicine. 2015.
- 96. Myklebust G, Bahr R. Return to play guidelines after anterior cruciate ligament surgery. British journal of sports medicine. 2005;39(3):127-31.
- 97. Ardern CL, Osterberg A, Sonesson S, Gauffin H, Webster KE, Kvist J. Satisfaction With Knee Function After Primary Anterior Cruciate Ligament Reconstruction Is Associated With Self-Efficacy, Quality of Life, and Returning to the Preinjury Physical Activity. Arthroscopy: the journal of arthroscopic & related surgery: official publication of the Arthroscopy Association of North America and the International Arthroscopy Association. 2016;32(8):1631-8.e3.
- 98. Sonesson S, Kvist J, Ardern C, Osterberg A, Silbernagel KG. Psychological factors are important to return to pre-injury sport activity after anterior cruciate ligament reconstruction: expect and motivate to satisfy. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2017;25(5):1375-84.
- 99. American College of Sports Medicine AAoFP, American Academy of Orthopaedic Surgeons ea. Psychological issues related to injury in athletes and the team physician: a consensus statement. Medicine and science in sports and exercise. 2006;38(11):2030-4.
- 100. Wiese-Bjornstal DM. Psychology and socioculture affect injury risk, response, and recovery in high-intensity athletes: a consensus statement. Scandinavian journal of medicine & science in sports. 2010;20 Suppl 2:103-11.
- 101. Ardern CL, Taylor NF, Feller JA, Whitehead TS, Webster KE. Psychological responses matter in returning to preinjury level of sport after anterior cruciate ligament reconstruction surgery. The American journal of sports medicine. 2013;41(7):1549-58.
- 102. te Wierike SCM, van der Sluis A, van den Akker-Scheek I, Elferink-Gemser MT, Visscher C. Psychosocial factors influencing the recovery of athletes with anterior cruciate ligament injury: A systematic review. Scandinavian journal of medicine & science in sports. 2013;23(5):527-40.
- 103. Ardern CL, Taylor NF, Feller JA, Whitehead TS, Webster KE. Sports participation 2 years after anterior cruciate ligament reconstruction in athletes who had not returned to

- sport at 1 year: a prospective follow-up of physical function and psychological factors in 122 athletes. The American journal of sports medicine. 2015;43(4):848-56.
- 104. Webster KE, Feller JA, Lambros C. Development and preliminary validation of a scale to measure the psychological impact of returning to sport following anterior cruciate ligament reconstruction surgery. Physical therapy in sport: official journal of the Association of Chartered Physiotherapists in Sports Medicine. 2008;9(1):9-15.
- 105. Thomee P, Wahrborg P, Borjesson M, Thomee R, Eriksson BI, Karlsson J. Self-efficacy of knee function as a pre-operative predictor of outcome 1 year after anterior cruciate ligament reconstruction. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2008;16(2):118-27.
- 106. Thomee P, Wahrborg P, Borjesson M, Thomee R, Eriksson BI, Karlsson J. A randomized, controlled study of a rehabilitation model to improve knee-function self-efficacy with ACL injury. Journal of sport rehabilitation. 2010;19(2):200-13.
- 107. Chmielewski TL, George SZ, Tillman SM, Moser MW, Lentz TA, Indelicato PA, et al. Low- Versus High-Intensity Plyometric Exercise During Rehabilitation After Anterior Cruciate Ligament Reconstruction. The American journal of sports medicine. 2016;44(3):609-17.
- 108. Abrams GD, Harris JD, Gupta AK, McCormick FM, Bush-Joseph CA, Verma NN, et al. Functional Performance Testing After Anterior Cruciate Ligament Reconstruction: A Systematic Review. Orthopaedic journal of sports medicine. 2014;2(1):2325967113518305.
- 109. Reid A, Birmingham TB, Stratford PW, Alcock GK, Giffin JR. Hop testing provides a reliable and valid outcome measure during rehabilitation after anterior cruciate ligament reconstruction. Physical therapy. 2007;87(3):337-49.
- 110. Engelen-van Melick N, van Cingel RE, Tijssen MP, Nijhuis-van der Sanden MW. Assessment of functional performance after anterior cruciate ligament reconstruction: a systematic review of measurement procedures. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2013;21(4):869-79.
- 111. Ferber R, Osternig LR, Woollacott MH, Wasielewski NJ, Lee JH. Bilateral accommodations to anterior cruciate ligament deficiency and surgery. Clinical biomechanics (Bristol, Avon). 2004;19(2):136-44.
- 112. Larsen JB, Farup J, Lind M, Dalgas U. Muscle strength and functional performance is markedly impaired at the recommended time point for sport return after anterior cruciate ligament reconstruction in recreational athletes. Human movement science. 2015;39:73-87.
- 113. Wellsandt E, Failla MJ, Snyder-Mackler L. Limb Symmetry Indexes Can Overestimate Knee Function After Anterior Cruciate Ligament Injury. The Journal of orthopaedic and sports physical therapy. 2017;47(5):334-8.

- 114. Logerstedt D, Arundale A, Lynch A, Snyder-Mackler L. A conceptual framework for a sports knee injury performance profile (SKIPP) and return to activity criteria (RTAC). Brazilian journal of physical therapy. 2015.
- 115. Thomee R, Kaplan Y, Kvist J, Myklebust G, Risberg MA, Theisen D, et al. Muscle strength and hop performance criteria prior to return to sports after ACL reconstruction. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2011;19(11):1798-805.
- 116. Gustavsson A, Neeter C, Thomee P, Silbernagel KG, Augustsson J, Thomee R, et al. A test battery for evaluating hop performance in patients with an ACL injury and patients who have undergone ACL reconstruction. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2006;14(8):778-88.
- 117. Neeter C, Gustavsson A, Thomee P, Augustsson J, Thomee R, Karlsson J. Development of a strength test battery for evaluating leg muscle power after anterior cruciate ligament injury and reconstruction. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2006;14(6):571-80.
- 118. Barfod KW, Feller JA, Clark R, Hartwig T, Devitt BM, Webster KE. Strength Testing Following Anterior Cruciate Ligament Reconstruction. A Prospective Cohort Study Investigating Overlap of Tests. Journal of strength and conditioning research / National Strength & Conditioning Association. 2018.
- 119. Horak FB. Postural orientation and equilibrium: what do we need to know about neural control of balance to prevent falls? Age and ageing. 2006;35 Suppl 2:ii7-ii11.
- 120. Trulsson A, Roos EM, Ageberg E, Garwicz M. Relationships between postural orientation and self reported function, hop performance and muscle power in subjects with anterior cruciate ligament injury. BMC musculoskeletal disorders. 2010;11:143.
- 121. Everhart JS, Best TM, Flanigan DC. Psychological predictors of anterior cruciate ligament reconstruction outcomes: a systematic review. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2015;23(3):752-62.
- 122. Kvist J, Ek A, Sporrstedt K, Good L. Fear of re-injury: a hindrance for returning to sports after anterior cruciate ligament reconstruction. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2005;13(5):393-7.
- 123. Wasserstein D, Huston LJ, Nwosu S, Kaeding CC, Parker RD, Wright RW, et al. KOOS pain as a marker for significant knee pain two and six years after primary ACL reconstruction: a Multicenter Orthopaedic Outcomes Network (MOON) prospective longitudinal cohort study. Osteoarthritis and cartilage / OARS, Osteoarthritis Research Society. 2015;23(10):1674-84.
- 124. Logerstedt D, Grindem H, Lynch A, Eitzen I, Engebretsen L, Risberg MA, et al. Single-legged hop tests as predictors of self-reported knee function after anterior cruciate ligament reconstruction: the Delaware-Oslo ACL cohort study. The American journal of sports medicine. 2012;40(10):2348-56.

- 125. Grindem H, Logerstedt D, Eitzen I, Moksnes H, Axe MJ, Snyder-Mackler L, et al. Single-legged hop tests as predictors of self-reported knee function in nonoperatively treated individuals with anterior cruciate ligament injury. The American journal of sports medicine. 2011;39(11):2347-54.
- 126. Pinczewski LA, Lyman J, Salmon LJ, Russell VJ, Roe J, Linklater J. A 10-year comparison of anterior cruciate ligament reconstructions with hamstring tendon and patellar tendon autograft: a controlled, prospective trial. The American journal of sports medicine. 2007;35(4):564-74.
- 127. Kyritsis P, Bahr R, Landreau P, Miladi R, Witvrouw E. Likelihood of ACL graft rupture: not meeting six clinical discharge criteria before return to sport is associated with a four times greater risk of rupture. British journal of sports medicine. 2016;50(15):946-51.
- 128. Christino MA, Fleming BC, Machan JT, Shalvoy RM. Psychological Factors Associated With Anterior Cruciate Ligament Reconstruction Recovery. Orthopaedic journal of sports medicine. 2016;4(3):2325967116638341.
- 129. Reinke EK, Spindler KP, Lorring D, Jones MH, Schmitz L, Flanigan DC, et al. Hop tests correlate with IKDC and KOOS at minimum of 2 years after primary ACL reconstruction. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2011;19(11):1806-16.
- 130. Ageberg E, Roos EM. The Association Between Knee Confidence and Muscle Power, Hop Performance, and Postural Orientation in People With Anterior Cruciate Ligament Injury. The Journal of orthopaedic and sports physical therapy. 2016;46(6):477-82.
- 131. Mokkink LB, Prinsen CA, Bouter LM, Vet HC, Terwee CB. The COnsensus-based Standards for the selection of health Measurement INstruments (COSMIN) and how to select an outcome measurement instrument. Brazilian journal of physical therapy. 2016;20(2):105-13.
- 132. Kise NJ, Risberg MA, Stensrud S, Ranstam J, Engebretsen L, Roos EM. Exercise therapy versus arthroscopic partial meniscectomy for degenerative meniscal tear in middle aged patients: randomised controlled trial with two year follow-up. BMJ (Clinical research ed). 2016;354:i3740.
- 133. Khan M, Evaniew N, Bedi A, Ayeni OR, Bhandari M. Arthroscopic surgery for degenerative tears of the meniscus: a systematic review and meta-analysis. CMAJ: Canadian Medical Association journal = journal de l'Association medicale canadienne. 2014;186(14):1057-64.
- 134. Roos EM, Hare KB, Nielsen SM, Christensen R, Lohmander LS. Better outcome from arthroscopic partial meniscectomy than skin incisions only? A sham-controlled randomised trial in patients aged 35-55 years with knee pain and an MRI-verified meniscal tear. BMJ open. 2018;8(2):e019461.
- 135. Aaronson N, Alonso J, Burnam A, Lohr KN, Patrick DL, Perrin E, et al. Assessing health status and quality-of-life instruments: attributes and review criteria. Quality of life

- research: an international journal of quality of life aspects of treatment, care and rehabilitation. 2002;11(3):193-205.
- 136. Bowling A. Measuring Disease: A Review of Disease-Specific Quality of Life Measurement Scales. 2nd ed. Buckingham: Open University Press; 2004.
- 137. Beaton DE, Bombardier C, Guillemin F, Ferraz MB. Guidelines for the process of cross-cultural adaptation of self-report measures. Spine. 2000;25(24):3186-91.
- 138. Mokkink LB, Terwee CB, Patrick DL, Alonso J, Stratford PW, Knol DL, et al. The COSMIN checklist for assessing the methodological quality of studies on measurement properties of health status measurement instruments: an international Delphi study. Quality of life research: an international journal of quality of life aspects of treatment, care and rehabilitation. 2010;19(4):539-49.
- 139. Mokkink LB, Terwee CB, Knol DL, Stratford PW, Alonso J, Patrick DL, et al. The COSMIN checklist for evaluating the methodological quality of studies on measurement properties: a clarification of its content. BMC medical research methodology. 2010;10:22.
- 140. Norman GR, Wyrwich KW, Patrick DL. The mathematical relationship among different forms of responsiveness coefficients. Quality of life research: an international journal of quality of life aspects of treatment, care and rehabilitation. 2007;16(5):815-22.
- 141. Cliff N. Dominance statistics: Ordinal analyses to answer ordinal questions. Psychological bulletin. 1993;114(3):494-509.
- 142. de vet HC, Terwee CB, Mokkink LB, Knol DL. Measurement in medicine A practical guide. New York: Cambridge University Press; 2011.
- 143. Terwee CB, Mokkink LB, van Poppel MN, Chinapaw MJ, van Mechelen W, de Vet HC. Qualitative attributes and measurement properties of physical activity questionnaires: a checklist. Sports medicine (Auckland, NZ). 2010;40(7):525-37.
- 144. Collins NJ, Prinsen CA, Christensen R, Bartels EM, Terwee CB, Roos EM. Knee Injury and Osteoarthritis Outcome Score (KOOS): systematic review and meta-analysis of measurement properties. Osteoarthritis and cartilage / OARS, Osteoarthritis Research Society. 2016.
- 145. Thomee P, Wahrborg P, Borjesson M, Thomee R, Eriksson BI, Karlsson J. A new instrument for measuring self-efficacy in patients with an anterior cruciate ligament injury. Scandinavian journal of medicine & science in sports. 2006;16(3):181-7.
- 146. Thomee P, Wahrborg P, Borjesson M, Thomee R, Eriksson BI, Karlsson J. Self-efficacy, symptoms and physical activity in patients with an anterior cruciate ligament injury: a prospective study. Scandinavian journal of medicine & science in sports. 2007;17(3):238-45.
- 147. Shapiro ET, Richmond JC, Rockett SE, McGrath MM, Donaldson WR. The use of a generic, patient-based health assessment (SF-36) for evaluation of patients with anterior

- cruciate ligament injuries. The American journal of sports medicine. 1996;24(2):196-200.
- 148. Sousa PL, Krych AJ, Cates RA, Levy BA, Stuart MJ, Dahm DL. Return to sport: Does excellent 6-month strength and function following ACL reconstruction predict midterm outcomes? Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2015.
- 149. Moller E, Weidenhielm L, Werner S. Outcome and knee-related quality of life after anterior cruciate ligament reconstruction: a long-term follow-up. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2009;17(7):786-94.
- 150. Bandura A. Self-efficacy: toward a unifying theory of behavioral change. Psychological review. 1977;84(2):191-215.
- 151. Marx RG, Stump TJ, Jones EC, Wickiewicz TL, Warren RF. Development and evaluation of an activity rating scale for disorders of the knee. The American journal of sports medicine. 2001;29(2):213-8.
- 152. Tegner Y, Lysholm J. Rating systems in the evaluation of knee ligament injuries. Clinical orthopaedics and related research. 1985(198):43-9.
- 153. Briggs KK, Lysholm J, Tegner Y, Rodkey WG, Kocher MS, Steadman JR. The reliability, validity, and responsiveness of the Lysholm score and Tegner activity scale for anterior cruciate ligament injuries of the knee: 25 years later. The American journal of sports medicine. 2009;37(5):890-7.
- 154. Briggs KK, Kocher MS, Rodkey WG, Steadman JR. Reliability, validity, and responsiveness of the Lysholm knee score and Tegner activity scale for patients with meniscal injury of the knee. The Journal of bone and joint surgery American volume. 2006;88(4):698-705.
- 155. Roos EM, Roos HP, Lohmander LS, Ekdahl C, Beynnon BD. Knee Injury and Osteoarthritis Outcome Score (KOOS)--development of a self-administered outcome measure. The Journal of orthopaedic and sports physical therapy. 1998;28(2):88-96.
- 156. Roos EM, Roos HP, Ekdahl C, Lohmander LS. Knee injury and Osteoarthritis Outcome Score (KOOS)--validation of a Swedish version. Scandinavian journal of medicine & science in sports. 1998;8(6):439-48.
- 157. Roos EM, Lohmander LS. The Knee injury and Osteoarthritis Outcome Score (KOOS): from joint injury to osteoarthritis. Health and quality of life outcomes. 2003;1:64.
- 158. Ware JE, Jr. SF-36 health survey update. Spine. 2000;25(24):3130-9.
- 159. Sullivan M, Karlsson J, Ware JE, Jr. The Swedish SF-36 Health Survey--I. Evaluation of data quality, scaling assumptions, reliability and construct validity across general populations in Sweden. Social science & medicine (1982). 1995;41(10):1349-58.
- 160. Sullivan M, Karlsson J. The Swedish SF-36 Health Survey III. Evaluation of criterion-based validity: results from normative population. Journal of clinical epidemiology. 1998;51(11):1105-13.

- 161. Saltin B, Grimby G. Physiological analysis of middle-aged and old former athletes. Comparison with still active athletes of the same ages. Circulation. 1968;38(6):1104-15.
- 162. Aires N, Selmer R, Thelle D. The validity of self-reported leisure time physical activity, and its relationship to serum cholesterol, blood pressure and body mass index. A population based study of 332,182 men and women aged 40-42 years. European journal of epidemiology. 2003;18(6):479-85.
- 163. Rodjer L, Jonsdottir IH, Rosengren A, Bjorck L, Grimby G, Thelle DS, et al. Self-reported leisure time physical activity: a useful assessment tool in everyday health care. BMC public health. 2012;12:693.
- 164. McCormack HM, Horne DJ, Sheather S. Clinical applications of visual analogue scales: a critical review. Psychological medicine. 1988;18(4):1007-19.
- 165. Hawker GA, Mian S, Kendzerska T, French M. Measures of adult pain: Visual Analog Scale for Pain (VAS Pain), Numeric Rating Scale for Pain (NRS Pain), McGill Pain Questionnaire (MPQ), Short-Form McGill Pain Questionnaire (SF-MPQ), Chronic Pain Grade Scale (CPGS), Short Form-36 Bodily Pain Scale (SF-36 BPS), and Measure of Intermittent and Constant Osteoarthritis Pain (ICOAP). Arthritis care & research. 2011;63 Suppl 11:S240-52.
- 166. van den Hoven LH, Gorter KJ, Picavet HS. Measuring musculoskeletal pain by questionnaires: the manikin versus written questions. European journal of pain (London, England). 2010;14(3):335-8.
- 167. Ohnmeiss DD. Repeatability of pain drawings in a low back pain population. Spine. 2000;25(8):980-8.
- 168. Drouin JM, Valovich-mcLeod TC, Shultz SJ, Gansneder BM, Perrin DH. Reliability and validity of the Biodex system 3 pro isokinetic dynamometer velocity, torque and position measurements. European journal of applied physiology. 2004;91(1):22-9.
- 169. Verdijk LB, van Loon L, Meijer K, Savelberg HH. One-repetition maximum strength test represents a valid means to assess leg strength in vivo in humans. Journal of sports sciences. 2009;27(1):59-68.
- 170. Ostenberg A, Roos E, Ekdahl C, Roos H. Isokinetic knee extensor strength and functional performance in healthy female soccer players. Scandinavian journal of medicine & science in sports. 1998;8(5 Pt 1):257-64.
- 171. Cohen J. Statistical power analysis for the behavioral sciences. 2nd ed. ed. New Jersey: Lawrence Erlbaum Associates, Inc.; 1988.
- 172. Terwee CB, Bot SDM, de Boer MR, van der Windt D, Knol DL, Dekker J, et al. Quality criteria were proposed for measurement properties of health status questionnaires. Journal of clinical epidemiology. 2007;60(1):34-42.
- 173. Bland JM, Altman DG. Statistical methods for assessing agreement between two methods of clinical measurement. Lancet. 1986;1(8476):307-10.

- 174. Vargha A, Delaney HD. A Critique and Improvement of the "CL" Common Language Effect Size Statistics of McGraw and Wong. Journal of Educational and Behavioral Statistics. 2000;25(2):101-32.
- 175. Negahban H, Mostafaee N, Sohani SM, Mazaheri M, Goharpey S, Salavati M, et al. Reliability and validity of the Tegner and Marx activity rating scales in Iranian patients with anterior cruciate ligament injury. Disability and rehabilitation. 2011;33(23-24):2305-10.
- 176. Collins NJ, Prinsen CA, Christensen R, Bartels EM, Terwee CB, Roos EM. Knee Injury and Osteoarthritis Outcome Score (KOOS): systematic review and meta-analysis of measurement properties. Osteoarthritis and cartilage / OARS, Osteoarthritis Research Society. 2016;24(8):1317-29.
- 177. Kvist J, Osterberg A, Gauffin H, Tagesson S, Webster K, Ardern C. Translation and measurement properties of the Swedish version of ACL-Return to Sports after Injury questionnaire. Scandinavian journal of medicine & science in sports. 2013;23(5):568-75.
- 178. Slagers AJ, Reininga IH, van den Akker-Scheek I. The Dutch language anterior cruciate ligament return to sport after injury scale (ACL-RSI) validity and reliability. Journal of sports sciences. 2016:1-9.
- 179. Kostogiannis I, Ageberg E, Neuman P, Dahlberg L, Friden T, Roos H. Activity level and subjective knee function 15 years after anterior cruciate ligament injury: a prospective, longitudinal study of nonreconstructed patients. The American journal of sports medicine. 2007;35(7):1135-43.
- 180. Letchford R, Button K, Sparkes V, van Deursen RWM. Assessing activity participation in the ACL injured population: a systematic review of activity rating scale measurement properties. Physical Therapy Reviews. 2012;17(2):99-109.
- 181. Patrick DL, Deyo RA. Generic and disease-specific measures in assessing health status and quality of life. Medical care. 1989;27(3 Suppl):S217-32.
- 182. Beaudart C, Biver E, Bruyere O, Cooper C, Al-Daghri N, Reginster JY, et al. Quality of life assessment in musculo-skeletal health. Aging clinical and experimental research. 2017.
- 183. Sernert N, Kartus J, Kohler K, Stener S, Larsson J, Eriksson BI, et al. Analysis of subjective, objective and functional examination tests after anterior cruciate ligament reconstruction. A follow-up of 527 patients. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 1999;7(3):160-5.
- 184. Neeb TB, Aufdemkampe G, Wagener JH, Mastenbroek L. Assessing anterior cruciate ligament injuries: the association and differential value of questionnaires, clinical tests, and functional tests. The Journal of orthopaedic and sports physical therapy. 1997;26(6):324-31.

- 185. Medicine ACoS. American College of Sports Medicine position stand. Progression models in resistance training for healthy adults. Medicine and science in sports and exercise. 2009;41(3):687-708.
- 186. Filbay SR, Crossley KM, Ackerman IN. Activity preferences, lifestyle modifications and re-injury fears influence longer-term quality of life in people with knee symptoms following anterior cruciate ligament reconstruction: a qualitative study. Journal of physiotherapy. 2016.
- 187. Tjong VK, Murnaghan ML, Nyhof-Young JM, Ogilvie-Harris DJ. A qualitative investigation of the decision to return to sport after anterior cruciate ligament reconstruction: to play or not to play. The American journal of sports medicine. 2014;42(2):336-42.
- 188. Möller E, Weidenhielm L, Werner S. Outcome and knee-related quality of life after anterior cruciate ligament reconstruction: a long-term follow-up. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2009;17(7):786-94.
- 189. Thomee P, Wahrborg P, Borjesson M, Thomee R, Eriksson BI, Karlsson J. Determinants of self-efficacy in the rehabilitation of patients with anterior cruciate ligament injury. Journal of rehabilitation medicine: official journal of the UEMS European Board of Physical and Rehabilitation Medicine. 2007;39(6):486-92.
- 190. Pihl K, Roos EM, Nissen N, JoRgensen U, Schjerning J, Thorlund JB. Over-optimistic patient expectations of recovery and leisure activities after arthroscopic meniscus surgery. Acta orthopaedica. 2016;87(6):615-21.
- 191. Ardern CL. Anterior Cruciate Ligament Reconstruction-Not Exactly a One-Way Ticket Back to the Preinjury Level: A Review of Contextual Factors Affecting Return to Sport After Surgery. Sports health. 2015;7(3):224-30.
- 192. Langford JL, Webster KE, Feller JA. A prospective longitudinal study to assess psychological changes following anterior cruciate ligament reconstruction surgery. British journal of sports medicine. 2009;43(5):377-81.
- 193. Myklebust G, Holm I, Maehlum S, Engebretsen L, Bahr R. Clinical, functional, and radiologic outcome in team handball players 6 to 11 years after anterior cruciate ligament injury: a follow-up study. The American journal of sports medicine. 2003;31(6):981-9.
- 194. Draugalis JR, Coons SJ, Plaza CM. Best practices for survey research reports: a synopsis for authors and reviewers. American journal of pharmaceutical education. 2008;72(1):11.
- 195. King MT. A point of minimal important difference (MID): a critique of terminology and methods. Expert review of pharmacoeconomics & outcomes research. 2011;11(2):171-84.
- 196. Terwee CB, Roorda LD, Dekker J, Bierma-Zeinstra SM, Peat G, Jordan KP, et al. Mind the MIC: large variation among populations and methods. Journal of clinical epidemiology. 2010;63(5):524-34.

- 197. Hegedus EJ, McDonough SM, Bleakley C, Baxter D, Cook CE. Clinician-friendly lower extremity physical performance tests in athletes: a systematic review of measurement properties and correlation with injury. Part 2-the tests for the hip, thigh, foot and ankle including the star excursion balance test. British journal of sports medicine. 2015;49(10).
- 198. Hegedus EJ, McDonough S, Bleakley C, Cook CE, Baxter GD. Clinician-friendly lower extremity physical performance measures in athletes: a systematic review of measurement properties and correlation with injury, part 1. The tests for knee function including the hop tests. British journal of sports medicine. 2015;49(10):642-8.
- 199. Negahban H, Mazaheri M, Kingma I, van Dieen JH. A systematic review of postural control during single-leg stance in patients with untreated anterior cruciate ligament injury. Knee surgery, sports traumatology, arthroscopy: official journal of the ESSKA. 2014;22(7):1491-504.
- 200. Dingenen B, Janssens L, Claes S, Bellemans J, Staes FF. Lower extremity muscle activation onset times during the transition from double-leg stance to single-leg stance in anterior cruciate ligament reconstructed subjects. Clinical biomechanics (Bristol, Avon). 2016;35:116-23.
- 201. Gokeler A, Welling W, Benjaminse A, Lemmink K, Seil R, Zaffagnini S. A critical analysis of limb symmetry indices of hop tests in athletes after anterior cruciate ligament reconstruction: A case control study. Orthopaedics & traumatology, surgery & research: OTSR. 2017;103(6):947-51.
- 202. Pietrosimone B, Lepley AS, Harkey MS, Luc-Harkey BA, Blackburn JT, Gribble PA, et al. Quadriceps Strength Predicts Self-reported Function Post-ACL Reconstruction. Medicine and science in sports and exercise. 2016;48(9):1671-7.