What are validated self-report adherence scales really measuring?: a systematic review

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AIMS

Medication non-adherence is a significant health problem. There are numerous methods for measuring adherence, but no single method performs well on all criteria. The purpose of this systematic review is to (i) identify self-report medication adherence scales that have been correlated with comparison measures of medication-taking behaviour, (ii) assess how these scales measure adherence and (iii) explore how these adherence scales have been validated.

METHODS

Cinahl and PubMed databases were used to search articles written in English on the development or validation of medication adherence scales dating to August 2012. The search terms used were *medication adherence*, *medication non-adherence*, *medication compliance* and names of each scale. Data such as barriers identified and validation comparison measures were extracted and compared.

RESULTS

Sixty articles were included in the review, which consisted of 43 adherence scales. Adherence scales include items that either elicit information regarding the patient's medication-taking behaviour and/or attempts to identify barriers to good medication-taking behaviour or beliefs associated with adherence. The validation strategies employed depended on whether the focus of the scale was to measure medication-taking behaviour or identify barriers or beliefs.

CONCLUSIONS

Supporting patients to be adherent requires information on their medication-taking behaviour, barriers to adherence and beliefs about medicines. Adherence scales have the potential to explore these aspects of adherence, but currently there has been a greater focus on measuring medication-taking behaviour. Selecting the 'right' adherence scale(s) requires consideration of what needs to be measured and how (and in whom) the scale has been validated.

Introduction

There are many effective medicines available to treat illness, but the benefits of these medicines will only accrue to the patients that take them. The World Health Organization [1] defines adherence as:

The extent to which a person's behaviour – taking medication, following a diet and/or executing lifestyle changes, corresponds with agreed recommendations from a health care provider.

Medication non-adherence is common, with studies in a range of settings identifying up to 50% of patients as non-adherent to a medicine [2–6]. Poor medication adherence results in adverse health outcomes [7–9] and increased health care costs [7].

Patients may be non-adherent due to different beliefs, barriers and a range of other factors. Patients may intentionally decide not to take their medicines based on well-informed or mistaken beliefs about the benefits and risks of their medicines [10, 11]. Patients can unintentionally non-adhere to medicines due to forgetfulness, careless-

ness, health literacy and socioeconomic factors. Non-adherence can also occur at different stages of the medication-taking process. A patient may exhibit non-adherence at the *initiation* of treatment, during treatment (where the patient may exhibit sub-optimal *implementation* of the treatment regimen) or the patient may *discontinue* the treatment early [12]. Strong evidence for any single approach to improve medication adherence is lacking, but interventions that are tailored to a patient's specific reasons and stage of non-adherence can be expected to better support good medication-taking behaviour [13–17].

Adherence to medicines is measured for different purposes. Common reasons to measure adherence include better informing the assessment of an intervention (as unrecognized non-adherence may lead to an underestimation of possible treatment effects), determining influences on adherence to medicines in people with specific disease states (such as hypertension or HIV) and identifying patients requiring education or support to improve medication use. Ideally, clinicians and researchers wanting a comprehensive assessment of adherence need measures that are inexpensive, relatively easy to administer, accurately identify the patient's current medication-taking behaviour and any barriers or beliefs that may influence the patient's use of medicines.

There are a number of ways of measuring adherence. Objective measures, including measurement of clinical outcomes, dose counts, pharmacy records, electronic monitoring of medication administration (e.g. the Medication Event Monitoring System, MEMS) and drug concentrations [18-21], seemingly provide the best measure of a patient's medication-taking behaviour in many contexts [22-27]. It is important to recognize that, while objective, most of these measures have drawbacks. MEMS, arguably the best objective measure of medication-taking behaviour, records package opening or device actuation, rather than actual medication-taking and the possibility of intentional dose dumping remains. MEMS, or MEMS-like devices, are also expensive and not readily available for some dose forms [21, 28-30]. While clinical outcomes are the ultimate aim of any intervention to improve adherence, the use of clinical outcomes as a proxy of adherence can be confounded by disease-specific factors independent of medication-taking behaviour.

Subjective measures of adherence include physician or family reports, patient interviews and self-report adherence scales [10, 31–34]. These measures have the potential to identify the specific reasons for a patient's non-adherence. Subjective measures can be relatively simple to use and are less expensive. However, they are prone to recall bias and the prospect that respondents provide answers that conform to their perceived expectations of their interviewer [35, 36]. There are a large number of adherence scales that are suitable for use in research or clinical settings. A number of well-validated adherence

scales have been strongly correlated with objective measures of adherence in several different populations of patients.

There is a need for scales that are easy to administer and correctly identify medication-taking behaviour, key barriers to adherence and beliefs associated with medication use that influence adherence. There have been few systematic attempts to describe the available self-report adherence scales and their benefits and limitations with respect to both medication-taking behaviour and the identification of barriers and beliefs associated with adherence [37, 38]. The aim of this review is to (i) identify self-report medication adherence scales that have been correlated with a comparison measure of medication-taking behaviour, (ii) assess how these scales measure adherence and (iii) explore how these adherence scales have been validated.

Methods

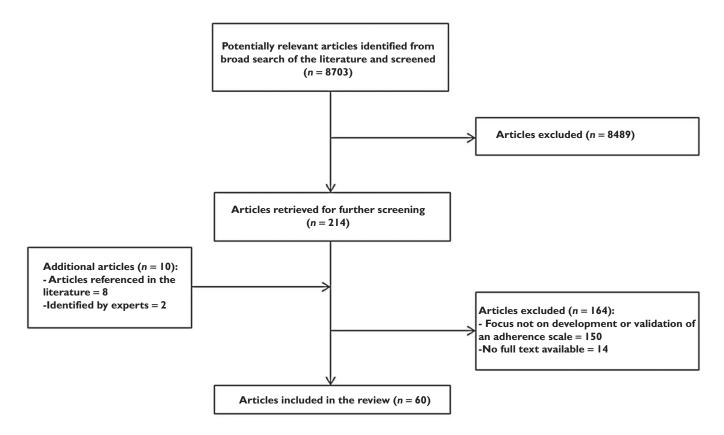
A literature search for adherence scales was conducted using Cinahl and PubMed electronic databases. The initial search terms used to identify the articles were: medication adherence, medication non-adherence, medication compliance and medication non-compliance. This broad database search identified the names of the self-report adherence scales, which were then searched individually. This search was limited to English language studies published between 1981 and 2012. The date of the last search was on 1 August 2012. The reference lists of the relevant studies were searched to identify additional articles.

Inclusion and exclusion criteria

Adherence scales were included if they had been correlated against a comparison measure (objective or subjective) of medication-taking behaviour. To be included there needed to be a full text article, written in the English language on the development and/or validation of the adherence scale. Studies that used the self-report adherence scale without correlating the adherence scale against a comparison measure of medication-taking behaviour were excluded. The list of scales was reviewed for completeness with two adherence researchers.

Data extraction and analysis

The data extracted from the studies included the number of items in the adherence scale, study setting, criteria for identifying non-adherence, response rate and time to complete the adherence scale. Each validated self-report adherence scale was categorized according to whether it contained items that elicited information on (i) specific medication-taking behaviours: dose taken, dose frequency, dose administration and prescription refills, (ii) barriers to adherence: e.g. forgetfulness, treatment



Flow chart of study selection process

complexity and side effects and/or (iii) beliefs associated with adherence: e.g. perceived necessity of medicines and concerns about medicines. Adherence scales were also assessed on whether or not the scale identified the initiation, implementation or discontinuation of treatment as per the taxonomy proposed by Vrijens *et al.* [12].

To assess the quality of the correlation study, the following criteria were extracted: how the adherence scale was administered, sample size, the adherence comparison measures, internal consistency and, where reported, the sensitivity and specificity of the scale against a standard of adherence. Information on criterion, content and construct validity was also extracted to assess the validation of the adherence scale. The results of the studies were reviewed and compared.

Results

Search strategy

The study selection process is illustrated in Figure 1. Twenty-one articles were retrieved using the Cinahl search engine and the remaining were identified using the same strategy in the PubMed database and from reference lists. Some adherence scales were excluded, as shown in Table 1.

Table 1Excluded self-report adherence scales

Excluded adherence scale	Reason for exclusion
Adult AIDS Clinical Trials Group (AACTG) Adherence Scale [90]	No validation studies were found in the literature search
Basel Assessment of Adherence Scale (BAAS) [91]	No validation studies were found in literature search.
Medication Adherence Evaluation Scale (MASS) [92]	No full-text article available
Medication Adherence Measure (MAM) interview [32]	Semi-structured interview and thus was not consistent between patients
Multicentre Aids Cohort Study (MACS) adherence form [93]	No adherence comparison measure

The literature search retrieved 60 articles that met the inclusion criteria (Figure 1). The sample size of the studies ranged from 40 to 1367 (Table 2) [39, 40]. The median sample size of the studies was 228. Twenty-two of the studies reported the response rate, ranging from 29 to 98%. The average response rate was 72% (Table 3). Forty-three self-report adherence scales were identified from the included studies.

Table 2

Comparison of the self-report adherence scales

Self-report adherence scale	Based on	Number of questions	Time to complete	Barriers identified	Stage of medication-taking identified	development and correlation studies
Group 1: Medication-taking behaviour						
Adherence Self-Report Questionnaire (ASRQ) [41]	CQR	_	Not reported	- Nil	Implementation	2
Adherence Visual Analogue Scale (VAS) Computerized [52]	Literature review	_	Not reported	- Nil	Implementation	_
Brief Adherence Rating Scale (BARS) [33]	CATIE trial	4	Not reported	- Nil	Implementation	_
Barroso et al. 30-day Adherence Question [53]	Did not specify	_	Not reported	- Nil	Implementation	_
Bell et al. Adherence Question [42]	Did not specify	_	Not reported	- Nil	Implementation	_
Centre for Adherence Support Evaluation (CASE) Adherence	Literature review	m	Not reported	- Nil	Implementation	_
index [34] Gehi <i>et al.</i> Adherence Question [9]	Expertise	-	Not reported	- Ni	Implementation	-
Grymonpre et al. Adherence Question [43]	Did not specify	_	Not reported	≡.Z ·	Implementation	_
Kerr et al. Adherence Question [44]	Expertise	-	Not reported	- Nil	Implementation	-
Medication Adherence Report Scale – 5 (MARS-5) [71, 85]	MARS-10	2	Not reported	- Nil	Implementation	m
					Discontinuation	
Stages of Change for Adherence (SOCA) [55]	Stages of Change model	2	Not reported	 Stages of change in regards to medication adherence 	Initiation Implementation	-
Group 2: Medication-taking behaviour and barriers						
Adherence to Refills and Medications Scale (ARMS) [51]	Literature review, MAQ and	12	Not reported	- Correct administration	Implementation	_
	Hill-Bone Compliance Scale			- Forgetfulness	Discontinuation	
				- Prescription refill ability		
Adherence Starts with Knowledge-12 (ASK-12) [45]	ASK-20	12	Not reported	 Patient-perceived barriers 	Implementation	_
				- Inconvenience	Discontinuation	
				- Forgetfulness		
Adherence Starts with Knowledge-20 (ASK-20) [46]	literature review nationt focus	20	Not reported	- Medication-taking behaviour	Implementation	2
	groups and expert panel input) I		- Patient-perceived barriers	Discontinuation	ı
Brief Medication Questionnaire [40]	Literature review and patient	6	Not reported	- Evaluates regimen	Implementation	_
	teedback			- Medication-taking problems	-	(
brooks Medication Adnerence Scale (bMAS) [30]	MAQ	4	Less than 5 min	- Forgettuiness - Carelessness	Implementation Discontinuation	7
				- Adverse effects and efficacy		
Choo et al. 5-item Questionnaire [57]	Brief Medication Questionnaire	5	Not reported	- Unintentional and intentional	Implementation	_
				non-adherence		
Fodor <i>et al.</i> Adherence Questionnaire [72]	Did not specify	6	Not reported	- Blood pressure awareness	Implementation	_
Godin et al. Self-Reported Adherence Ouestionnaire [47]	Expertize	9	5 min	- Barriers to adherence	Implementation	-
Hill-Bone Compliance Scale – 10 [8]	The Hill-Bone Compliance	10	Not reported	- Medication adherence	Implementation	_
	Scale – 14			- Appointment-making		
				- Sodium intake		



National	Hill-Bone Compliance Scale – 14 [62]	Literature review and Clinical	14	5 min	- Medication adherence	Implementation	_
Adherence Scale (TAS) [89] MAQ 4 Less than 5 min - Foogetuines Implementation ament Tool (MAAX) [80] Litreature review and Expert 12 Not reported - Foogetuines Implementation c Scale (MMAS) [34] MAQ and behavioural aspects 8 Not reported - Advance effects and problems Implementation A Medication Adherence MAMAS 8 Not reported - Foogetuines Implementation A Medication Adherence MAMAS 8 Not reported - Foogetuines Implementation A Litreature review, expertise and recision are foods and problems 1-3 min - Medication-taking behaviour Discontinuation ALONDS (RAM) Countinuation (RAM) Scale [10] Litreature review, expertise and 66 1-3 min - Medication-taking behaviour Discontinuation ALONDS (RAM) Countinuation (RAM) Scale [10] Litreature review, expertise 6 1-3 min - Medication-taking problems Implementation ALONDS (RAM) Countinuation (RAM) (RA		expertise			- Appointment-making	-	
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MAQ and behavioural aspects Not reported - Access to medications Implementation - Access to medications Implementation Discontinuation Disco		clinicians			- Adverse effects		
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Health Belief Model, Health 28 Not reported - Cognitive functioning Promotion Model, Promotion Model, Reasoned Action Promotion Model, Reasoned Action Seasoned Action Seaso	[60]				- Medication-taking strategy		
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Reasoned Action - Self-efficacy - Commitment Comm		Promotion Model,			- Patient-Provider		
- Commitment - Forgetfulness and carelessness et al. 164] Patient interviews 26 Not reported - Forgetfulness and efficacy Discontinuation - Adverse effects and efficacy Discontinuation - Adverse effects and efficacy Discontinuation - Adverse effects - Adverse effects - Lifestyle barriers - MASES 13 Not reported - Lifestyle barriers - Adverse effects		Reasoned Action			- Self-efficacy		
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MASES 13 Not reported - Medication self-efficacy - Lifestyle barriers - Adverse effects					- Lifestyle barriers		
	Medication Adherence Self-Efficacy Scale Revised	MASES	13	Not reported	- Medication self-efficacy	Implementation	-
- Adverse effects	(MASES-R) [79]				- Lifestyle barriers		
					- Adverse effects		

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Self-report adherence scale	Based on	Number of questions	Time to complete	Barriers identified	Stage of medication-taking identified	Number of development and correlation studies
Medication Adherence Reasons Scale [66]	Literature review	75	Not reported	 Managing issues Beliefs Multiple medication issues Availability issues Forgetfulness 	Implementation Discontinuation	-
The Self-Efficacy for Appropriate Medication Use Scale (SEAMS) [65]	Literature, expertise and patient interviews	13	Not reported	- Specific problem areas - Self-efficacy	Ē	-
Group 4: Beliefs associated with adherence Beliefs about Medicines Questionnaire [10]	Health Belief Model and Patient Beliefs	18	Not reported	- Medication necessity beliefs - Medication concerns	Nil	4
Drug Attitude Inventory (DAI) [4]	Literature review and patient reports	30	Not reported	- Attitudes towards medications - Beliefs on medications	Discontinuation	-
Group 5: Barriers and beliefs						
Beliefs and Behaviour Questionnaire (BBQ) [67]	Qualitative interviews with Chronic Obstructive Pulmonary Disease (COPD) patients	30	Not reported	- Beliefs - Experiences	Ē	-
Brief Evaluation of Medication Influences and Beliefs (BEMIB) [68]	Health Belief Model and Patient/Investigator feedback	00	Less than 5 min	- Forgetfulness - Access to medications - Support network - Benefits of medication	Implementation	-
Compliance Questionnaire Rheumatology (CQR) [75]	Patient interviews	19	Approximately 12 min	- Forgetfulness - Value of doctor instructions	Implementation	2
Maastricht Utrecht Adherence in Hypertension (MUAH) Questionnaire [69]	Patient interviews	25	Average 25 min	 Medication attitude Discipline and aversions Coping with health problems 	Implementation	-
Medication Adherence Report Scale (MARS) [70]	MAQ and DAI	10	Not reported	- Forgetfulness - Adverse effects - Value of medication - Behaviour and attitudes	Implementation Discontinuation	2



Self-report adherence scales – administration, response and validation data

Self-report adherence scale	Reference number	How scale was administered	Response rate	Internal consistency	Sensitivity	Specificity	Correlation (criterion validity)	Comparison measure of adherence	Sample size
Group 1: Medication-taking behaviour	aviour								
Adherence Self-Report Questionnaire (ASRQ)	[41]	Clinician-administered (Other HCP)	Not reported	Not reported	Not reported	Not reported	Significant	Electronic monitoring (MEMS)	245
	[94]	Clinician-administered (Primary physician)	53%	Not reported	Not reported	Not reported	Not significant (all)	Electronic monitoring (MEMS) and Other scale (VAS)	78
Adherence Visual Analogue Scale (VAS) Computerized	[52]	Self-administered (Home, Internet)	Not reported	Not reported	Not reported	Not reported	Significant (all)	Dose counts and Clinical outcome (Viral load)	290
Brief Adherence Rating Scale (BARS)	[33]	Researcher-administered (Clinic)	Not reported	$\alpha = 0.92$	0.73	0.74	Significant	Electronic monitoring (MEMS)	61
Barroso <i>et al.</i> 30-day Adherence Question	[53]	Researcher-administered (Clinic)	Not reported	Not reported	Not reported	Not reported	Significant	Clinical outcome (Viral load)	93
Bell <i>et al.</i> Adherence Question	[42]	Self-administered (Clinic, Absent)	%08	Not reported	Not reported	Not reported	Not significant (all)	Electronic monitoring (MEMS) and Dose count	08
Centre for Adherence Support Evaluation (CASE) Adherence Index	[54]	Clinician-administered (Primary physician)	Not reported	Not reported	0.70	0.71	Significant (all)	Clinical outcome (Viral load) and Self-report	524
Gehi <i>et al.</i> Adherence Question	[6]	Researcher-administered (Clinic)	Not reported	Not reported	Not reported	Not reported	Significant	Clinical outcome (Cardiovascular events)	1015
Grymonpre <i>et al.</i> Adherence Question	[43]	Researcher-administered (Home)	Not reported	Not reported	Not reported	Not reported	Significant (all)	Pharmacy records and Dose counts	135
Kerr <i>et al.</i> Adherence Question	[44]	Researcher-administered (Clinic)	Not reported	Not reported	Not reported	Not reported	Not significant	Pharmacy records	80
Medication Adherence Report Scale – 5 (MARS-5)	[2]	Clinician-administered (Primary physician)	Not reported	Not reported	0.085	0.97	Not significant	Pharmacy records	128
[82]	[71]	Self-administered (Clinic, Present)	Not reported	$\alpha = 0.78$	Not reported	Not reported	Significant (all)	Drug levels and Self-report	255
Stages of Change for Adherence (SOCA)	[55]	Self-administered (Clinic, Absent)	Not reported	Not reported	Not reported	Not reported	Significant (all)	Electronic monitoring (MEMS) and Other scale (BMAS)	731
Group 2: Medication-taking behaviour and barriers Adherence to Refills and [51] Rese Medications Scale (ARMS)	aviour and ba [51]	uriers Researcher-administered (Clinic)	%68	$\alpha = 0.81$	Not reported	Not reported	Significant (all)	Clinical outcome (BP control), Pharmacy records and Other	435
Adherence Starts with	[45]	Self-administered (Clinic, Absent)	Not reported	$\alpha = 0.75$	Not reported	Not reported	Significant (all)	Scale (MAC) Pharmacy record and Other scale	112
Adherence Starts with Knowledge-20 (ASK-20)	[46] [95]	Self-administered (Home, Internet) Self-administered (Clinic, Present)	Not reported Not reported	$\alpha = 0.85$ $\alpha = 0.76$	Not reported Not reported	Not reported Not reported	Significant Significant	Self-report Pharmacy records and Other scale	605
Brief Medication	[40]	Researcher-administered (Clinic)	95%	Not reported	0.80	1.00	(Scale only) Significant	(MAQ) Electronic monitoring (MEMS)	43
Questionnaire Brooks Medication	[56]	Self-administered (Unclear)	%98	$\alpha = 0.69$	Not reported	Not reported	1	Nil	294
Adherence Scale (BMAS)	[96]	Self-administered (Home)	63%	$\alpha = 0.86$	Not reported	Not reported	Significant (all)	Pharmacy records and Self-report	86
Choo et al. 5-item	[22]	Researcher-administered	Not reported	Not reported	Not reported	Not reported	Significant	Electronic monitoring (MEMS)	286
Agestionianie									

172 Researcher-administered (Clinic, Absem) Not reported Significant Clinic Self-administered (Clinic, Absem) Not reported Clinic, Absem) Not reported Not reported Not reported Not reported Significant Clinic Self-administered (Clinic, Absem) Not reported Not reported Not reported Significant Self-administered (Clinic, Absem) Not reported Not reported Not reported Significant Self-administered (Clinic, Absem) Not reported Not reported Not reported Significant Self-administered (Clinic, Absem) Not reported Not reported Not reported Significant Self-administered (Clinic, Absem) Not reported Not reported Not reported Significant Self-administered (Clinic, Absem) Not reported Not reported Not reported Significant Self-administered (Clinic, Absem) Not reported Not reported Not reported Not reported Significant Self-administered (Clinic, Absem) Not reported Not reported Not reported Not reported Not reported Significant Significant Not reported		Reference	How scale was		Internal			Correlation	Comparison measure of	
(27] Researcher-administered (Clinic, Absent) Not reported Not reported O.71 Or. Spriftcant Significant [8] Researcher-administered (Clinic, Absent) Not reported 0.71 0.72 Not significant [62] Researcher and self-administered (Clinic) Not reported α = 0.84 Not reported Not reported Significant [63] Self-administered (Clinic) 29% α = 0.84 Not reported Not reported Significant [48] Self-administered (Clinic) 29% α = 0.84 Not reported Not reported Not reported Not reported Not reported Significant [49] Self-administered (Clinic, Absent) Not reported 0.53 0.91 Significant [40] Self-administered (Clinic, Absent) Not reported α = 0.82 Not reported Significant [50] Self-administered (Clinic, Absent) Not reported 0.53 0.91 Significant [50] Self-administered (Clinic, Absent) Not reported Not reported Not reported Not reported Sign	Self-report adherence scale	number	administered	Response rate	consistency	Sensitivity	Specificity	(criterion validity)	adherence	Sample
Researcher-administered (Clinic, Absent) Not reported Not reported 0.71 0.72 Not significant (Clinic) Researcher-administered (Clinic, Absent) Not reported 0.71 0.72 Not significant (Clinic) Self-administered (Clinic, Absent) Not reported Not reported Not reported Significant (Clinic) Self-administered (Clinic, Absent) Not reported Not reported Not reported Significant (Significant (Clinic, Absent) Not reported Not reported Not reported Significant (Significant (Signific	Fodor <i>et al.</i> Adherence	[72]	Researcher-administered (Clinic)	Not reported	Not reported	Not reported	Not reported	Significant	Clinical outcome (BP control)	359
8 Researcher and self-administered Not reported Cinici) Cinici) Cinici) Self-administered (Finici) 29% Cinici) Cinici) Self-administered (Finici) 29% Cinici) Cinician-administered (Finician-administered (Finician-administere	Questionnaire Godin <i>et al.</i> Self-Reported	[47]	Self-administered (Clinic, Absent)	Not reported	Not reported	0.71	0.72	Not significant	Clinical outcome (Viral	256
[62] Researcher and self-administered (Clinic) 29% $\alpha = 0.84$ Not reported Not reported Significant (all) Self-administered (Home) 91% $\alpha = 0.81$ Not reported Not reported Significant (all) Self-administered (Clinic) 29% $\alpha = 0.83$ 0.93 0.93 Self-administered (Clinic, Abzent) Not reported Not reported Not reported Significant Self-administered (Clinic, Abzent) Not reported Not reported Not reported Significant Not reported Not reported Not reported Significant Not reported Not reported Not reported Not reported Significant Not reported Not reported Not reported Not reported Significant Not reported	Adnerence Questionnaire Hill-Bone Compliance Scale –	[8]	Researcher- and self-administered	Not reported	$\alpha = 0.79$	Not reported	Not reported	Significant	suppression) Clinical outcome (BP control)	86
Self-administered (Home) 91% α = 0.81 Not reported Not reported Significant (all) Self-administered (Clinic) 29% α = 0.83 0.93 0.53 Significant (all) Self-administered (Linic, Absent) Not reported Not reported Not reported Significant (all) Self-administered (Clinic, Absent) Not reported Not reported Not reported Significant Self-administered (Clinic, Absent) Not reported Not reported Not reported Significant Self-administered (Clinic, Absent) Not reported α = 0.72 Not reported Significant Self-administered (Clinic, Absent) Not reported α = 0.72 Not reported Not reported Significant (all) Self-administered (Clinic, Absent) Not reported Not reported Not reported Significant (all) Self-administered (Clinic, Absent) Not reported Not reported Not reported Significant (all) Self-administered (Clinic, Absent) Not reported Not reported Not reported Significant (all) Self-administered (Clinic, Absent) Not reported Not reported Not reported Significant (all) Self-administered (Clinic, Absent) Not reported Not reported Not reported Significant (all) Self-administered (Clinic, Absent) Not reported α = 0.75 Not reported Not reported Significant (all) Self-administered (Clinic, Absent) Not reported α = 0.32 Not reported Not reported Significant Significant (all) Self-administered (Clinic) Not reported Not reported Not reported Significant Significant Significant Significant Significant Significant Significant Significant Significant Significant Significant Significant Significant Sig	Hill-Bone Compliance Scale – 14	[62]	(Clinic) (Clinic)	Not reported	$\alpha = 0.84$	Not reported	Not reported	Significant	Clinical outcome (BP control)	718
(39] Self-administered (Clinic) 29% α = 0.83 0.93 0.53 Significant (30] Self-administered (Clinic) 98% α = 0.83 0.93 0.53 Significant (30] Self-administered (Unclea) 82% α = 0.82 Not reported Not reported Significant (48) Self-administered (Hone) 39% α = 0.56 Not reported Not reported Significant (10) Self-administered (Clinic, Absent) Not reported 0.63 0.91 Significant (10) Self-administered (Clinic, Absent) Not reported Not reported Not reported Significant (59) Researcher-administered (Hone) Not reported Not reported Not reported Significant (59) Self-administered (Hone) Not reported Not reported Not reported Significant (50) Self-administered (Clinic, Absent) Not reported Not reported Not reported Not reported Significant (50) Self-administered (Clinic, Absent) Not reported Not repo	Immunosuppressant Therapy Adherence Scale (ITAS)	[58]	Self-administered (Home)	91%	$\alpha = 0.81$	Not reported	Not reported	Significant (all)	Pharmacy records, Drug levels and Clinical outcome (Rejection)	222
34] Researcher-administered (Clinic, Absent) 39% $\alpha = 0.83$ 0.93 0.53 Significant 120] Self-administered (Clinic, Absent) Not reported Significant (all) Physician) Not reported Not reported Not reported Not reported Significant (all) Not reported Not reported Not reported Not reported Not reported Not reported Significant (all) Not reported	Medication Adherence Assessment Tool (MAAT)	[80]	Self-administered (Clinic)	29%	$\alpha = 0.80$	Not reported	Not reported	Significant	Pharmacy records	289
Self-administered (Unidear) 66% Not reported Not reported Not reported Significant Assarcher-administered (Clinic, Absent) Not reported Not reported Not reported Significant Self-administered (Clinic, Absent) Not reported Not reported Not reported Not reported Significant Not reported Not reported Not reported Significant Not reported Not reported Not reported Significant (31) physician) Not reported Not reported Not reported Not reported Not reported Not reported Significant (31) physician) Not reported Not reported Not reported Not reported Not reported Significant (31) physician) Not reported Not reported Not reported Not reported Significant (31) physician) Not reported Not reported Not reported Significant (31) Not reported Not reported Not reported Significant (31) Not reported Not reported Not reported Significant (31) Not reported Not reported Significant (31) Not reported Not reported Not reported Significant Not reported Not reported Significant Not reported Not reported Not reported Significant (31) Not reported Not reported Not reported Significant Dhysician) Not reported Not reported Not reported Not reported Significant (31) Not reported Not reported Not reported Not reported Significant Not reported Not reported Not reported Significant (31) Not reported N	Morisky Medication	[34]	Researcher-administered (Clinic)	%86	$\alpha = 0.83$	0.93	0.53	Significant	Clinical outcome (BP control)	1367
Self-administered (Home) 39% $\alpha = 0.82$ Not reported Not reported Significant Self-administered (Clinic, Absent) Not reported Not reported Not reported Not reported Significant (Self-administered (Clinic, Absent) Not reported Not reported Not reported Not reported Significant (all) Physician) [59] Researcher-administered (Primary Not reported Not reported Not reported Significant (all) physician) [60] Self-administered (Clinic, Absent) Not reported $\alpha = 0.75$ 0.72 0.91 Significant (all) physician) [61] Researcher-administered (Clinic, Absent) Not reported $\alpha = 0.90$ Not reported Not reported Significant (all) Not reported (Clinic) Not reported $\alpha = 0.90$ Not reported Not reported Significant (all) Not reported (Clinic) Not reported Not reported Not reported Significant (all) Not reported (Clinic) Not reported Not reported Not reported Significant (all) Not reported (Clinic) Not reported Not reported Not reported Significant (all) Not reported (Clinic) Not reported Not reported Significant (all)	Adherence Scale (MMAS)	[20] [97]	Self-administered (Unclear) Researcher-administered (Clinic)	66% 82%	Not reported $\alpha = 0.56$	Not reported 0.73	Not reported 0.36	Significant Significant	Pharmacy records Pharmacy records	87
19 Self-administered (Clinic, Absent) Not reported Not reported 0.63 0.91 Significant 10 Self-administered (Clinic, Absent) Not reported Not reported Not reported Not reported Significant (all) 19 Researcher-administered (Home) Not reported Not reported Not reported Significant (all) 19 Clinician-administered (Primary Not reported $\alpha = 0.75$ 0.72 0.91 Significant (all) 19 Self-administered (Home) 70% Not reported Not reported Not reported Significant (all) 10 Researcher-administered (Clinic) Not reported $\alpha = 0.90$ Not reported Not reported Significant 10 Researcher-administered (Clinic) Not reported Not reported Not reported Significant 11 Researcher-administered (Clinic) Not reported Not reported Not reported Significant 12 Clinician-administered (Clinic) Not reported Not reported Not reported Significant 12 Researcher-administered (Clinic) Not reported Not reported Not reported Significant 12 Researcher-administered (Clinic) Not reported Not reported Not reported Significant 13 Researcher-administered (Clinic) Not reported Not reported Not reported Significant 14 Significant 15 Self-administered (Clinic) Not reported Not reported Significant 16 Self-administered (Clinic) Not reported Not reported Significant 17 Self-administered (Clinic) Not reported Not reported Not reported Significant 18 Self-administered (Clinic) Not reported Not reported Significant	Osteoporosis-Specific Morisky Medication Adherence Scale (OS-MMAS)	[48]	Self-administered (Home)	%68	$\alpha = 0.82$	Not reported	Not reported	Significant	Other scale (Beliefs about Medicines Questionnaire)	197
10 Self-administered (Clinic, Absent) Not reported $\alpha = 0.72$ Not reported Significant (all) 139 Researcher-administered (Primary Not reported Not reported Not reported Not reported Significant (all) 139 Clinician-administered (Primary Not reported $\alpha = 0.75$ 0.72 0.91 Significant (all) 130 Self-administered (Clinic, Absent) Not reported $\alpha = 0.90$ Not reported Not	Pediatric Inhaler Adherence Questionnaire (PIAQ)	[49]	Self-administered (Clinic, Absent)	Not reported	Not reported	0.63	0.91	Significant	Dose counts	64
[59] Researcher-administered (Clinic, Absent) Not reported α = 0.75 0.72 0.91 Significant (all) [60] Self-administered (Home) 70% Not reported Not reported Not reported Not reported Not reported Not reported Significant (all) [63] Self-administered (Clinic, Absent) Not reported α = 0.61 Not reported Not reported Not reported Not reported Not reported Significant principant [73] Researcher-administered (Clinic) 73% α = 0.61 0.32 Not reported Not repor	Reported Adherence to Medicine (RAM) Scale	[10] [50]	Self-administered (Clinic, Absent) Self-administered (Home)	Not reported Not reported	$\alpha = 0.72$ Not reported	Not reported Not reported	Not reported Not reported	– Significant	Nil Other scale (Beliefs about Madicines Ouestionnaire)	524 271
[59] Clinician-administered (Primary Not reported $\alpha = 0.75$ 0.72 0.91 Significant (all) physician) [60] Self-administered (Home) 70% Not reported $\alpha = 0.90$ Not reported Not reported Significant (all) [63] Self-administered (Clinic, Absent) Not reported $\alpha = 0.90$ Not reported Not reported Significant [78] Researcher-administered (Clinic) Not reported $\alpha = 0.61$ 0.81 0.44 Significant physician) [73] Researcher-administered (Clinic) Not reported $\alpha = 0.32$ Not reported Significant physician) [74] Researcher-administered (Clinic) Not reported Not reported Significant Significant (All) Researcher-administered (Clinic) Not reported Not reported Not reported Significant Significant Significant Significant Significant Not reported Not reported Significant Signific	Self-Reported Adherence (SERAD) Questionnaire	[65]	Researcher-administered (Clinic)	Not reported	Not reported	Not reported	Not reported	Significant (all)	Electronic monitoring (MEMS), Dose counts and Drug levels	530
[60] Self-administered (Home) 70% Not reported Not reported Not reported Significant (all) Ph [63] Self-administered (Clinic, Absent) Not reported $\alpha = 0.90$ Not reported Not reported Not reported Not significant (all) Ph [73] Researcher-administered (Clinic) Not reported $\alpha = 0.61$ 0.81 0.44 Significant Ph [74] Researcher-administered (Primary Not reported Not reported 0.32 0.73 Not significant Ph physician) Not reported Not reported Not reported Significant Ph [77] Researcher-administered (Clinic) Not reported Not reported Not reported Significant (all) Ele	Simplified Medication Adherence Questionnaire (SMAQ)	[39]	Clinician-administered (Primary physician)	Not reported	$\alpha = 0.75$	0.72	0.91	Significant (all)	Electronic monitoring (MEMS) and Clinical outcome (Viral load)	40
[63] Self-administered (Clinic, Absent) Not reported α = 0.90 Not reported Not significant Se [73] Researcher-administered (Clinic) 73% α = 0.61 0.81 0.44 Significant Ph [76] Researcher-administered (Primary Not reported Not reported 0.32 0.73 Not significant Ph physician) [77] Researcher-administered (Clinic) Not reported Significant (all) Ele	The Patterns of Asthma Medication Use Questionnaire	[60]	Self-administered (Home)	%02	Not reported	Not reported	Not reported	Significant (all)	Pharmacy records and Clinical outcome (Asthma)	176
(73) Researcher-administered (Clinic) 73% α = 0.61 0.81 0.44 Significant Clinician-administered (Primary physician) (76) Researcher-administered (Primary physician) Not reported α = 0.32 Not reported Not reported 0.73 Not significant Physician physician (77) Researcher-administered (Clinic) Not reported Not reported Not reported Not reported Not reported Significant (all) Ele	Group 3: Barriers to adherence Adherence Attitude Inventory (AAI)	[63]	Self-administered (Clinic, Absent)	Not reported	$\alpha = 0.90$	Not reported	Not reported	Not significant	Self-report	165
Clinician-administered (Primary Not reported Not reported 0.32 0.73 Not significant Phphysician) Researcher-administered (Clinic) Not reported Not reported Not reported Significant (all) Ele	Medication Adherence Questionnaire (MAQ)	[73]	Researcher-administered (Clinic) Researcher-administered (Clinic)	73% Not reported	$\alpha = 0.61$ $\alpha = 0.32$	0.81 Not reported	0.44 Not reported	Significant Significant	Clinical outcome (BP control) Pharmacy records	290
Researcher-administered (Clinic) Not reported Not reported Not reported Not reported Significant (all) Ele		[5]	Clinician-administered (Primary physician)	Not reported	Not reported	0.32	0.73	Not significant	Pharmacy records	128
		[77]	Researcher-administered (Clinic)	Not reported	Not reported	Not reported	Not reported	Significant (all)	Electronic monitoring (MEMS), Dose counts and Drug levels	385

	[86]	Researcher-administered (Clinic)	32%	Not reported	Not reported	Not reported	Significant	Clinical outcome (Diabetes control)	186
	[66]	Researcher-administered (Home)	%88	$\alpha = 0.42$	Not reported	Not reported	Not significant	Dose counts	319
	[78]	Researcher-administered (Clinic)	Not reported	Not reported	Not reported	Not reported	Significant	Dose counts	413
	[100]	Researcher-administered (Clinic)	81%	$\alpha = 0.62$	Not reported	Not reported	Significant	Clinical outcome (Diabetes	294
Medication Adherence Self-Efficacy Scale (MASES)	[64]	Self-administered (Unclear)	Not reported	$\alpha = 0.95$	Not reported	Not reported	Not significant	Clinical outcome (BP control)	72
Medication Adherence Self-Efficacy Scale Revised (MASES-R)	[62]	Self-administered (Clinic, Present)	Not reported	$\alpha = 0.91$	Not reported	Not reported	Significant (all)	Electronic monitoring (MEMS) and Other scale (MAQ)	168
Medication Adherence Reasons Scale	[99]	Self-administered (Home, Internet)	Not reported	$\alpha = 0.65$	Not reported	Not reported	Significant	Other scale (MAQ)	840
Self-Efficacy for Appropriate Medication Use Scale (SEAMS)	[65]	Self-administered (Clinic)	Not reported	$\alpha = 0.89$	Not reported	Not reported	Significant	Other scale (MAQ)	436
Group 4: Beliefs associated with adherence	adherence			0			9	A 4 4 4 7 - 1 - 1 - 1 - 1 - 1 - 1 - 1 - 1 - 1 -	
Beliefs about Medicines Ouestionnaire	[01]	self-administered (Clinic, Absent) Self-administered (Clinic, Absent)	Not reported 57%	$\alpha = 0.70$ Not reported	Not reported Not reported	Not reported Not reported	Significant	Other scale (MARS-5)	324
•	[101]	Self-administered (Clinic)	Not reported	$\alpha = 0.73$	Not reported	Not reported	Significant	Other scale (MAQ)	192
	[102]	Self-administered (Clinic, Present)	93%	Not reported	Not reported	Not reported	Significant	Other scale (MMAS)	275
Drug Attitude Inventory (DAI)	[4]	Self-administered (Home)	Not reported	$\alpha = 0.93$	0.72	0.63	Significant	Therapist report	150
Group 5: Barriers and beliefs									
Beliefs and Behaviour Questionnaire (BBQ)	[67]	Self-administered (Home, Post)	53%	$\alpha = 0.65$	Not reported	Not reported	Significant	Other scale (MARS)	276
Brief Evaluation of Medication Influences and	[89]	Self-administered (Clinic, Absent)	Not reported	$\alpha = 0.63$	0.83	0.71	Not significant (all)	Pharmacy records and Other scale (DAI)	63
Beliefs (BEMIB)							Ì		
Compliance Questionnaire	[61]	Researcher-administered (Home)	Not reported	$\alpha = 0.71$	86.0	0.67	Significant	Self-report	127
Rheumatology (CQR)	[75]	Self-administered (Home)	82%	Not reported	0.62	0.95	Significant	Electronic monitoring (MEMS)	127
Maastricht Utrecht Adherence in Hypertension (MUAH) Ouastionnaire	[69]	Researcher-administered (Clinic)	%06	$\alpha = 0.74$	Not reported	Not reported	Significant (all)	Electronic monitoring (MEMS), Pharmacy records and Other scale (Brief Medication Ottestionnaira)	255
Medication Adherence Report Scale (MARS)	[70]	Self-administered (Clinic, Absent)	Not reported	$\alpha = 0.75$	Not reported	Not reported	Significant (Drug levels only)	Drug levels or Caregiver report	99
	[103]	Self-administered (Clinic, Absent)	Not reported	$\alpha = 0.62$	Not reported	Not reported	Significant	Caregiver report	277

Home—patients completed adherence scale at home; Clinic—patients completed adherence scale in clinic, researcher or clinician may be present or absent while patient fills out form; Telephone — patients completed adherence scale over the telephone; Primary physician = Patient's usual physician; Other HCP = Other health care professionals looking after patient; $\underline{\alpha}$ = Cronbach's alpha of reliability; \underline{BP} = Blood pressure.

Content of the scales

The adherence scales can be categorized into five groups based on the information they seek to elicit (the number of scales is given in parentheses, full details in Table 2). Group 1 scales seek information only on medication-taking behaviour (11), group 2 scales seek information on medication-taking behaviour and barriers to adherence (19), group 3 scales seek information only on barriers to adherence (6), group 4 scales seek information only on beliefs associated with adherence (2) and group 5 scales seek information on barriers and beliefs associated with adherence (5).

Thirty of the 43 scales contained items that asked specific questions about medication-taking behaviour (groups 1 and 2). Most of these adherence scales measure the number of doses taken [9,33,41–50] and contain items such as 'how many days over the past month did you take less than prescribed?' [33] and 'did you miss a tablet yesterday?' [42]. Other adherence scales measuring medication-taking behaviour do so through exploring the frequency of patients not refilling their prescription on time [40, 45, 46, 51].

Twenty adherence scales measuring medication-taking behaviour specified a timeframe for the questions. The timeframe specified ranged from 1 day to 12 months [9, 33, 40, 41, 47, 52–58] [34, 39, 42, 44, 48, 49, 59, 60].

Thirty scales contained items that elicited information on barriers to, and determinants of adherence (groups 2, 3 and 5). Some of these adherence scales are disease-specific and thus explore common barriers that may influence adherence in these disease populations [48, 49, 56, 60, 61]. For example, the Pediatric Inhaler Adherence Questionnaire (PIAQ) explores adherence in patients with asthma and assesses the patient's difficulty in using asthma inhalers and the cost of inhalers [49]. Most of these adherence scales explore forgetfulness as a barrier to adherence and identify some of the situations where forgetfulness may be more common, such as when working or travelling [8, 62–65]. Some adherence scales also explore physical barriers to adherence, such as vision problems, dexterity issues and dysphagia [40, 49, 66].

Seven scales elicited information on the patient's beliefs about their medicines that may relate to adherence (groups 4 and 5). These scales included items identifying beliefs that medicines are necessary, harmful and unnatural [4, 10, 61, 67–70]. For example, the Beliefs about Medicines Questionnaire explores whether the patient holds beliefs that their medicines are necessary as well as whether they have any concerns about their medicines [10].

Forty of the 43 scales contained items that sought to identify aspects of adherence that are consistent with the taxonomy provided by Vrijens *et al.* [12]. Most scales contain items that seek to assess the extent of implementation of a dosing regimen (39/43) (Table 2). Thirteen of the scales also contain items that seek to identify the dis-

continuation of treatment [34, 39, 45, 46, 48, 51, 56, 58, 66, 71–74]. The DAI contained items that sought information on discontinuation (alone) [4] and the SOCA scale identified the initiation of treatment [55]. Three adherence scales do not contain any items that seek to identify the initiation, implementation or discontinuation of treatment [10, 65, 67].

Administration of the scales

The adherence scales have been administered in different ways. Indeed, for the scales with more than one validation or correlation study, the additional studies often administered the scale in a slightly different way. Details of who completed the scale (i.e. patient, clinician or researcher) and where the scale was administered are provided in Table 3. There was a roughly even split between studies that requested the patient to complete the scale and those that had the researcher or clinician complete the scale in consultation with the patient. The location of administration (clinic, home, via telephone or internet) varied between the scales (as reported in Table 3). The time to complete the scale was reported in eight of the 43 adherence scales. Reported times varied from less than 5 min [49, 56, 58, 68] to approximately 25 min (Table 2) [69]. The scales taking less than 5 min to complete consisted of 4 to 14 items. Twelve min was required to complete a 19-item scale and 25 min to complete a 25-item scale [69, 75].

Figure 2 illustrates the conditions in which the adherence scales have been validated. Most of the adherence scales have been validated in a single disease population [4, 8, 9, 40, 42, 44, 47, 49, 51, 53, 54, 56–59, 62, 72] (Figure 2). The Medication Adherence Questionnaire (MAQ), which is a simple four-item questionnaire, has been validated in a broad range of diseases, including hypertension, dyslipidaemia, heart failure and Parkinson's disease [73, 76–78].

Approaches to assessing self-report adherence scales

Assessing the validity of self-report adherence scales differed among the 60 included studies. Details of the studies, assessment of internal consistency, comparison measures and whether the scale was significantly correlated with the comparison measure are provided in Table 3. Similar approaches to validation were seen from scales with similar content.

Medication-taking behaviour The primary method for assessing group 1 and group 2 scales was to determine the correlation between the scale and an objective measure of adherence. Twenty-eight of the 30 scales included in groups 1 and 2 assessed how well the scale correlated with an objective measure of adherence, eight of these scales have been assessed against MEMS and 12 against clinical outcomes (Figure 3).

			•					•			•	•		•			•			•	Maastricht Utrecht Adherence in Hypertension Medication Adherence Report Scale - 10 Compliance Questionnaire Rheumatology	Group 5	Barriers AND beliefs
								•			•	•		•			•				Brief Evaluation of Medication Influences and Beliefs		Вап
		•											Н			\dashv	•				Beliefs and Behaviour Questionnaire Drug Attitude Inventory	4	-Je
																					Beliefs about Medicines Questionnaire	Group,	Beliefs
_	Н									•		-	•			•				•	Self-Efficacy for Appropriate Medication Use Scale	G	_
													•					•			Medication Adherence Reasons Scale		
																				•	Medication Adherence Self-Efficacy Scale-Revised	Group 3	iers
																				•	Medication Adherence Self-Efficacy Scale	Gro	Barriers
						•			•				•		•	•				•	Medication Adherence Questionnaire	•	_
																			•		Adherence Attitude Inventory		
																		•			The Patterns of Asthma Medication Use Questionnaire		
																			•		Simplified Medication Adherence Questionnaire		
																			•		Self-Reported Adherence Questionnaire		
					•		•	•	•	•	•	•	•	•	•	•	•	•		•	Reported Adherence to Medication Scale		2
																		•			Pediatric Inhaler Adherence Questionnaire		rri
				•																	Osteoporosis-Specific Morisky Medication Adherence Scale		2
•																				•	Morisky Medication Adherence Scale		Z
																	•				Medication Adherence Assessment Tool		es
	•																				Immunosuppressant Therapy Adherence Scale	p 2	Adherence scales
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																			•		Godin et al. Self-Reported Adherence Questionnaire		Adi
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											•				•	•		:			Adherence Starts with Knowledge - 20		
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	-														-				•	•	Stages of Change for Adherence Measure		
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Diseases

Disease populations used to validate self-report adherence scales

Figure 2

•	•		•	•	•	•	•		Maastricht Utrecht Adherence in Hypertension Medication Adherence Report Scale - 10 Compliance Questionnaire Rheumatology Brief Evaluation of Medication Influences and Beliefs Beliefs and Behaviour Questionnaire Drug Attitude Inventory Beliefs about Medicines Questionnaire Self-Efficacy for Appropriate Medication Use Scale Medication Adherence Reasons Scale Medication Adherence Self-Efficacy Scale-Revised Medication Adherence Self-Efficacy Scale Medication Adherence Questionnaire	Group 3 Group 4 Group 5	Barriers Beliefs Barriers AND beliefs
		•	•	•	• • •	• • • • • • • • • • • • • • • • • • • •			Adherence Attitude Inventory The Patterns of Asthma Medication Use Questionnaire Simplified Medication Adherence Questionnaire Self-Reported Adherence Questionnaire Reported Adherence to Medication Scale Pediatric Inhaler Adherence Questionnaire Osteoporosis-Specific Morisky Medication Adherence Scale Morisky Medication Adherence Scale Medication Adherence Assessment Tool Immunosuppressant Therapy Adherence Scale Hill-Bone Compliance Scale - 14 Hill-bone Compliance Scale - 10 Godin et al. Self-Reported Adherence Questionnaire Fodor et al. Adherence Questionnaire Choo et al. Questionnaire Brooks Medication Adherence Scale Brief Medication Questionnaire Adherence Starts with Knowledge - 20 Adherence Starts with Knowledge - 12	Group 2	Medication-taking behaviour AND barriers
Caregiver reports	Provider reports	Drug concentrations	Dose counts	Self-report • • •	Electronic monitoring • • • •	Pharmacy records	Other scale •	Clinical outcome	Adherence to Refills and Medications Scale Stages of Change for Adherence Measure Medication Adherence Rating Scale - 5 Kerr et al. Adherence Question Grymonpre et al. Adherence Question Gehi et al. Adherence Question Centre for Adherence Support Evaluation Adherence Bell et al. Adherence Question Barosso et al. 30-day Adherence Question Brief Adherence Rating Scale Adherence Visual Analogue Scale (VAS) Adherence Self-Report Questionnaire	Group I	Medication-taking behaviour

Figure 3Comparison measures of medication-taking behaviour used to validate the self-report adherence scales

Adherence scales

Comparison measure

Barriers and beliefs Scales in groups 3–5 were more likely to rely on alternative approaches to validation. Content validity was typically assessed via a panel of subject matter experts. A range of approaches was utilized for construct validity, including item analysis against tools validated to elicit specific types of health beliefs and factor analyses of responses to other scales or semi-structured interviews.

Three of the six group 3 scales (scales that contain items that elicit information on barriers to adherence only) have been assessed against an objective measure of adherence (one using MEMS, one using clinical outcomes and one using MEMS and clinical outcomes). All six of these scales have been tested for content validity [63–66, 73, 79] and four have also been tested for construct validity [63, 64, 66, 73].

Adherence scales that solely focus on eliciting information regarding a patient's beliefs about their medicines (group 4) have not been assessed against objective measures of adherence. The Beliefs about Medicines Questionnaire and DAI have been significantly correlated with other adherence scales (Table 3). Both scales have been tested for content validity, in addition the Beliefs about Medicines Questionnaire has also been tested for construct validity [4, 10].

Two out of the five group 5 scales (beliefs and barriers) have assessed the correlation between the scale and an objective measure of adherence (both against MEMS). All five of the adherence scales have been correlated with subjective measures: other scales (n=3), self-report (n=1) and caregiver reports (n=1). Four of these adherence scales have also been correlated with objective measures of adherence (Figure 3). All of these adherence scales have been tested for content validity and three (BBQ, BEMIB and MARS-10) have been tested for construct validity [67, 68, 70].

Identifying non-adherence Many self-report adherence scales have recommended cut-offs for identifying nonadherent patients. Twenty-eight scales categorized medication adherence by determining the overall score and separating the population into two groups: adherent and non-adherent [4, 9, 33, 39, 42-47, 49-51, 53, 54, 56-63, 65, 68, 72, 79, 80]. Where reported, the cut-off point to identify non-adherence is most commonly the score that corresponds to patients that took 80% of their medicines as ascertained by an objective measure of adherence such as MEMS. Some scales, such as the Beliefs and Behaviour Questionnaire (BBQ) [67], suggest a cut-off point that corresponds to the score of another self-report adherence scale which has been seen to correspond to patients that took 80% of their medicines according to an objective measure. Other adherence scales, such as the DAI, AAI and MASES-R first split the population into adherent and nonadherent based upon responses to questions about whether medicines were taken or not, and then compared the mean scores of the adherence scales to determine the

cut-off [4, 63, 79]. The SERAD and Gehi *et al.* Adherence Question contain direct medication-taking behaviour questions and answers to these questions are utilized to determine the percentage of adherence and thus dichotomize adherence [9, 59].

A small number of adherence scales have taken a different approach to assigning the adherence cut-off. The MAQ, MMAS, Brief Medication Questionnaire, ASRQ and VAS divided non-adherence into more than two groups, ranging from three to seven [34, 40, 41, 52, 73]. This categorization further differentiated between different levels of patient's adherence to their medicines. The MAQ and MMAS categorized the population into high, medium and low levels of adherence [34, 73]. The MMAS cut-off points were selected based on the correlation with blood pressure control. The Brief Medication Questionnaire grouped the study population into repeat, sporadic and no non-adherence [40]. The ASRQ and VAS classified non-adherence into six and seven levels, respectively based on the researchers' expertise [41, 52].

A small number of scales (12) have assessed the sensitivity and specificity of their cut-off against an objective measure of adherence. The results of these studies are reported in Table 3.

Discussion

We identified 43 adherence scales that have been correlated with a comparison measure of adherence. The identified adherence scales elicit information regarding different facets of adherence including medication-taking behaviour, barriers to and determinants of adherence and beliefs associated with adherence. This information, where accurate, can be put to different uses. Self-report adherence scales can (i) measure medication-taking behaviour, where use of the scale either complements objective measures, or is used as an alternative to objective measures and/or (ii) identify reasons for a patient's nonadherence, by identifying patient-specific barriers or beliefs that impede adherence. The data obtained in this systematic review provide information on how well specific adherence scales can be expected to perform these tasks.

Most of the scales identified as group 1–3 focus on measuring medication-taking behaviour by asking direct questions about medication-taking behaviour or eliciting barriers to good medication-taking behaviour. Group 3 scales focus on barriers to adherence and have the potential to both measure medication-taking behaviour and identify barriers to adherence. The purpose of some group 3 scales is to measure medication-taking behaviour by eliciting information on barriers, as opposed to providing a comprehensive assessment of patient barriers to adherence. The MAQ, for example, is a short four-item group 3 scale that has been well-validated against objective

measures of adherence. The demonstration of a significant correlation between the adherence scale and a suitable objective measure in patients with the same disease seems a reasonable minimum requirement on the use of a scale as an alternative to an objective measure. Of the 36 group 1-3 adherence scales, 20 have been significantly correlated with either MEMS or clinical outcomes. Nine of the 36 adherence scales exploring medication-taking behaviour significantly correlated with the MEMS. The MEMS can record the time of dose actuation and can provide detailed information on medication-taking behaviour over time [28-30]. Fifteen group 1-3 adherence scales have been correlated with clinical outcomes. Few scales have been shown to correlate with MEMS or clinical outcomes in multiple disease states, making the choice of a scale more difficult in patient groups other than those included in the validation studies.

A link between specific levels of adherence and clinical outcomes has been demonstrated in some disease states (e.g. HIV [25, 53, 81] and cardiovascular disease [9, 82, 83]). For the vast majority of disease states, however, no such link has been made. Most adherence scales provide suggested cut-offs for identifying 'non-adherent' patients. Cut-offs permit the identification of patients who may be non-adherent and benefit from education or support. However, the arbitrary nature of the cut-offs provided for most self-report adherence scales needs to be kept in mind. Dichotomizing adherence does not differentiate between types of non-adherence, repeat vs. sporadic adherence or patients at different stages of the medication-taking process. Recent taxonomies of adherence recognize the dynamic nature of patient medicationtaking behaviour. Vrijens et al. acknowledges that the process of medication-taking starts when the patient takes the first dose of medicine (initiation) continues with the implementation of the regimen and ends when the patient discontinues the medicine [12]. Gearing et al. propose a six-phase dynamic model of adherence: treatment initiation, treatment trial, partial treatment acceptance, intermittent treatment adoption, premature discontinuation and full adherence [84]. An important area for future research is the use of self-report adherence scales to identify the different types of non-adherence suggested by Vrijens et al. [12] and Gearing et al. [84].

A substantial number of scales have been validated against clinical outcomes, but no direct measure of medication-taking behaviour such as MEMS; examples include the Barroso 30-day Adherence Question and the Hill-Bone Compliance Scales. A demonstrated correlation between a self-report adherence scale and clinical outcomes in a specific patient population has relatively clear benefits for use of the scale in similar populations of patients. Knowing when this evidence is transferrable into new populations of patients, however, is challenging. For most disease states there are influences on clinical outcomes in addition to medication-

taking behaviour. Factors that influence clinical outcomes play a part in addition to the many factors that may separate measures of adherence by self-report adherence scales from actual medication-taking behaviour. No doubt some of these scales have focused on clinical outcomes due to the availability of clinical data and the relative cost or availability of MEMS. However, validation of a scale against both clinical outcomes and direct measures of medication-taking behaviour is beneficial.

Scales included in groups 2 to 5 include items that elicit reasons a patient may be non-adherent. These scales may identify barriers the patient is experiencing to good medication-taking behaviour, and any patient-specific beliefs about their medicines that may influence adherence. While some Group 3 scales focus more on measuring medication-taking behaviour (e.g. the MAQ), others seek more detailed information on barriers that an individual may be experiencing (e.g. the Pediatric Inhaler Adherence Questionnaire). Scales included in groups 4 and 5 seek to identify patient beliefs about medicines that may influence adherence. Of these scales, the most extensively assessed is the Beliefs about Medicines Questionnaire (BMQ). The BMQ-Specific identifies whether patients hold the belief that their medicine is necessary as well as whether the patient has concerns about their medicine [10].

Scales that focus on identifying reasons for non-adherence appropriately employ validation strategies focused on content and construct validity. The BMQ is a good example of a self-report adherence scale focused on measuring an aspect of adherence other than medication-taking behaviour. The items of the BMQ have been validated through confirmatory principle components analysis and the criterion and divergent validity assessed against similar items in the Illness Perceptions Questionnaire and the Sensitive Soma Scale [10]. The BMQ-Specific has been shown to correlate well with medication-taking behaviour measured by self-report adherence scales. Patients who believe their medicine to be necessary and have fewer concerns have consistently been shown to be more adherent in a range of diseases [85–89].

Scales such as the BMQ are not stand-alone comprehensive adherence scales but, like scales that focus on identifying barriers to adherence, they provide the opportunity for a more comprehensive assessment of a patient's adherence, and the drivers behind that adherence, than subjective or objective measures that focus on measuring medication-taking behaviour. The information provided by self-report adherence scales that seek to identify barriers and beliefs that are influencing adherence may prove useful in addition to accurate information on the patient's medication-taking behaviour. Specifically, these scales may help inform tailored interventions to improve medication adherence, but their use for this purpose is yet to be assessed.



Limitations

This systematic review only included studies of self-report adherence scales that included a comparison measure of medication-taking behaviour. This was deemed appropriate given the importance of measuring medication-taking behaviour in assessing adherence. A consequence of this criterion is that this study does not provide a comprehensive analysis of the validation of self-report adherence scales.

Conclusions

Self-report adherence scales have the potential to measure both medication-taking behaviour, and/or identify barriers and beliefs associated with adherence. Selecting an adherence scale requires consideration of what the adherence scale measures and how well it has been validated. Research on validating and using the existing self-report adherence scales as a measure of medication-taking behaviour is relatively strong. There has been less focus on assessing how information gained from scales that identify patient-specific barriers and beliefs associated with adherence may be used to support wise medicine use. This presents an important and exciting avenue for further research.

Competing Interests

All authors have completed the Unified Competing Interest form at http://www.icmje.org/coi_disclosure.pdf (available on request from the corresponding author) and declare no support from any organization for the submitted work, no financial relationships with any organizations that might have an interest in the submitted work in the previous 3 years and no other relationships or activities that could appear to have influenced the submitted work.

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