**CBT for depression and anxiety adapted for psychosis risk in primary care:**

**A controlled trial to assess feasibility, acceptability and signals of efficacy**

**Supplementary material**

**S1: Socio-demographic characteristics**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
|  | N | Whole sample | CBT-BP | CBT-PR | Between group differences |
|  | N | % | N | % | N | % | *X2* | *p* |
| Gender | 99 |  |  |  |  |  |  | 0.87(2) | 0.65 |
|  Female |  | 57 | 57.6 | 32 | 58.2 | 25 | 56.8 |  |  |
|  Male |  | 41 | 41.1 | 22 | 40.0 | 19 | 43.2 |  |  |
|  Non-binary |  | 1 | 1.0 | 1 | 1.8 | 0 | 0 |  |  |
| Ethnicity | 99 |  |  |  |  |  |  | 4.31 (5) | 0.51 |
|  Asian |  | 1 | 1.0 | 1 | 1.8 | 0 | 0 |  |  |
|  Black |  | 1 | 1.0 | 0 | 0 | 1 | 2.3 |  |  |
|  Mixed |  | 5 | 5.1 | 2 | 3.6 | 3 | 6.8 |  |  |
|  Not stated |  | 4 | 4.0 | 1 | 1.8 | 3 | 6.8 |  |  |
|  Other |  | 2 | 2.0 | 1 | 1.8 | 1 | 2.3 |  |  |
|  White |  | 86 | 86.9 | 50 | 90.9 | 36 | 81.8 |  |  |
| Presenting problem | 99 |  |  |  |  |  |  | 13.00 (4) | 0.01\* |
|  Anxiety |  | 11 | 11.1 | 9 | 16.4 | 2 | 4.5 |  |  |
|  Anxiety & depression |  | 29 | 29.3 | 9 | 16.4 | 20 | 45.5 |  |  |
|  Depression |  | 52 | 52.5 | 66 | 60.0 | 19 | 43.2 |  |  |
|  Other |  | 1 | 1.0 | 0 | 0 | 1 | 2.3 |  |  |
|  Unknown |  | 6 | 6.1 | 4 | 7.3 | 2 | 4.5 |  |  |
| Long term condition | 99 |  |  |  |  |  |  | 0.08 (1) | 0.78 |
|  Reported |  | 39 | 39.4 | 21 | 38.2 | 18 | 40.9 |  |  |
|  None reported |  | 60 | 60.6 | 34 | 61.8 | 26 | 59.1 |  |  |
| Disability | 99 |  |  |  |  |  |  | 0.09(1) | 0.80 |
|  Reported |  | 17 | 17.2 | 10 | 18.2 | 7 | 15.9 |  |  |
|  None reported |  | 82 | 82.8 | 45 | 81.8 | 37 | 84.1 |  |  |
| Employment | 99 |  |  |  |  |  |  | 12.17 (5) | 0.03\* |
|  Employed |  | 35 | 35.4 | 13 | 23.6 | 22 | 50.0 |  |  |
|  Unemployed |  | 29 | 29.3 | 22 | 40.0 | 7 | 15.9 |  |  |
|  Student |  | 24 | 24.2 | 13 | 23.6 | 11 | 25.0 |  |  |
|  Long-term sick |  | 6 | 6.1 | 4 | 7.3 | 2 | 4.5 |  |  |
|  Maternity leave |  | 2 | 2.0 | 2 | 3.6 | 0 | 4.5 |  |  |
|  Not stated |  | 3 | 3.0 | 1 | 1.8 | 2 | 50.0 |  |  |

*Note.* CBT-BP = best practice cognitive behaviour therapy. CBT-PR = cognitive behaviour therapy adapted for psychosis risk. \**p* < 0.05.

**S2: Baseline comparisons**

|  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- |
|  | N | Whole sample | CBT-BP | CBT-PR | Between group differences |
|  |  | M | SD | M | SD | M | SD | *t* | *p* |
| Age | 99 | 28.05 | 11.65 | 28.45 | 11.55 | 27.55 | 11.89 | 0.38 | 0.70 |
| PHQ-9 first | 99 | 18.70 | 5.35 | 19.14 | 4.68 | 18.14 | 6.10 | 0.93 | 0.35 |
| GAD-7 first | 99 | 15.10 | 4.20 | 15.25 | 3.95 | 14.09 | 4.52 | 0.41 | 0.69 |
| WSAS first | 99 | 23.62 | 9.12 | 23.76 | 9.29 | 23.43 | 9.02 | 0.18 | 0.86 |
| Sessions attended | 99 | 10.64 | 6.97 | 9.44 | 7.41 | 12.14 | 6.12 | -1.94 | 0.05\* |
| Sessions not attended | 99 | 1.03 | 1.31 | 1.11 | 1.29 | 0.93 | 1.35 | 0.67 | 0.51 |
| Sessions cancelled  | 99 | 1.93 | 1.96 | 1.36 | 1.79 | 2.64 | 1.97 | -3.37 | 0.001\*\* |

*Note.* PHQ-9 = Physical Health Questionnaire-9 measure of depression. GAD-7 = Generalised Anxiety Disorder-7 measure of anxiety. WSAS = Work and Social Adjustment Scale measure of functioning. CBT-BP = best practice cognitive behaviour therapy. CBT-PR = cognitive behaviour therapy adapted for psychosis risk. \**p* < 0.05. \*\**p* < 0.01.

**S3: Associations between socio-demographic factors and therapy engagement**

|  |  |  |  |  |
| --- | --- | --- | --- | --- |
|  |  | Sessions attended | Sessions DNA | Sessions cancelled |
|   | N | M  | SD  | *r* | *p* | *r* | *p* | *r* | *p* |
| Age | 44 | 27.55 | 11.89 | 0.083 | 0.592 | -.350 | 0.020\* | 0.096 | 0.534 |

|  |  |  |  |  |
| --- | --- | --- | --- | --- |
|  |  | Sessions attended | Sessions DNA | Sessions cancelled |
|   | N | M  | SD  | *t* | *p* | M  | SD  | *t* | *p* | M  | SD  | *t* | *p* |
| Gender Male Female | 1925 | 12.9511.52 | 6.425.95 | 0.76 | 0.45 | 1.370.60 | 1.800.76 | 1.74 | 0.06 | 3.002.36 | 2.161.80 | 1.07 | 0.29 |
| Long term condition Reported None reported | 1826 | 14.0610.81 | 5.596.22 | -1.77 | 0.08 | 0.880.96 | 1.571.22 | 0.17 | 0.86 | 3.002.38 | 1.912.00 | -1.21 | 0.31 |
| Disability Reported None reported | 377 | 12.2411.57 | 5.628.87 | 0.26 | 0.79 | 0.861.29 | 1.291.70 | -0.75 | 0.46 | 2.622.71 | 2.101.11 | -0.11 | 0.91 |

|  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- |
|  | Ethnicity | N | M | SD | *F* | *p* |
| Sessions attended | Black | 1 | 11.00 | . | 1.21 | 0.32 |
| Mixed | 3 | 11.33 | 5.132 |
| Not stated | 3 | 5.00 | 4.359 |
| Other ethnicity | 1 | 15.00 | . |
| White | 36 | 12.75 | 6.194 |
| Sessions not attended  | Black | 1 | 1.00 | . | 0.78 | 0.55 |
| Mixed | 3 | 0.33 | .58 |
| Not stated | 3 | 2.00 | 1.73 |
| Other ethnicity | 1 | 2.00 | . |
| White | 36 | 0.86 | 1.38 |
| Sessions cancelled | Black | 1 | 1.00 | . | 0.25 | 0.91 |
| Mixed | 3 | 2.33 | 2.52 |
| Not stated | 3 | 2.33 | 1.53 |
| Other ethnicity | 1 | 2.00 | . |
| White | 36 | 2.75 | 2.03 |

|  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- |
|  | Employment status | N | M | SD | *F* | *p* |
| Sessions attended | Employed | 22 | 13.05 | 5.67 | 3.21 | 0.02\* |
| Unemployed | 7 | 9.14 | 5.24 |
| Student | 11 | 12.73 | 5.29 |
| Long-term sick | 2 | 19.50 | 9.19 |
| Not stated | 2 | 2.00 | 2.83 |
| Sessions not attended | Employed | 22 | 0.64 | 0.85 | 3.35 | 0.02\* |
| Unemployed | 7 | 0.29 | 0.49 |
| Student | 11 | 1.82 | 1.94 |
| Long-term sick | 2 | 0.00 | 0.00 |
| Not stated | 2 | 2.50 | 2.12 |
| Sessions cancelled | Employed | 22 | 2.45 | 1.65 | 0.35 | 0.84 |
| Unemployed | 7 | 3.43 | 2.51 |
| Student | 11 | 2.45 | 2.42 |
| Long-term sick | 2 | 2.50 | 2.12 |
| Not stated | 2 | 3.00 | 1.41 |

*Note.* \**p* < 0.05.

**S4: Associations between clinical factors and therapy engagement**

|  |  |  |  |
| --- | --- | --- | --- |
|  | Sessions attended | Sessions not attended | Sessions cancelled |
|    | N | M  | SD  | *r* | *p* | *r* | *p* | *r* | *p* |
| PHQ-9 first | 44 | 18.14 | 6.10 | 0.090 | 0.560 | -0.433 | 0.003\*\* | -0.331 | 0.028\* |
| GAD-7 first | 44 | 14.91 | 4.53 | -0.158 | 0.305 | -0.327 | 0.030\* | -0.137 | 0.375 |
| PQ-16 first | 42 | 22.36 | 8.44 | -0.213 | 0.176 | 0.077 | 0.628 | -0.108 | 0.495 |

|  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- |
|  | Presenting problem | N | M | SD | *F* | *p* |
| Sessions attended | Anxiety | 2 | 8.00 | 8.48 | 1.57 | 0.20 |
| Anxiety and depression | 20 | 10.95 | 5.07 |
| Depression | 19 | 14.47 | 6.36 |
| Other diagnosis | 1 | 10.00 | . |
| Unknown | 2 | 7.00 | 9.90 |
| Sessions DNA | Anxiety | 2 | 0.00 | 0.00 | 0.62 | 0.65 |
| Anxiety and depression | 20 | 1.05 | 1.50 |
| Depression | 19 | 0.79 | 1.13 |
| Other diagnosis | 1 | 1.00 | . |
| Unknown | 2 | 2.00 | 2.83 |
| Sessions cancelled | Anxiety | 2 | 1.50 | 0.71 | 0.57 | 0.68 |
| Anxiety and depression | 20 | 2.50 | 2.50 |
| Depression | 19 | 2.95 | 1.43 |
| Other diagnosis | 1 | 4.00 | . |
| Unknown | 2 | 1.50 | 0.71 |

*Note.* PHQ-9 = Physical Health Questionnaire-9 measure of depression. GAD-7 = Generalised Anxiety Disorder-7 measure of anxiety. PQ-16 = Prodromal Questionnaire measure of clinical high risk for psychosis. \**p* < 0.05. \*\**p* < 0.01.

**S5: Associations between socio-demographic factors and therapy outcomes**

|  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- |
|  |  |  |  | PHQ-9 change | GAD-7 change | PQ-16 change |
|    | N | M   | SD   | *r* | *p*  | *r*  | *p*  | *r*  | *p*  |
| Age | 44 | 27.55 | 11.89 | 0.16 | 0.28 | 0.33 | 0.03\* | -0.29 | 0.08 |

|  |  |  |  |  |
| --- | --- | --- | --- | --- |
|  |  | PHQ-9 change | GAD-7 change | PQ-16 change |
|    | N | M   | SD   | *t* | *p*  | M | SD   | *t* | *p*  | N | M   | SD   | *t*  | *p*  |
| Gender Male Female | 1925 | -7.05-5.20 | 6.396.15 | -0.97 | 0.34 | -5.58-4.80 | 5.766.14 | -0.43 | 0.67 | 1521 | 0.874.38 | 9.426.06 | -1.36 | 0.18 |
| Long term condition |   |   |   | 0.83 | 0.41 |   |   | 0.65 | 0.52 |   |   |   | 0.50 | 0.62 |
|  Reported | 26 | -5.35 | 4.78 | -4.65 | 5.42 | 19 | 3.53 | 6.19 |
|  None reported | 18 | -6.94 | 7.97 | -5.83 | 6.68 | 17 | 2.24 | 9.29 |
| Disability |  |  |  | -0.99 | 0.33 |  |  | -0.55 | 0.59 |  |  |  | 0.41 | 0.69 |
|  Reported | 7 | -3.86 | 8.15 | -4.00 | 8.70 | 5 | 1.60 | 8.09 |
|  None reported | 37 | -6.41 | 5.86 | -5.35 | 5.38 | 31 | 3.13 | 7.77 |

*Note.* PHQ-9 = Physical Health Questionnaire-9 measure of depression. GAD-7 = Generalised Anxiety Disorder-7 measure of anxiety. PQ-16 = Prodromal Questionnaire measure of clinical high risk for psychosis. \**p* < 0.05.

**S6: Associations between clinical factors and therapy outcomes**

|  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- |
|  | Ethnicity | N | M | SD | *F* | *p* |
| PHQ-9 change | Black | 1 | -5.00 | . | 0.93 | 0.46 |
| Mixed | 3 | -4.00 | 7.81 |
| Not stated | 3 | -1.00 | 1.00 |
| Other ethnicity | 1 | .00 | . |
| White | 36 | -6.78 | 6.35 |
| GAD-7 change | Black | 1 | -6.00 | . | 0.58 | 0.68 |
| Mixed | 3 | -4.33 | 8.50 |
| Not stated | 3 | -3.33 | 6.66 |
| Other ethnicity | 1 | 3.00 | . |
| White | 36 | -5.56 | 5.83 |
| PQ-16 change | Black | 1 | 4.00 | . | 0.24 | 0.92 |
| Mixed | 3 | 5.33 | 8.14 |
| Not stated | 2 | 7.00 | 4.24 |
| Other ethnicity | 1 | 3.00 | . |
| White | 29 | 2.34 | 8.18 |

|  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- |
|  | Employment status | N | M | SD | *F* | *p* |
| PHQ-9 change | Employed | 22 | -8.23 | 6.50 | 1.87 | 0.34 |
| Unemployed | 7 | -5.71 | 3.50 |
| Student | 11 | -3.27 | 4.08 |
| Long-term sick | 2 | -.50 | 16.26 |
| Not stated | 2 | -3.00 | 4.24 |
| GAD-7 change | Employed | 22 | -6.45 | 5.46 | 0.79 | 0.54 |
| Unemployed | 7 | -2.14 | 6.34 |
| Student | 11 | -5.00 | 5.04 |
| Long-term sick | 2 | -3.00 | 15.56 |
| Not stated | 2 | -4.00 | 5.66 |
| PQ-16 change | Employed | 18 | 3.67 | 8.02 | 0.44 | 0.73 |
| Unemployed | 6 | -.17 | 6.24 |
| Student | 10 | 3.80 | 8.07 |
| Long-term sick | 2 | 1.00 | 11.31 |
| Not stated | 0 | . | . |

|  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- |
|  | Presenting problem | N | M | SD | *F* | *p* |
| PHQ-9 change | Anxiety | 2 | -5.00 | 4.24 | 0.25 | 0.91 |
| Anxiety and depression | 20 | -5.90 | 5.82 |
| Depression | 19 | -6.63 | 7.33 |
| Other diagnosis | 1 | -6.00 | . |
| Unknown | 2 | -2.00 | 2.83 |
| GAD-7 change | Anxiety | 2 | 2.50 | .71 | 1.13 | 0.36 |
| Anxiety and depression | 20 | -4.90 | 6.16 |
| Depression | 19 | -6.32 | 5.82 |
| Other diagnosis | 1 | -7.00 | . |
| Unknown | 2 | -3.00 | 4.24 |
| PQ-16 change | Anxiety | 2 | -1.50 | 6.36 | 0.63 | 0.65 |
| Anxiety and depression | 15 | 1.47 | 8.98 |
| Depression | 17 | 4.06 | 6.89 |
| Other diagnosis | 1 | 8.00 | . |
| Unknown | 1 | 9.00 | . |

*Note.* PHQ-9 = Physical Health Questionnaire-9 measure of depression. GAD-7 = Generalised Anxiety Disorder-7 measure of anxiety. PQ-16 = Prodromal Questionnaire measure of clinical high risk for psychosis.

**S7: Associations between relational measures and therapy engagement and outcomes**

|  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- |
|  | PAM-R disorganised | PAM-R avoidance | PAM-R anxious | ECR-12 avoidance | ECR-12 anxious | DWM-S(A) |
|   | N | M  | SD  | *r* | *p* | *r* | *p* | *r* | *p* | *r* | *p* | *r* | *p* | *r* | *p* |
| Sessions attended | 42 | 10.64 | 7.00 | 0.02 | 0.94 | -0.02 | 0.88 | -0.24 | 0.12 | -0.03 | 0.83 | -0.11 | 0.48 | 0.42 | 0.006\*\* |
| Sessions DNA | 42 | 1.03 | 1.31 | -0.11 | 0.48 | -0.35 | 0.03\* | 0.18 | 0.25 | -0.30 | 0.053 | 0.19 | 0.23 | 0.13 | 0.41 |
| Sessions cancelled | 42 | 1.93 | 1.97 | -0.30 | 0.054 | -0.15 | 0.34 | 0.03 | 0.87 | 0.02 | 0.92 | -0.10 | 0.52 | -0.06 | 0.72 |
| PHQ-9 difference | 41 | -5.08 | 6.84 | 0.04 | 0.82 | 0.003 | 0.98 | 0.09 | 0.56 | -0.02 | 0.92 | -0.14 | 0.38 | -0.40 | 0.008\*\* |
| GAD-7 difference | 42 | -4.36 | 5.71 | 0.15 | 0.35 | 0.11 | 0.49 | 0.07 | 0.68 | 0.03 | 0.83 | -0.18 | 0.24 | -0.34 | 0.03\* |
| PQ-16 at 3 months | 36 | 19.64 | 9.58 | 0.25 | 0.14 | -0.04 | 0.83 | 0.26 | 0.12 | 0.03 | 0.87 | 0.11 | 0.53 | -0.50 | 0.002\*\* |

*Note.* PHQ-9 = Physical Health Questionnaire-9 measure of depression. GAD-7 = Generalised Anxiety Disorder-7 measure of anxiety. PQ-16 = Prodromal Questionnaire measure of clinical high risk for psychosis (CHR-P). PAM-R = Psychosis Attachment Measure-Revised measure of attachment. ECR-12 = Experiences in Close Relationships-Short measure of attachment. DWM-S(A) = Dysfunctional Working Models Scale (adapted) measure of problematic interpersonal beliefs. \*p < 0.05. \*\*p < 0.01.

**CONSORT 2010 checklist of information to include when reporting a pilot or feasibility trial\***

|  |  |  |  |
| --- | --- | --- | --- |
| **Section/Topic** | **Item No** | **Checklist item** | **Reported on page No** |
| **Title and abstract** |
|  | 1a | Identification as a pilot or feasibility randomised trial in the title | 1 |
| 1b | Structured summary of pilot trial design, methods, results, and conclusions (for specific guidance see CONSORT abstract extension for pilot trials) | 3 |
| **Introduction** |
| Background and objectives | 2a | Scientific background and explanation of rationale for future definitive trial, and reasons for randomised pilot trial | 4, 5, 6 |
| 2b | Specific objectives or research questions for pilot trial | 11,12 |
| **Methods** |
| Trial design | 3a | Description of pilot trial design (such as parallel, factorial) including allocation ratio | 6, 7 |
| 3b | Important changes to methods after pilot trial commencement (such as eligibility criteria), with reasons | N/A |
| Participants | 4a | Eligibility criteria for participants | 7 |
| 4b | Settings and locations where the data were collected | 7 |
|  | 4c | How participants were identified and consented | 8, 9 |
| Interventions | 5 | The interventions for each group with sufficient details to allow replication, including how and when they were actually administered | 8 |
| Outcomes | 6a | Completely defined prespecified assessments or measurements to address each pilot trial objective specified in 2b, including how and when they were assessed | 9, 10 |
| 6b | Any changes to pilot trial assessments or measurements after the pilot trial commenced, with reasons | N/A |
|  | 6c | If applicable, prespecified criteria used to judge whether, or how, to proceed with future definitive trial | N/A |
| Sample size | 7a | Rationale for numbers in the pilot trial | 7 |
| 7b | When applicable, explanation of any interim analyses and stopping guidelines | N/A |
| Randomisation: |  |  |  |
| Sequence generation | 8a | Method used to generate the random allocation sequence | N/A |
| 8b | Type of randomisation(s); details of any restriction (such as blocking and block size) | N/A |
| Allocationconcealmentmechanism | 9 | Mechanism used to implement the random allocation sequence (such as sequentially numbered containers), describing any steps taken to conceal the sequence until interventions were assigned | N/A |
| Implementation | 10 | Who generated the random allocation sequence, who enrolled participants, and who assigned participants to interventions | N/A |
| Blinding | 11a | If done, who was blinded after assignment to interventions (for example, participants, care providers, those assessing outcomes) and how | N/A |
| 11b | If relevant, description of the similarity of interventions | N/A |
| Statistical methods | 12 | Methods used to address each pilot trial objective whether qualitative or quantitative | 11. 12, |
| **Results** |
| Participant flow (a diagram is strongly recommended) | 13a | For each group, the numbers of participants who were approached and/or assessed for eligibility, randomly assigned, received intended treatment, and were assessed for each objective | 18 |
| 13b | For each group, losses and exclusions after randomisation, together with reasons | N/A |
| Recruitment | 14a | Dates defining the periods of recruitment and follow-up | 8 |
| 14b | Why the pilot trial ended or was stopped | N/A |
| Baseline data | 15 | A table showing baseline demographic and clinical characteristics for each group | 13 Table 1 & SM Table 1 |
| Numbers analysed | 16 | For each objective, number of participants (denominator) included in each analysis. If relevant, these numbersshould be by randomised group | 13, 14 |
| Outcomes and estimation | 17 | For each objective, results including expressions of uncertainty (such as 95% confidence interval) for anyestimates. If relevant, these results should be by randomised group | 13, 14 |
| Ancillary analyses | 18 | Results of any other analyses performed that could be used to inform the future definitive trial | 13, 14, 15, |
| Harms | 19 | All important harms or unintended effects in each group (for specific guidance see CONSORT for harms) | 3 |
|  | 19a | If relevant, other important unintended consequences | N/A |
| **Discussion** |
| Limitations | 20 | Pilot trial limitations, addressing sources of potential bias and remaining uncertainty about feasibility | 16 |
| Generalisability | 21 | Generalisability (applicability) of pilot trial methods and findings to future definitive trial and other studies | 18 |
| Interpretation | 22 | Interpretation consistent with pilot trial objectives and findings, balancing potential benefits and harms, andconsidering other relevant evidence | 16, 17 |
|  | 22a | Implications for progression from pilot to future definitive trial, including any proposed amendments | 17 |
| **Other information** |  |
| Registration | 23 | Registration number for pilot trial and name of trial registry | 6 |
| Protocol | 24 | Where the pilot trial protocol can be accessed, if available | N/A |
| Funding | 25 | Sources of funding and other support (such as supply of drugs), role of funders | 2 |
|  | 26 | Ethical approval or approval by research review committee, confirmed with reference number | 6 |

Citation: Eldridge SM, Chan CL, Campbell MJ, Bond CM, Hopewell S, Thabane L, et al. CONSORT 2010 statement: extension to randomised pilot and feasibility trials. BMJ. 2016;355. This is an Open Access article distributed in accordance with the terms of the Creative Commons Attribution (CC BY 3.0) license (<http://creativecommons.org/licenses/by/3.0/>), which permits others to distribute, remix, adapt and build upon this work, for commercial use, provided the original work is properly cited.

\*We strongly recommend reading this statement in conjunction with the CONSORT 2010, extension to randomised pilot and feasibility trials, Explanation and Elaboration for important clarifications on all the items. If relevant, we also recommend reading CONSORT extensions for cluster randomised trials, non-inferiority and equivalence trials, non-pharmacological treatments, herbal interventions, and pragmatic trials. Additional extensions are forthcoming: for those and for up-to-date references relevant to this checklist, see [www.consort-statement.org](http://www.consort-statement.org).