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Research Paper

Effect of a complex intervention to improve post-vision screening referral compliance among pre-school children in China: A cluster randomized clinical trial

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ABSTRACT

Background: We investigated whether specific appointments for quality-assured care could increase referral uptake, often low in China, in children's vision screening.

Methods: We randomized children aged 4–7 years in Yudu, Jiangxi, China, by school to Control (free school-based eye screening, parents of children failing screening recommended for further examination [usual practice]) or Intervention (identical examinations, with parents additionally provided with specific appointments for further examinations by quality-assured doctors at a designated local hospital). Both groups could select any hospital for referral exams, which were not free. Six months after screening, parents were interviewed for referral compliance at any hospital (primary outcome) and potential determinants. This trial is registered at the ClinicalTrials.gov, number NCT 03251456.

Findings: Among 9936 children at 63 schools randomized to Intervention (32 schools, 5053 [50.9%] children) or Control (31 schools, 4883 [49.1%] children), 1114 children (11.2% failed screening. Among 513 referred Intervention children (46.1%, 32 schools, mean age 5.36 years, 53% boys) and 601 referred Control children (53.9%, 31 schools, mean age 5.30 years, 57.7% boys), 104 (20.3%) and 135 (22.5%) were lost to follow-up respectively. Under Intention to Treat analysis, assuming children lost to follow-up were non-compliant, Intervention children had significantly higher compliance than Controls (308/513 = 60.6% vs. 225/601 = 37.4%, P < 0.001). In regression models, Intervention group membership (Relative risk [RR] 1.71, 95% confidence interval, 1.36–2.12), travel time to hospital (RR: 0.97, 0.95–0.999), baseline glasses wear (RR: 1.37, 1.17–1.60), strabismus (RR: 1.17, 1.01–1.36) and worse uncorrected vision (RR: 1.41, 1.03–1.92) were associated with compliance.

Interpretation: Providing specific appointments for quality-assured eye care improved referral compliance in this setting.

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1. Introduction

Screening is an essential public health service, aiming to identify high-risk populations and patients at early stages of disease. Timely and appropriate follow-up is imperative to achieve the goal of disease prevention and control. Children detected in vision screening programs are often asymptomatic or at an early clinical stage, thus post-screening clinical follow-up rates are often low [1,2].

Good vision is important for children's health and well-being. The reported prevalence of amblyopia and ocular abnormalities among preschoolers were 1.6% and 2%, respectively in recent large America studies [3,4]. High quality data are unavailable for China, but there is no reason to expect a lesser burden. Preschool vision screening has the potential to reduce the burden of visual impairment on society, and also to improve educational opportunities for children [5,6]. Many countries include preschool
screening as part of regional or national government health care, but practices vary [7]. However in China, with 118.5 million preschool-aged children [8], there is no established national preschool screening or general practitioner-based referral system. Studies are needed to investigate appropriate vision screening and referral models for preschool children in China, and provide high-quality evidence to inform public policy.

A common short-coming of community screening programs is low referral compliance among individuals identified with disease [9]. Several reasons have been identified for lack of health-seeking behavior in Chinese population-based vision surveys, which include lack of knowledge about eye disease and its treatment, and concerns about the quality of locally-available care [10], with the latter proven particularly difficult to overcome [10,11]. A common feature in the Chinese health system, especially at rural, county-level facilities, is not to provide specific appointment dates, with patients choosing their own date to return and attempting to secure a place on the list of appointment dates and offering quality-assured eye care, in a vision screening program among preschool children in China.

### 2. Methods

#### 2.1. Study design

The Yudu Preschool Study is a cluster-randomized, school-based trial conducted in Yudu county, Jiangxi province, China from August 2017 to June 2018. The study was approved by the Ethical Review Committee of the Zhongshan Ophthalmic Center (ZOC), Sun Yat-sen University, Guangzhou, China, and was conducted in accordance with the tenets of the Declaration of Helsinki. Before baseline data collection and screening, principals, and head teachers in participating kindergartens were fully informed of the study process and provided consent for participation. The parents or legal guardians of all participants provided written informed consent, and the trial was registered online (ClinicalTrials.gov ID: NCT 03251456) prior to participant enrollment. At the end of the screening, children and families received printed screening reports. All authors vouch for the accuracy and completeness of the data and for the fidelity to the trial protocol of this report. This study was reported according to the CONSORT extended guidelines.

#### 2.2. Participants, randomization and masking

In 2017, there were 198 registered kindergartens in Yudu county with 31,812 children in attendance. After excluding kindergartens with fewer than 30 children, a total of 189 kindergartens (110 rural and 79 urban kindergartens) remained. A total of 63 kindergartens (32 rural and 31 urban) were selected randomly and assigned with a random number generating program ([www.randomization.com](http://www.randomization.com)) to the Intervention or Control group with a block size of four. School-based cluster randomization was used in this study to prevent contamination, due to the enhanced feasibility of masking in this way, while the children’s parents were aware whether they were part of the intervention or control arm and investigators, but not interviewers, were aware of cluster allocation. All children received vision screening per protocol, and those with abnormal screening results were provided with different referral suggestions in the Intervention and Control group (see below). After vision screening and before referral, the following children were further excluded from the study: (1) Those aged less than four or greater than seven years. (It is not uncommon for Chinese children aged seven years to still attend kindergarten, especially in rural areas). (2) Those whose parents failed to receive the screening report, based on a questionnaire interview.

#### 2.3. Procedures

It is compulsory for preschool children in China to receive a VA screening and a general physical examination within the first semester of school as part of the national public health service program [12]. In addition to the routine screening performed by local health centers, we added free examinations as follows: refraction, examination of the red reflex in both eyes, cover-uncover test to detect strabismus and flashlight examination of the anterior segment, all of which were performed by local nurses, ocular technicians and ophthalmologists who had been trained by doctors from ZOC. General doctors and nurses from the local health center performed general physical examinations at the same time.

Uncorrected visual acuity (UCVA) was assessed by three trained ophthalmic nurses using standard logarithm of the minimum angle of resolution (logMAR) VA charts with tumbling E optotypes at five meters per standardized protocol [13]. For children habitually wearing spectacles, presenting VA (wearing spectacles) was also measured. Refractive power in each eye without cycloplegia was measured using a handheld autorefractor (Mobile Vision Screener plusoptix S12C, Nuremberg, Germany) by two trained technicians per standardized protocol. The red reflex was also examined using the handheld autorefractor.
The cover-uncover test was performed by ophthalmologists to detect the presence of strabismus at both 40 cm and 5 m. Tropias were categorized as esotropia, exotropia, or vertical tropia. The ophthalmologists also examined the appearance of both eyes, lids, corneas, and lenses with a handheld flashlight.

An abnormal screening result was defined as the presence of at least one of the following: (1) strabismus identified by the cover-uncover test; (2) other ocular abnormality detected on flashlight examination of the anterior segment; or (3) finding of any of the below on handheld automated refraction, based on recommendations by the American Academy of Ophthalmology’s PPP: [14]

1) Spherical power $\geq +1.75$ diopters (D) or $\leq -1.50$D in either eye
2) Absolute inter-eye difference of spherical power $\geq 1.50$D
3) Cylindrical power $\geq 1.50$D in either eye
4) Absolute inter-eye difference of cylindrical power $\geq 1.00$D
5) Inability to complete automated refraction in either eye
6) Abnormal red reflex during automated refraction.

In both study groups, we provided printed screening reports and informed parents of any abnormal results (see definition above) at the end of the screening. In the Control group, study personnel informed the parents of children with abnormal screening results that they should take their children to a hospital of their choice for further examination. In the Intervention group, parents were additionally provided with a referral card at the examination site with a medical appointment for further assessments within a designated time period at a referral center in a local secondary hospital (The People’s Hospital of Yudu County), if they elected to go there. Parents in the Intervention group were also informed that doctors at the referral center had received professional training by doctors from ZOC, which would also be present at the referral center for management. Parents in both groups could select any hospital for referral exams, which were not free. Six months after the screening, all parents in both groups were interviewed for compliance assessment.

Two questionnaires were administered in this study. The vision history questionnaire was sent to each parent and collected by teachers on the next day when parents brought their children to school. Six months after the screening, parents were interviewed by phone with another questionnaire concerning factors potentially influencing compliance.

2.4. Outcomes

The primary outcome of the study was compliance with suggested referral eye examinations at any hospital within six months after receiving an abnormal screening report, in both the Intervention and Control groups. The numerator was number of participants who went for referral at any hospital within six months after receiving an abnormal screening report, and the denominator was the total number of participants who were recommend to receive referral eye examinations, whether they actually went for referral or not. Two independent interviewers from the local public health service who were familiar with the local language were trained before the study according to standardized questionnaire interview protocols. The interviewers were unaware of the study group assignment of participating children/families. Parents were asked whether they had taken their children for further examinations as suggested by telephone using the standard questionnaire above. If parents could not be contacted after three phone calls, teachers helped contact the parents and inform them of the phone interview. If parents still could not be contacted after a further phone call, home visits were conducted by two interviewers accompanied by local community health workers or government staff. Participants were deemed lost to follow-up if they could not be contacted using the above methods.

2.5. Statistical analysis

Based on the cluster-randomized design and an assumed compliance rate of 40% in the Control and 70% in the Intervention group [6], we determined that 30 schools (15 schools per group, with an average of 15 children expected to have abnormal vision screening test results at each), would provide 90% power at an alpha error of 0.05, intra-class correlation (ICC) of 0.15 and variation in cluster sizes of 0-60. Assumptions were based on previous screening programs in the area and our prior published trials [6]. Accounting for the stratified sampling by rural versus urban setting, a participation rate of 90% and loss to follow-up of 5%, a total of 60 schools (30 per study group) was required.

Results were presented as median and inter quartile range (IQR) for continuous variables with non-normal distributions and number (proportions) for categorical variables. Spherical equivalent was defined as sphere plus half the negative cylinder. Statistical tests were performed for comparisons of baseline characteristics between the Intervention and Control groups: ordinal logistic regression for age ranging from 4 to 7 years and Somers’ D test for non-normally distributed continuous variables (travel time, uncorrected visual acuity and spherical equivalent refraction; logistic regression for binary variables: gender, urban dwelling, wearing glasses, ophthalmic history including head tilt present, squint or photophobia, ever had visual acuity testing, ever had ocular examination or refraction, ever diagnosed with eye disease, relative diagnosed with eye disease, and eye examination results: strabismus at 0.4 m or 5.0 m and any abnormality in routine examination.) All comparisons took cluster effects within kindergartens into account. The difference between groups in compliance rate was calculated using generalized linear models, with Poisson regression to estimate relative risk (RR) and 95% confidence intervals (CI) [15,16]. All variables significant at the $P < 0.10$ level in simple regression models were included in the multiple regression model. All statistical analyses were performed according to the trial protocol using a commercially available software package (Stata 13.1, StataCorp, College Station TX, USA).

For the intention to treat analysis, which required that all randomized participants to be included in analyses, we estimated the compliance rate in both groups in the most conservative fashion by assuming that all children lost to follow-up did not present for referral. For the risk factor analysis, we used multiple imputation in Stata to impute missing data, including for children lost to follow-up, using logistic regression models for binary variables and linear regression model for continuous variables, selecting the independent variables based on predictive value and availability of data. The multiple imputation approach created 20 copies of the data, in which missing values were imputed by chained equations. Final results were obtained by averaging these 20 datasets using Rubin’s rules [17], which ensured that the standard errors for all regression coefficients took reflected uncertainty in the imputations as well as uncertainty in the estimation.

2.6. Role of the funding source

The funders of the study had no role in study design, data collection, data analysis, data interpretation, or writing of the report. The corresponding author had full access to all the data in the study and had final responsibility for the decision to submit for publication.

3. Results

A total of 63 schools with 9936 children underwent randomization; 32 kindergartens with 5053 children (50.9%) were assigned to the Intervention and 31 kindergartens with 4883 children (49.1%) to Control group. (Fig. 1) A total of 1114 children (11.2%) had abnormal screening results, including 94 children (0.95%) with astigmatism and 116 (1.17%) with an ocular abnormality. Of the 540 children with abnormal vision screening results in the Intervention group, we
Cluster randomized selection

Randomization

Screening

Follow-Up

Analysis

Enrollment

Excluded 9 (4.55%) kindergartens with < 30 children

198 Chinese kindergartens in Yudu, 31812 children

Excluded 4282 (87.7%) children:
- 4242 (99.1%) children with normal screening result
- 16 (0.37%) aged <4 or ≥7 years
- 24 (0.56%) Referral notice not received

189 (95.5%) kindergartens, 30360 (95.4%) children
- Rural: 110 (58.2%) kindergartens, 17001 (56.0%) children
- Urban: 79 (41.8%) kindergartens, 13359 (44.0%) children

63 (33.3%) kindergartens selected. 9936 (32.7%) children
- Rural: 317 (52.7%) children
- Urban: 290 (56.5%) children

Intervention group: 32 (50.8%) kindergartens, 5053 (50.9%) children

Control group: 31 (49.2%) kindergartens, 4883 (49.1%) children

Included in intention-to-treat analysis: 513 children (100%)

Included in intention-to-treat analysis: 601 children (100%)

Lost to follow-up: 104 (20.3%) children, could not be contacted: 104 children

Lost to follow-up: 135 (22.5%) children, could not be contacted: 135 children

Further excluded 22 children (4.07%) aged less than four or greater than seven years and five children (0.93%) whose parents failed to receive the screening report. Thus, a total of 513 Intervention children (95.0%, 223 rural, and 290 urban) were eligible and included in the final analysis. Similarly, 601 (93.8%, 317 rural and 284 urban) of the 641 children with abnormal screening results in the Control group were included in the final analysis. (Fig. 1)

Table 1 illustrates the baseline characteristics of children in the Intervention (mean [SD] age 5.36 [0.94] years) and Control group (5.30 [0.91] years). Children in the two study groups did not differ in age, gender, travel time to hospital, proportion of urban dwellers, or rate of baseline glasses wear. At the time of baseline screening, rates of detection of ocular abnormalities among children in the Intervention and Control groups were similar (Table 1), as were the mean logMAR UCVA (both 0.10 [6/7.5] [0–0.22], P = 0.288). The median (inter-quartile range) spherical equivalent refraction in the better-seeing eye in the Control and Intervention groups were 0.50 (0.00–1.13) D and 0.38 (0.00–1.00) D, respectively (P = 0.350). Six months after receiving an abnormal eye screening report and being informed of the need for further examination, 104 (20.3%) and 135 (22.5%) children were lost to follow-up in the Intervention and Control group, respectively. The baseline characteristics of children who completed and failed to complete follow-up did not differ significantly (Supplementary Table 1).

Under Intention to Treat analysis (assuming all children lost to follow-up were non-compliant), Intervention children had significantly higher compliance than Controls (308/513 = 60.0% vs. 225/601 = 37.4%, P < 0.001). Further analysis showed rural and urban-dwelling children had similar compliance rates in both the Control (35.3% vs. 39.8%, RR=1.13, 95% CI: 0.87–1.45, P = 0.361) and Intervention groups (53.8% vs. 64.8%, RR = 1.20, 95% CI: 0.99–1.47, P = 0.067). Membership in the Intervention group (RR: 1.53, 1.36–1.72, P < 0.001), travel time from kindergarten to hospital (RR: 0.97, 0.95–0.999, P = 0.039), wearing glasses at baseline (RR: 1.37, 95%CI: 1.17–1.60, P < 0.001), signs of strabismus (RR: 1.17, 95%CI: 1.01–1.36, P = 0.032) and UCVA (RR: 1.41, 95%CI: 1.03–1.92, P = 0.031) were significantly associated with successful referral compliance. (Table 2)

Based on the questionnaire interview at six months, among parents who took their children to hospital for further examination, the top two factors influencing choice of hospital were doctor’s perceived professional skill (61.6% and 68.6% in the Control and Intervention group, respectively), and level and size of the hospital (21.8% and 15.4% in the Control and Intervention group, respectively). Among parents who did not take their children for further examination (342 children, 39%), the top three barriers were the same in the Control and Intervention groups: lack of awareness, inconvenience and lack of access (39.1%, 25.2%, 17.2% and 30.2%, 36.5% and 8.33% in the Control and Intervention group, respectively) (Supplementary Table 2).

4. Discussion

Disease screening is a basic public health service strategy worldwide, aimed at identifying individuals at increased risk of certain diseases in the general population for early intervention. Effective screening and early intervention for various ocular conditions have been proven to benefit patients’ prognosis, reduce the cost of treatment, and improve the
models are needed. A practical and comprehensive screening and referral system. Proven and increasing number of children with vision impairment [21], but lacks a practical and comprehensive screening and referral system. Proven models are needed.

We found that 11.2% of the children had abnormal screening results, which is higher than the reported prevalence of vision impairment or amblyopia in previous studies of preschool-aged children [3, 22], suggesting a potentially increasing demand for eye health care in this age range. We found that providing specific appointments for quality-assured eye care at the conclusion of screening could significantly improve referral compliance in both urban and rural areas. Our finding that doctors' perceived professional skill and level of hospital were the most common finding that availability of well-trained doctors can drive service uptake. The impact on the medical system of perceived professional skill and level of hospital were the most common finding that availability of well-trained doctors can drive service uptake. The impact on the medical system of vision screening has been widely acknowledged: 's vision screening has been widely acknowledged:

Table 1
Baseline characteristics of the study participants.

| Characteristics                        | All (N = 1114) | Control Group (N = 601, 53.9%) | Intervention Group (N = 513, 46.1%) | P-value* | Missing data, N (%) |
|----------------------------------------|---------------|--------------------------------|-----------------------------------|----------|---------------------|
| **Demographics**                       |               |                                |                                   |          |                     |
| Mean Age (SD), years                   | 5.32 (0.93)   | 5.30 (0.91)                    | 5.36 (0.94)                       | 0.529    | 0 (0.00)            |
| Boys, No. (%)                          | 619 (55.6)    | 347 (57.7)                     | 272 (53.0)                        | 0.090    | 0 (0.00)            |
| Urban dwellers, No. (%)                | 574 (51.5)    | 284 (47.3)                     | 290 (56.5)                        | 0.529    | 0 (0.00)            |
| Travel time from kindergarten to hospital, median (IQR), minutes | 20 (10–55) | 30 (10–55) | 20 (10–45) | 0.638 | 0 (0.00) |
| Wearing glasses, No. (%)               | 44 (3.98)     | 22 (3.69)                      | 22 (4.32)                         | 0.710    | 8 (0.72)            |
| **Ophthalmic questionnaire, No. (%)**  |               |                                |                                   |          |                     |
| Head tilt present                      | 84 (8.27)     | 46 (8.20)                      | 38 (8.35)                         | 0.929    | 98 (8.80)           |
| Squint or photophobia                  | 79 (7.78)     | 40 (7.13)                      | 39 (6.57)                         | 0.512    | 98 (8.80)           |
| Ever had visual acuity testing         | 318 (31.3)    | 163 (29.1)                     | 155 (34.1)                        | 0.471    | 99 (8.89)           |
| Ever had ocular examination or refraction | 188 (16.6) | 92 (16.4)                     | 96 (21.2)                         | 0.386    | 102 (9.16)          |
| Ever diagnosed with eye disease        | 118 (11.9)    | 60 (10.9)                      | 58 (13.0)                         | 0.503    | 120 (10.8)          |
| Relative diagnosed with eye disease    | 54 (5.47)     | 27 (4.95)                      | 27 (6.11)                         | 0.572    | 127 (11.4)          |

Table 2
Intention to treat analysis of potential predictors of compliance with suggested eye examinations, adjusting for cluster effect within kindergartens. *

| Variables in the simple regression with y-value Relative Risk (95% CI) | Multiple regression* |
|---------------------------------------------------------------|----------------------|
| Simple regression                                              |                      |
| Intervention group (Control group as reference)                | 1.56 (1.37, 1.78)    | -0.001   |
| Relative Risk (95% CI)                                        | 1.53 (1.36, 1.72)    | -0.001   |
| Age (Years)                                                    | 1.01 (0.94, 1.08)    | 0.827     |
| Male sex                                                       | 1.01 (0.92, 1.11)    | 0.809     |
| Urban dwelling                                                 | 1.18 (1.01, 1.39)    | 0.040     |
| Travel time to hospital, minutes                               | 0.96 (0.93, 0.99)    | 0.011     |
| Relative Risk (95% CI)                                        | 0.97 (0.95, 0.999)   | 0.039     |
| Wearing glasses at baseline                                   | 1.02 (1.46, 1.81)    | -0.001   |
| Relative Risk (95% CI)                                        | 1.17 (1.17, 1.60)    | -0.001   |
| Results of ophthalmic examination/questionnaire                |                      |
| Head tilt                                                      | 1.33 (1.15, 1.53)    | -0.001   |
| Squint or photophobia                                          | 1.27 (1.11, 1.45)    | -0.001   |
| Ever had visual acuity test                                   | 1.16 (1.04, 1.29)    | 0.009     |
| Ever had ocular examination or refraction                      | 1.28 (1.15, 1.42)    | -0.001   |
| Ever diagnosed with eye disease                                | 1.31 (1.16, 1.47)    | -0.001   |
| Relative diagnosed with eye disease                            | 0.89 (0.70, 1.14)    | 0.360     |
| Strabismus present                                            |                      |
| At 0.4 m                                                       | 1.00 (0.79, 1.27)    | 0.992     |
| At 5.0 m                                                       | 0.82 (0.63, 1.06)    | 0.122     |
| Uncorrected visual acuity in the better-seeing eye             | 1.88 (1.38, 2.57)    | -0.001   |
| Relative Risk (95% CI)                                        | 1.41 (1.03, 1.92)    | 0.031     |
| Spherical equivalent refraction in the better-seeing eye, dipters | 0.99 (0.93, 1.05)    | 0.651     |
| Abnormality present on eye examination                         | 0.87 (0.73, 1.03)    | 0.101     |

CI: Confidence interval. The significance of the bold entries was shown by the p-values which were less than 0.05.

* Ordinal logistic regression was used to compare age between the control and intervention groups; Somers' D test was used to compare travel time, uncorrected visual acuity and spherical equivalent refraction; Logistic regression was used for all binary variables. All comparisons took account of cluster effect within kindergartens.

We found that 11.2% of the children had abnormal screening results, which is higher than the reported prevalence of vision impairment or amblyopia in previous studies of preschool-aged children [3, 22], suggesting a potentially increasing demand for eye health care in this age range. We found that providing specific appointments for quality-assured eye care at the conclusion of screening could significantly improve referral compliance in both urban and rural areas. Our finding that doctors' perceived professional skill and level of hospital were the most common finding that availability of well-trained doctors can drive service uptake. The impact on the medical system of vision screening has been widely acknowledged: 's vision screening has been widely acknowledged:

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The compliance rate for further eye examinations in our study was only 37.4% in the Control group, suggesting that merely informing parents of eye problems detected in their children leaves nearly two-thirds untreated. Very few previous trials have investigated compliance after vision screening in children, and studies other than trials report varying rates under different healthcare systems [24]. Alison et al. found that 34.4% and 48.1% of children failed to attend scheduled hospital eye services and local optometrist, respectively [25]. Tjiam et al. observed that 23% of children who had abnormal screening results did not present for further examinations as suggested [26]. Reported risk factors for non-adherence with post-eye screening referral include unawareness of the disease severity, poor access to care, financial burdens and inconvenient transport, similar to studies of screening for other diseases [27,28].

In our own study, longer travel time to the hospital decreased compliance, while the presence of symptomatic conditions and evidence of previous contact with the eye care system (e.g. astigmatism, poor vision or baseline glass wear) significantly increased it. Data from our questionnaire showed that lack of knowledge about eye disease was the greatest barrier to referral compliance, indicating that education programs might be beneficial.

Previous studies of interventions to improve post-screening referral compliance have mostly focused on cancer screening [18,25], and few studies have reported on interventions to enhance compliance after eye screening in children [19,28]. One study in India reported a 23-step invention, which also included a specific appointment for children needing refraction, effective in increasing referral compliance [29]. Another school-based study in the United States found that including school nurses in the screening pathway significantly improved referral compliance among children with abnormal screening results [2]. Education programs and follow-up phone calls have also been reported effective in enhancing post-screening referral compliance [26,29]. To the best of our knowledge, the current study is the only randomized clinical trial on referral compliance after eye screening in children.

Implications of our findings for program planners depend upon the practicality of this model for scale-up. Vision screening programs should not only inform patients of abnormal screening results, but also provide specific appointments, which can be easily accomplished, particularly when electronic medical record systems are available. Our simple intervention for quality certification is also scalable and sustainable, as it relied only on modest training and subsequent limited periods of training and oversight by junior doctors from tertiary facilities. Incentives to junior doctors to participate might include the opportunity for professional enhancement through training of trainers, and opportunities to participate in research, as was offered with the current paper.

Strengths of this study include the randomized, controlled design, a representative sample of kindergartens selected at random from a designated area, and a reasonable follow-rate in contacting parents. Limitations should also be acknowledged. It was not possible to mask the examiners, as patients in China expect feedback after examinations, including the follow-up plan which constituted our intervention, to be delivered directly by caregivers. A fifth of children was lost to follow-up in this study. To minimize the possible impact of this, we applied the most conservative possible assumption in our ITT analysis, namely that all such children defaulted on follow-up. We still observed significantly higher compliance rates in the Intervention compared to the Control group, which adds to the robustness of our conclusions. The sensitivity and specificity of vision screening for amblyopia and related risk factors were not assessed in the current study, as the primary outcome was post-screening compliance. A proper screening protocol is also of vital importance for vision screening among preschool children, and this warrants further investigation. Automated refraction without cycloplegia, as we performed here, is likely to result in some inaccuracies due to accommodation in young children. We felt this was acceptable for a screening examination, as cycloplegia can lead to high rates of parental refusal of examinations in China [30]. Further, these inaccuracies are unlikely to affect the reliability of our primary outcome of referral compliance. Lastly, this study only included Chinese children aged 4–7 years in a single county, and the application of our results to other populations can only be made with caution.

Despite these limitations, our study is one of the first designed to address a significant problem, that of poor follow-up referral after eye screening in children, particularly in a middle-income country. Our results suggest that providing specific appointments for quality-assured eye care in eye screening programs can significantly increase service uptake in this setting.

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Declaration of Competing Interest

All authors declare that they have no competing interests.

Supplementary materials

Supplementary material associated with this article can be found in the online version at doi:10.1016/j.eclinm.2020.100258.

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