3| Results of the trial

Overall effect of the intervention

On an intention-to-treat analysis, the adherence to cervical cancer screening was significantly higher among women assigned to intervention than to control group (39.0% vs. 25.7%, p<0.001), corresponding to a difference of 13.3% (95% Confidence Interval [95% CI] 8.1 to 18.5), an Odds Ratio (OR) of 1.85 (95% CI 1.45 to 2.37) and a number needed to screen of 8.

Effect of the intervention by recruiting site and population subgroups

The superiority of the intervention was homogeneous across all the participant primary care units (Figure 3). For <u>USF Espaço Saúde</u> the adherence was 22.2% in the intervention and 19.5% in the control groups, corresponding to a difference of 3.4% (95% CI -10.5 to 17.3) and an OR of 1.17 (95% CI 0.61 to 2.25).

The effectiveness of the intervention was higher among women aged 35-49 years, living in a more deprived area and among those who participated previously in organized screening (Figure 4).

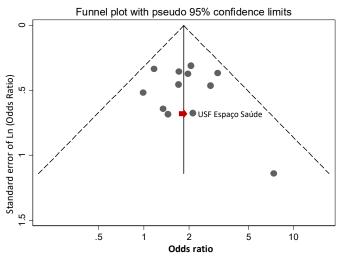


Figure 3 – Funnel plot* representing the odds ratios for the association between the intervention and adherence to cervical cancer screening, across the primary care units involved.

*Dots represent each of the participant primary care units (PCU). The vertical line depicts the overall effect. Results from each PCU are significantly different from the overall effect only when falling outside the triangle defined by the dashed lines.

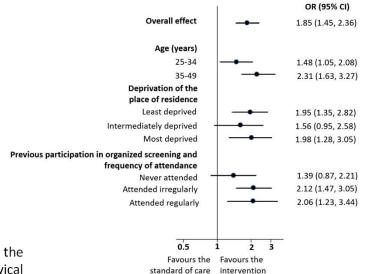


Figure 4 – Forest plot representing the effect of the intervention stratified by age group, deprivation and previous participation in organized screening.

4 Internal and external validity

<u>Internal validity</u>: this investigation was based on a robust study design, and no major threats to the internal validity were identified.

<u>External validity</u>: this trial only included women aged 25 to 49 years, although cervical cancer screening is recommended till 65 years; therefore, the conclusions may not apply to older women, with expectedly more limited digital literacy. The proposed intervention was tested in health care units with characteristics similar to the primary health care settings that may be targeted by this intervention, and was tailored to the specificities of each center. The recruitment and follow-up procedures have replicated the standard practice in each center.

5 | Summary of main findings

A strategy based on automated and customized text messages, phone calls and reminders increased the adherence to cervical cancer screening in 13.3%, in relation to the standard of care (written letter). For <u>USF Espaço Saúde</u>, the proposed intervention increased the adherence in 3.4%. This invitation strategy is operator-independent, and therefore has the potential to be easily implemented on a regular basis.

6 | Funding and conflict of interest

This was an academic study supported by the institutions involved: *Instituto de Saúde Pública da Universidade do Porto, ACeS Porto Ocidental* and *ACeS Marão e Douro Norte*. The authors have no conflict of interest to declare.

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