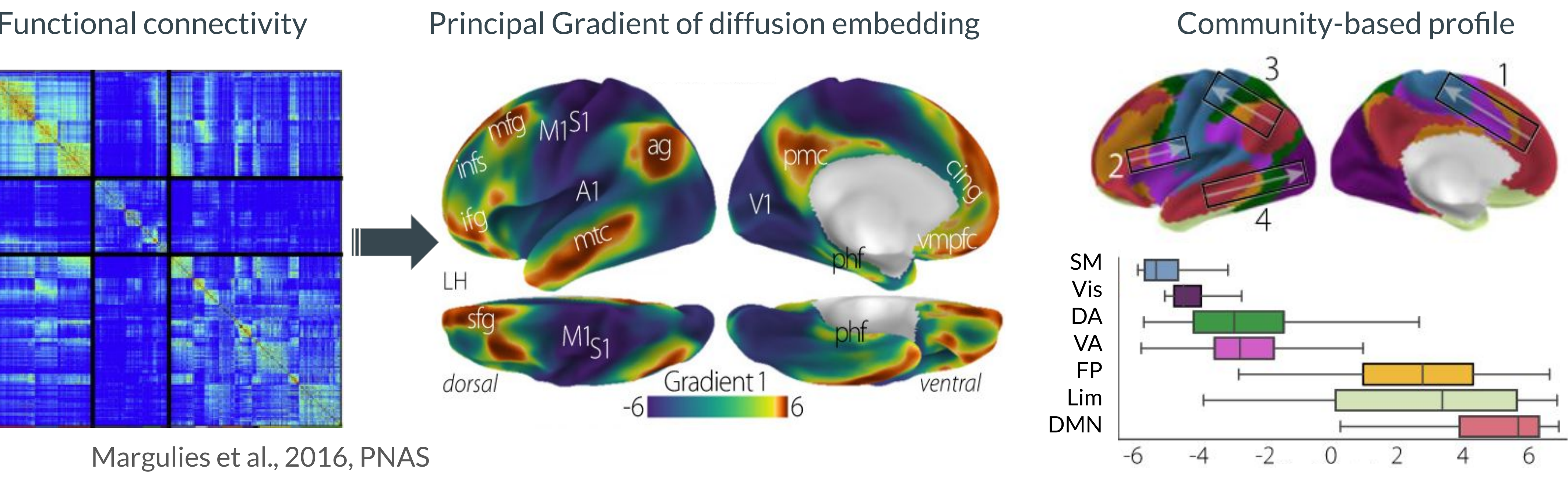


Long-term meditation and meditation related traits increase the connectivity between functional networks



Methods



Mindfulness meditation state and traits modulate the brain gradient connectome in expert and novice meditators

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Background

Mindfulness meditation practice is associated with specific functional relationships between central-executive-network (CEN), salience-network (SN), and default-mode-network (DMN), but also the sensory-perceptual cortices on which they exert top-down modulation

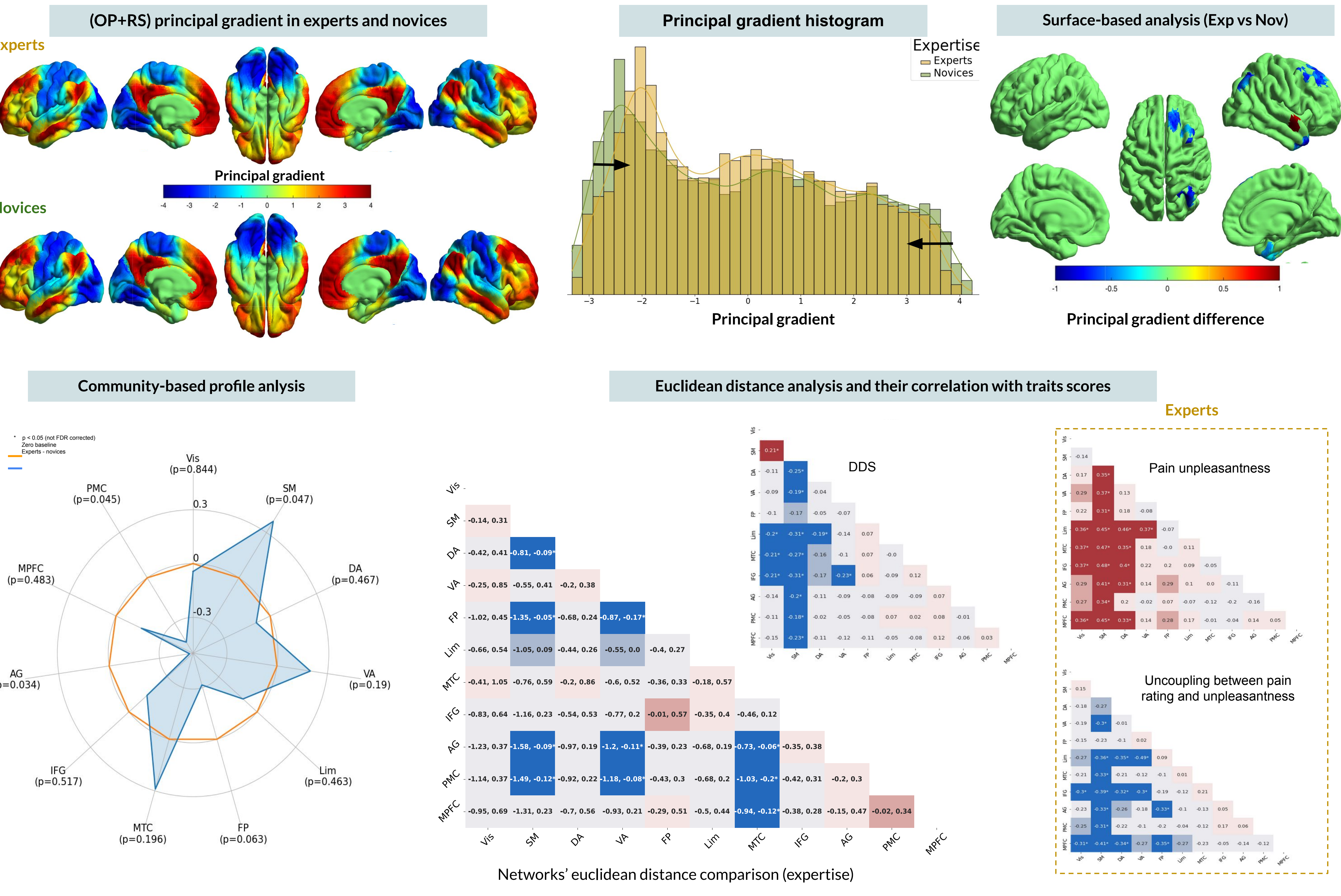
Objective

Methods

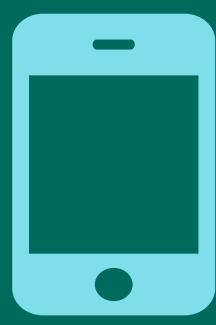
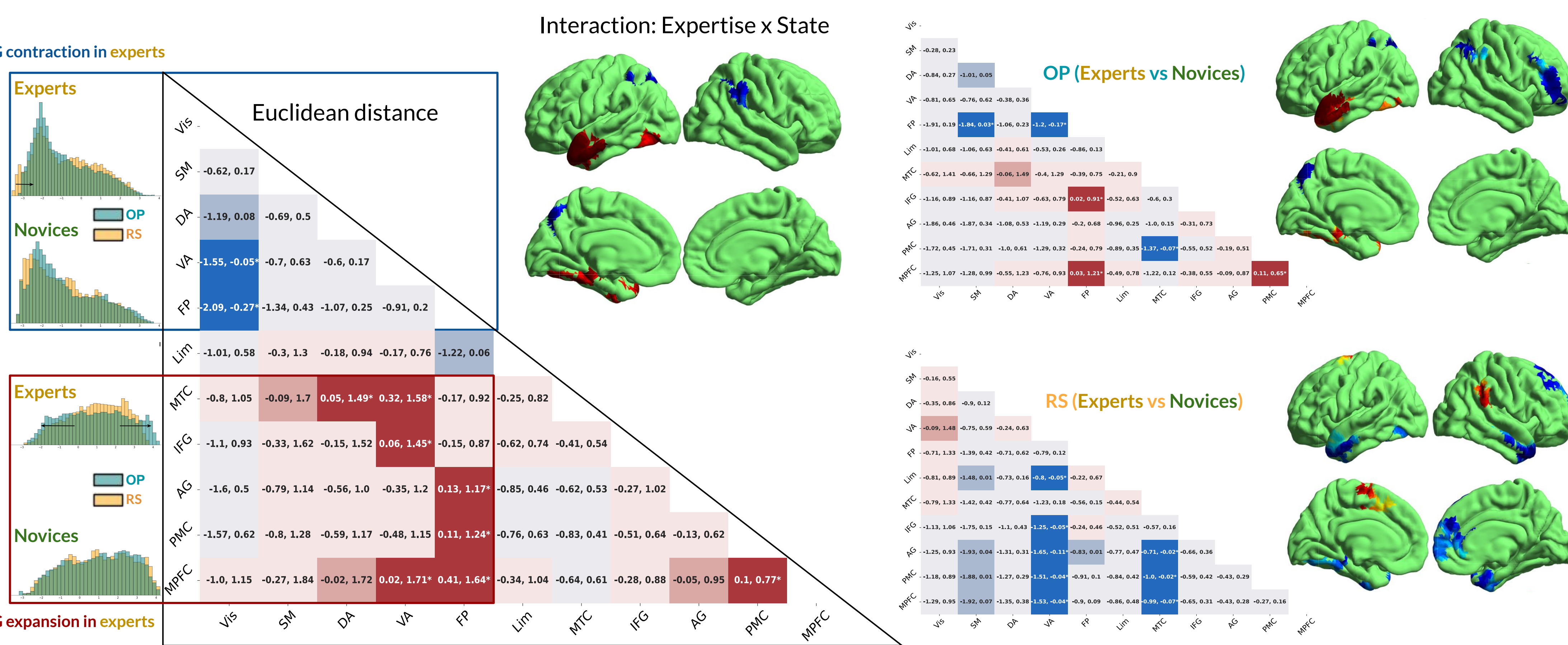
Results

Conclusion

Traits-level analysis



Interaction-level analysis



What doctors can do against inappropriate prescribing and drug overuse in polypharmacy – a rapid review of clinical trials

Objective

To identify from randomized controlled trials strategies for the management of polypharmacy that practicing physicians can employ to achieve meaningful endpoints in multimorbid patients.

An a priori protocol of a sensitive search strategy for interventional trials indexed in MEDLINE and CENTRAL from 2014 to 2018, including related primary sources, was submitted to internal peer review. Studies qualified for eligibility according to prespecified inclusion criteria. The authors independently screened the results, extracted data and assessed the risk of bias using Cochrane methodology. Reporting followed the PRISMA guidelines. Certainty of evidence was appraised using the GRADE approach.

Of a total of 4381 hits, 10 RCT met the inclusion criteria (**Figure 1**). A majority of prespecified PICO criteria were represented (**Table 1**). Overall risk of bias was judged as very serious (**Figure 2**). Incomplete reporting for a priori declared outcomes was detected in 6/10 studies (**Table 2**). Results from 3 RCT (1,324 patients) with high risk of bias on the effects on falls in older patients are unclear (OR 0.99, 95% CI 0.7-1.41). In 1 RCT with high risk of bias, patients (n = 732) experienced a lower rate of adverse drug events (ARR 9.7%, 95% CI 13.4-3.6). 5/7 outcomes did not allow for metanalysis. Certainty of the effect estimates was very low for all outcomes (**Table 3**).

Despite the growing challenges of care for patients with polypharmacy and multimorbidity, evidence from clinical trials that address critical outcomes is limited. It is unclear whether the identified interventions that can be by individual physicians to reduce drug overuse and inappropriate prescribing in multimorbid patients, such as protocols for medication review or educational interventions, resulted in clinical improvements.

| * represented, - not represented in review | |
|----------------------------------------------------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| Population | <ul style="list-style-type: none"> Patients with polypharmacy Patients with multiple chronic diseases Elderly patients (>65 years) Patients with inappropriate prescribing |
| Interventions feasible/applicable to individual physicians | <ul style="list-style-type: none"> Anamnesis/Medication reconciliation Medication therapy management Deprescribing Lists/tools for potentially inappropriate medications Electronic drug management tools Drug databases Peer Review Medical education |
| Control | <ul style="list-style-type: none"> Standards of care |
| Outcomes | <ul style="list-style-type: none"> Hospitalizations Falls Adverse drug events Health-related quality of life Potentially inappropriate medications Drug overuse Drug underuse |
| Settings | <ul style="list-style-type: none"> Primary care Hospitals Admission Outpatient clinics Internal medicine wards Nursing homes |

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graph TD
    A[4381 references imported for screening] --> B[4323 studies screened]
    A --> C[58 duplicates removed]
    B --> D[1431 full-text studies assessed for eligibility]
    B --> E[2892 studies irrelevant]
    D --> F[10 studies included]
    D --> G[1421 studies excluded]
    G --> H["1081 wrong study design  
151 interventions focused on pharmacists  
64 multiprofessional interventions  
47 trial protocol  
39 wrong population  
3 case report  
1 pediatric population"]
  
```

| Count | Description |
|-------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
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| | 1081 wrong study design 151 interventions focused on pharmacists 64 multiprofessional interventions 47 trial protocol 39 wrong population 3 case report 1 pediatric population |
| 10 | studies included |

| Item | Low risk of bias (%) | Unclear risk of bias (%) | High risk of bias (%) |
|-----------------------------------------------------------|----------------------|--------------------------|-----------------------|
| Random sequence generation (selection bias) | 40 | 0 | 60 |
| Allocation concealment (selection bias) | 50 | 0 | 50 |
| Blinding of participants and personnel (performance bias) | 0 | 0 | 100 |
| Blinding of outcome assessment (detection bias) | 30 | 20 | 50 |
| Incomplete outcome data (attrition bias) | 40 | 10 | 50 |
| Selective reporting (reporting bias) | 0 | 40 | 60 |
| Other bias | 50 | 25 | 25 |

| Study | Type | Intervention | Outcomes (♥ fully reported, ○ incompletely reported, - not measured, ✕ not reported) | | | | | | |
|----------------------|--------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------|-------|-----|-------|-----|---------|----------|
| | | | Hospitalizations | Falls | ADE | hrGoL | PIM | Overuse | Underuse |
| Boye, 2017 | RCT, N=612 elderly patients visiting the ED because of a fall (Netherlands) | Discontinuation or dose reduction of fall-risk-increasing drugs (FRID) | ♥ | ♥ | ✕ | ✕ | - | - | - |
| Cullinan, 2017 | RCT, 146 hospital doctors (Ireland) | Short e-learning course, doctor training tool (SCRIPT) including a specific module for prescribing in older patients | - | - | - | - | ○ | - | - |
| Dalleur, 2014 | RCT, N=146 frail elderly inpatients (Belgium) | Review of medications list according to explicit criteria (STOPP criteria) | - | - | - | - | ○ | ○ | - |
| Eveleigh, 2014 | Cluster RCT, N=146 patients with long-term antidepressant use from 46 family practices (Netherlands) | Antidepressant cessation advice in case of inappropriate long-term use in primary care | - | - | - | ○ | - | - | - |
| Frankenthal, 2014 | RCT, N=306 elderly residents at a chronic care geriatric facility prescribed with at least one medication (Israel) | Review of medications list according to explicit criteria (STOPP/START criteria) | ♥ | ♥ | - | ○ | ♥ | - | ♥ |
| Gallagher, 2011 | RCT, N=382 elderly hospitalized patients (Ireland) | Review of medications list according to explicit criteria (STOPP/START criteria) | - | - | - | - | ♥ | ♥ | ♥ |
| García-Górrate, 2014 | Cluster RCT, N=1018 nursing home residents (Spain) | 10 hours educational program on drug use, followed by on-demand support by phone | ♥ | ♥ | - | - | ♥ | ♥ | ♥ |
| O'Connor, 2016 | Cluster RCT, N=732 acutely ill elderly patients admitted to the ED (Ireland) | Single time point presentation to physicians of potentially inappropriate medications according to START/STOPP criteria | ✕ | - | ♥ | - | - | - | - |
| Schäfer, 2018 | Cluster RCT, N=604 elderly multimorbid patients from 55 primary care practices (Germany) | 3 individual narrative doctor-patient dialogues (30 minutes each) over 12-month period | ♥ | - | - | ♥ | - | - | - |
| Wehling, 2016 | RCT, N=409 patients from two geriatric clinics (Germany) | Review of medications list according to explicit criteria (FORTA list) | - | ♥ | - | - | ♥ | ♥ | ♥ |

| Outcome Nº of participants (studies) | Relative effect (95% CI) | Anticipated absolute effects (95% CI) | | | Certainty (GRADE) |
|------------------------------------------------------------------------------------------------------------|-----------------------------|---------------------------------------|-------------------------|-----------------------------------------|----------------------|
| | | Control | Intervention | Difference | |
| Hospitalizations Nº of participants: 1324 (4 RCTs) ^{1,2,3,4} | - | - | - | not pooled | ⊕○○○ VERY LOW |
| Falls Nº of participants: 2049 (3 RCTs) ^{1,3,5} | OR 0.99 (0.70 to 1.41) | 23.1% | 22.9% (17.4 to 29.8) | 0.2% fewer (5.7 fewer to 6.7 more) | ⊕○○○ VERY LOW |
| Adverse drug events Nº of participants: 732 (1 RCT) ⁶ | OR 0.48 (0.31 to 0.79) | 21.0% | 11.3% (7.6 to 17.3) | 9.7% fewer (13.4 fewer to 3.6 fewer) | ⊕○○○ VERY LOW |
| Health-related quality of life Nº of participants: 1053 (3 RCTs) ^{2,7,8} | - | - | - | not pooled | ⊕○○○ VERY LOW |
| Potentially inappropriate medications Nº of participants: 2407 (6 RCTs) ^{3,4,5,9,10} | - | - | - | not pooled | ⊕○○○ VERY LOW |
| Drug overuse Nº of participants: 1955 (4 RCTs) ^{3,3,10} | - | - | - | not pooled | ⊕○○○ VERY LOW |
| Drug underuse Nº of participants: 1908 (4 RCTs) ^{3,4,5} | - | - | - | not pooled | ⊕○○○ VERY LOW |

1. Heyes, A., et al. Effectiveness of medication withdrawal in order to limit health care from the impenetrable "Machete" that is long to reduce the risk of COVID-19. *PLoS ONE* 2020; 15(4): e0230464. <https://doi.org/10.1371/journal.pone.0230464>.
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7. "Effectiveness of Medication Withdrawal in Order to Limit Health Care from the impenetrable 'Machete' that is long to reduce the risk of COVID-19." *PLoS ONE* 2020; 15(4): e0230464. <https://doi.org/10.1371/journal.pone.0230464>.
8. "Effectiveness of Medication Withdrawal in Order to Limit Health Care from the impenetrable 'Machete' that is long to reduce the risk of COVID-19." *PLoS ONE* 2020; 15(4): e0230464. <https://doi.org/10.1371/journal.pone.0230464>.
9. "Effectiveness of Medication Withdrawal in Order to Limit Health Care from the impenetrable 'Machete' that is long to reduce the risk of COVID-19." *PLoS ONE* 2020; 15(4): e0230464. <https://doi.org/10.1371/journal.pone.0230464>.
10. "Effectiveness of Medication Withdrawal in Order to Limit Health Care from the impenetrable 'Machete' that is long to reduce the risk of COVID-19." *PLoS ONE* 2020; 15(4): e0230464. <https://doi.org/10.1371/journal.pone.0230464>.