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We thank Vega-Morales et al (ref) for their interest in our proposed approach for identifying individuals at-risk of rheumatoid arthritis (RA) in a primary care setting [1,2]. The authors agree that primary care is usually the first point of contact for RA patients when they initially develop symptoms. For this reason, they agree that general practitioners (GPs) are well placed to be involved in screening strategies to identify individuals at risk of progressing to RA (ref).

Vega-Morales et al advocate the squeeze test (Gaenslen’s compression test) as a screening tool to aid GPs in identifying at risk individuals for onward referral to a rheumatologist. The rationale for the squeeze test is that compression of the MCP joints evokes pain in an individual with active synovitis. We agree that the squeeze test is useful for identifying early arthritis [3], but its sensitivity is limited and it may need to be combined with other screening tests in order to be sufficiently discriminatory [4]. Nonetheless it is cheap, quick and easy to perform and therefore an appropriate test for the GP who suspects a patient may have developed RA. Automation of the squeeze test, as proposed by Vega-Morales et al, may indeed have a role in this setting (ref). Whether the squeeze test is sensitive enough to identify subclinical synovitis detected by ultrasound will be an important question to address.

Despite its use in early arthritis, we would argue that the squeeze test is inappropriate for identifying at risk individuals, including those with imminent, but not yet established, synovitis. This important group, who have risk factors for RA (including systemic autoimmunity) but crucially do not have arthritis, must be identified before synovitis develops in order to benefit from potential preventative intervention. In line with this, the EULAR taskforce definition for arthralgia suspicious for progression to RA is aimed at identifying homogeneous patient groups that are at risk of developing arthritis before they progress to RA [5].

Testing for anti-CCP antibodies in individuals who present to their GP with a new musculoskeletal complaint can effectively identify individuals at high risk of developing RA,
before the onset of synovitis, and without the need for specialist assessment [2]. In those
that have a positive anti-CCP test, further risk assessment using clinical, serological and
imaging tests can then quantify the risk of progression to arthritis more accurately [6]. Thus
at risk individuals may be followed in longitudinal studies and further stratified for risk
appropriate intervention. The cost-effectiveness of this approach will need to be
determined, but must be considered in the correct context; it may be a key advance toward
preventing a disease that is associated with considerable morbidity, treatment and societal
costs (7).

Vega-Morales et al’s response highlights the changing paradigm of early RA. The concept of
early disease is changing from early arthritis to pre-arthritis while the target changes from
early remission to prevention. As such, novel screening approaches are required. These
must effectively identify at risk individuals prior to the onset of synovitis for bone fide
prevention to be feasible.

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