THE LANCET Rheumatology

Supplementary appendix

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Appendix 1. Demographic and clinical characteristics of patients with RMD after two-dose SARS-CoV-2 mRNA vaccination, stratified by anti-SARS-CoV-2 RBD antibody-response after three- months.

| | Overall¹ (n=809) | Negative (n=56) ¹ | Low-positive ¹ (n=77) | High-positive (n=676) ¹ |
|---|---------------------|------------------------------|----------------------------------|------------------------------------|
| Age, median (IQR) | 49 (39,59) | 50.5 (41·4, 58·8) | 52·3 (40·0, 61·7) | 48.0 (38.6, 58.7) |
| Female sex, no. (%) ² | 745 (92·0) | 51 (91·1) | 72 (93·5) | 622 (92·3) |
| Non-white, no. (%) ² | 61 (7·5) | 8 (14-3) | 7 (9·3) | 46 (6·9) |
| Diagnosis, no. (%) | | | | |
| Inflammatory arthritis ³ | 355 (43·8) | 14 (25·0) | 31 (40.3) | 310 (45·9) |
| Systemic lupus erythematosus | 147 (18·2) | 13 (23·2) | 12 (15·6) | 122 (18·0) |
| Sjögren's syndrome | 39 (4·8) | 3 (5·4) | 2 (2·6) | 34 (5·0) |
| Myositis | 44 (5·4) | 5 (8·9) | 8 (10·4) | 31 (4·6) |
| Vasculitis ⁴ | 16 (1·9) | 4 (7·1) | 3 (3.9) | 9 (1·3) |
| Systemic sclerosis | 8 (1) | 2 (3·6) | 0 (0.0) | 6 (0.9) |
| Overlap connective tissue disease 5 | 188 (23-2) | 12 (21·4) | 20 (26·0) | 156 (23·1) |
| Other ⁶ | 12 (1.5) | 3 (5·4) | 1 (1·3) | 8 (1·2) |
| Vaccine, no. (%) | | | | |
| Pfizer/BioNTech | 436 (53.9) | 34 (61) | 50 (65) | 352 (52·1) |
| Moderna | 376 (46·4) | 22 (39) | 27 (35) | 327 (47-9) |
| Days from D2 to 1mo testing, median (IQR) | 29 (28,32) | 29 (28,33) | 29 (28, 32) | 29 (28,32) |
| Days from D2 to 3mo testing, median (IQR) | 91 (87,94) | 91 (87, 94) | 91 (87, 93) | 91 (87, 94) |
| Therapy included in regimen, no. (%) ⁷ | | | | |
| Conventional DMARD | | | | |
| Azathioprine | 58 (7·2) | 6 (10·7) | 6 (7·8) | 46 (6·8) |
| Hydroxychloroquine | 322 (39·8) | 17 (30·4) | 27 (35·1) | 278 (41·1) |
| Leflunomide | 33 (4·1) | 2 (3·6) | 3 (3.9) | 28 (4·1) |
| Methotrexate | 209 (25·8) | 14 (25·0) | 19 (24·7) | 176 (26·0) |
| Mycophenolate ⁸ | 114 (14·1) | 16 (28·5) | 19 (24·7) | 79 (11·6) |
| Sulfasalazine | 35 (4·3) | 2 (3·6) | 2 (2·6) | 31 (4.6) |
| Tacrolimus | 11 (1·3) | 0 (0.0) | 5 (6·5) | 6 (0.9) |
| Biologic | | | | |
| Belimumab | 83 (10·2) | 3 (5·4) | 8 (10·4) | 72 (10·7) |
| Interleukin inhibitors | | | | |
| Anakinra | 2 (0·3) | 0 (0.0) | 0 (0.0) | 2 (0·3) |
| IL-6 inhibitors ⁹ | 10 (1·2) | 0 (0.0) | 2 (2·6) | 8 (1·2) |
| IL-17 inhibitors ¹⁰ | 19 (2.3) | 0 (0.0) | 1 (1·3) | 18 (2·7) |

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| Ustekinumab | 14 (1·7) | 0 (0-0) | 0 (0.0) | 14 (2·1) |
|--------------------------------|------------|-----------|-----------|------------|
| JAK inhibitor ¹¹ | 50 (6·1) | 1 (1·7) | 1 (1·3) | 48 (7·1) |
| Rituximab | 52 (6·4) | 33 (58·9) | 11 (14·3) | 8 (1·2) |
| TNF inhibitor ¹² | 173 (21-4) | 5 (8·9) | 23 (29·9) | 145 (21·4) |
| Immunomodulatory ¹³ | 43 (5·3) | 2 (3·6) | 14 (18·2) | 27 (4·0) |
| Glucocorticoid ¹⁴ | 227 (28) | 26 (46·4) | 32 (41·6) | 169 (25) |

- 1 The percentages in these columns are shown as percent of each category in the overall column. Negative antibody response was defined per manufacturer data as Roche Elecsys anti-RBD pan Ig <0.8 units/mL. Low-positive antibody response was defined as anti-RBD pan Ig 0.8-50 units/mL. High-positive antibody response was defined as anti-RBD pan Ig >50 units/mL.
- 2 The denominators for these categories differ from the total N as 1 participant selected "prefer not to answer" for sex, 8 selected "prefer not to answer" for race, 2 did not respond to the vaccine manufacturer question, and 1 did not respond to the prior COVID-19 diagnosis question.
- 3 Rheumatoid arthritis, ankylosing spondylitis, psoriatic arthritis, reactive arthritis, or inflammatory bowel disease associated arthritis
- 4 Polyarteritis nodosa, Behcet's syndrome, polymyalgia rheumatica, temporal arteritis, eosinophilic granulomatosis polyangiitis, granulomatous polyangiitis, Henoch-Schonlein purpura, microscopic polyangiitis, or Takayasu arteritis
- 5 Denotes combination of two or more of inflammatory arthritis, systemic lupus erythematosus, Sjögren's syndrome, myositis, vasculitis and systemic sclerosis
- 6 Other denotes participants with Adult onset Still's disease, undifferentiated connective tissue disease, or sarcoidosis
- 7 Patients could select >1 medication, thus n>809
- 8 Mycophenolate includes mycophenolic acid and mycophenolate mofetil.
- 9 Interleukin (IL)-6 inhibitors include sarilumab and tocilizumab.
- 10 IL-17 inhibitors include ixekizumab and secukinumab.
- 11 Janus kinase (JAK) inhibitors include tofacitinib and upadacitinib.
- 12 TNF inhibitors include adalimumab, certolizumab, etanercept, golimumab, and infliximab.
- 13 Immunomodulatory includes intravenous immunoglobulin (IVIg) and subcutaneous Immunoglobulin (SCIg).
- 14 Glucocorticoid includes prednisone and prednisone equivalents.

Appendix 2. Clinical characteristics of 15 patients with de novo antibody response at three months following two-dose SARS-CoV-2 vaccination

| Age | Sex | Diagnosis | Therapy | 1 Month Titer | 3 Month Titer |
|-----|--------|--|--|------------------|------------------|
| 62 | Female | Overlap connective tissue disease ¹ | Azathioprine Belimumab Glucocorticoid ² Hydroxychloroquine | <0.40 | 1.30 |
| 37 | Female | Myositis | Azathioprine Glucocorticoid¹ Tacrolimus | <0.40 | 2.70 |
| 63 | Female | Scleroderma, Vasculitis ³ | Mycophenolate ⁴ | <0.40 | 2.70 |
| 50 | Female | Systemic lupus erythematosus | Glucocorticoid ² Hydroxychloroquine IVIg Mycophenolate ⁴ Rituximab | <0.40 | 6-00 |
| 40 | Female | Systemic lupus erythematosus | Belimumab Hydroxychloroquine Glucocorticoid ² Mycophenolate ⁴ | <0.40 | 6-40 |
| 70 | Female | Myositis | Hydroxychloroquine IVIg Mycophenolate⁴ | <0.40 | 7.50 |
| 39 | Female | Systemic lupus erythematosus | Hydroxychloroquine Mycophenolate ⁴ Glucocorticoid ² | 0.76 | 7.70 |
| 34 | Female | Myositis | Azathioprine IVIg | <0.40 | 10·10 |
| 37 | Female | Vasculitis ³ | Mycophenolate⁴ Tacrolimus | <0.40 | 15.80 |
| 60 | Female | Vasculitis ³ | Mycophenolate⁴ | <0.40 | 60-30 |
| 30 | Female | Overlap connective tissue disease ¹ | Hydroxychloroquine Mycophenolate⁴ | <0.40 | 80.70 |
| 43 | Female | Overlap connective tissue disease ¹ | Glucocorticoid ² Mycophenolate ⁴ Tacrolimus | <0.40 | 146.00 |
| 48 | Female | Systemic lupus erythematosus | Glucocorticoid ² Hydroxychloroquine, Mycophenolate ⁴ Tacrolimus | 0.43 | 174-30 |
| 52 | Female | Myositis | Hydroxychloroquine Mycophenolate ³ Glucocorticoid ² | <0.40 | 216-90 |
| 65 | Female | Overlap connective tissue disease ¹ | Mycophenolate ⁴ | <0.40 | 249.00 |

¹ Denotes combination of two or more of inflammatory arthritis, systemic lupus erythematosus, Sjögren's syndrome, myositis, vasculitis and systemic sclerosis

² Glucocorticoid includes prednisone and prednisone equivalents.

³ Polyarteritis nodosa, Behcet's syndrome, polymyalgia rheumatica, temporal arteritis, eosinophilic granulomatosis polyangiitis, granulomatous polyangiitis, Henoch-Schonlein purpura, microscopic polyangiitis, or Takayasu arteritis

⁴ Mycophenolate includes mycophenolic acid and mycophenolate mofetil.