

Supplemental Table 1. Full cohort summary, probands and first-degree relatives by time period and location

| | Mayo 1 | | | Mayo 2 | | | Combined (Mayo 1 & Mayo 2) | | |
|----------|--------|-------|-------|--------|-----|---------------|----------------------------|-----|---------------|
| | Other | Other | Other | Other | MN | All locations | Other | MN | All locations |
| Probands | 118 | 118 | 118 | 125 | 100 | 225 | 243 | 187 | 430 |
| FDRs | 327 | 327 | 327 | 295 | 265 | 560 | 622 | 557 | 1179 |

Abbreviations: MGUS, monoclonal gammopathy of undetermined significance; MM, multiple myeloma; SMM, smoldering multiple myeloma. Values are counts. Other=Proband resided outside of Minnesota at diagnosis. MN=Proband resided in Minnesota at diagnosis. All locations=Minnesota and all other locations.

Mayo 1=participants recruited from Vachon et al. 2009, 2005-2009.

Mayo 2=participants recruited from 2009-2015

Supplemental Table 2. Minnesota Region: Demographic characteristics of Myeloma (MM/SMM) probands and their first degree relatives

| Characteristics | Probands (n=187) | First-degree relatives of probands (n=557) |
|--------------------------------|---------------------|---|
| Sex | | |
| Male | 108 (57.8) | 236 (42.4) |
| Female | 79 (42.2) | 321 (57.6) |
| Age category (years) | | |
| <40 | 3 (1.6) | --- |
| 40-49 | 14 (7.5) | 126 (22.6) |
| 50-59 | 51 (27.3) | 148 (26.6) |
| 60-69 | 67 (35.8) | 138 (24.8) |
| 70-79 | 43 (23.0) | 101 (18.1) |
| 80+ | 9 (4.8) | 44 (7.9) |
| Relationship to proband | | |
| Parent | - | 52 (9.3) |
| Sibling | - | 367 (65.9) |
| Child | - | 138 (24.8) |

Abbreviations: MM, multiple myeloma; SMM, smoldering multiple myeloma; FDRs, first degree relatives. Values reported are n (%).

Supplemental Table 3. Demographic characteristics of Myeloma (MM /SMM) probands and First-degree relatives (all locations)

| Characteristics | Mayo 1 ¹ | | Mayo 2 | |
|--------------------------------|---------------------|------------------------------------|-------------------|---------------------------------|
| | Probands n=205 | First-degree relatives n=619 | Probands n=225 | First-degree relatives n=560 |
| Sex | | | | |
| Male | 107 (52.2) | 249 (40.2) | 134 (59.6) | 238 (42.5) |
| Female | 98 (47.8) | 370 (59.8) | 91 (40.4) | 322 (57.5) |
| Age (years) | | | | |
| 18-39 | 6 (2.9) | | 3 (1.3) | |
| 40-49 | 24 (11.7) | 202 (32.6) | 16 (7.1) | 115 (20.5) |
| 50-59 | 62 (30.2) | 132 (21.3) | 73 (32.4) | 148 (26.4) |
| 60-69 | 71 (34.6) | 139 (22.5) | 79 (35.1) | 153 (27.3) |
| 70-79 | 37 (18.0) | 89 (14.4) | 47 (20.9) | 92 (16.4) |
| 80+ | 5 (2.4) | 57 (9.2) | 7 (3.1) | 52 (9.3) |
| Relationship to proband | | | | |
| Parent | - | 87 (14.1) | - | 52 (9.3) |
| Sibling | - | 366 (59.1) | - | 375 (67.0) |
| Child | - | 166 (26.8) | - | 133 (23.8) |

Abbreviations: MM, multiple myeloma; SMM, smoldering multiple myeloma. Values reported are n (%).

¹Samples from 2009 Cohort: Vachon CM, et al.

Supplemental Table 4. Prevalence of MGUS in first-degree relatives of Myeloma probands (MM/SMM) compared with Olmsted County residents. (Mayo 1 compared to Mayo 2 and Olmsted County)

| | Mayo 1 ⁽¹⁾ | | | Mayo 2 | | | Olmsted County | | |
|-----------------------------|------------------------------------|------------------------------|------------------------|------------------------------------|------------------------------|------------------------|---------------------|-------------------------------|------------------------|
| | First-degree relatives of probands | | | First-degree relatives of probands | | | | | |
| Age (years) | Total (n=619) | Total with MGUS (n=39) | Prevalence (95% CI) | Total (n=560) | Total with MGUS (n=36) | Prevalence (95% CI) | Total (n= 21463) | Total with MGUS (n=694) | Prevalence (95% CI) |
| 40-49 | 202 | 3 | 1.5 (0.3,4.3) | 115 | 3 | 2.6 (0.5,7.6) | | | |
| 50-59 | 132 | 5 | 3.8 (1.2,8.8) | 148 | 6 | 4.1 (1.5,8.8) | 8373 | 141 | 1.7 (1.4,2.0) |
| 60-69 | 139 | 11 | 7.9 (4.0,14.2) | 153 | 10 | 6.5 (3.1,12.0) | 6019 | 178 | 3.0 (2.5,3.4) |
| 70-79 | 89 | 14 | 15.7 (8.6,26.4) | 92 | 8 | 8.7 (3.8,17.1) | 4508 | 205 | 4.6 (4.0,5.2) |
| 80+ | 57 | 6 | 10.5 (3.9,22.9) | 52 | 9 | 17.3 (7.9,32.9) | 2563 | 170 | 6.6 (5.7,7.7) |
| Total, adjusted (40+) | 619 | 39 | 6.1 (4.1,8.0) | 560 | 36 | 5.8 (3.7,7.8) | | | |
| Total, adjusted (50+) | 417 | 36 | 8.5 (5.7,11.4) | 445 | 33 | 7.3 (4.7,9.8) | 21463 | 694 | 3.2 (3.0,3.5) |

Abbreviations: MGUS, monoclonal gammopathy of undetermined significance; MM, multiple myeloma; SMM, smoldering multiple myeloma; 95% CI, 95% confidence interval. Rates for MGUS determined by serum protein electrophoresis with immunofixation for confirmation.

Adjusted=Age and sex adjusted to 2000 US total population.

¹Samples from 2009 Cohort: Vachon CM, et al.

Supplemental Table 5. Minnesota Region: Prevalence of MGUS in first-degree relatives of Myeloma (MM / SMM) probands compared with Olmsted County residents

| Age (years) | First-degree relatives of probands | | | Olmsted County | | |
|---|------------------------------------|---------------------------|------------------------|--------------------|----------------------------|------------------------|
| | Total (n=557) | Total with MGUS (n=43) | Prevalence (95% CI) | Total (n=21463) | Total with MGUS (n=694) | Prevalence (95% CI) |
| 40-49 | 126 (22.6) | 1 (2.3) | 0.8 (0.0,4.4) | - | - | - |
| 50-59 | 148 (26.6) | 6 (14.0) | 4.1 (1.5,8.8) | 8373 (39.0) | 141 (20.3) | 1.7 (1.4,2.0) |
| 60-69 | 138 (24.8) | 13 (30.2) | 9.4 (5.0,16.1) | 6019 (28.0) | 178 (25.6) | 3.0 (2.5,3.4) |
| 70-79 | 101 (18.1) | 16 (37.2) | 15.8 (9.1,25.7) | 4508 (21.0) | 205 (29.5) | 4.6 (4.0,5.2) |
| 80+ | 44 (7.9) | 7 (16.2) | 15.9 (6.4,32.8) | 2563 (11.9) | 170 (24.5) | 6.6 (5.7,7.7) |
| Total, adjusted (Ages 40+) | 557 (100) | 43 (100) | 6.6 (4.5,8.6) | - | - | - |
| Total, adjusted (Ages 50+) | 431 (77.4) | 42 (97.7) | 9.6 (6.6,12.5) | 21463 (100) | 694 (100) | 3.2 (3.0,3.5) |

Abbreviations: MGUS, monoclonal gammopathy of undetermined significance; MM, multiple myeloma; SMM, smoldering multiple myeloma; 95% CI, 95% confidence interval. Values reported are n (%). Rates for MGUS determined by serum protein electrophoresis with immunofixation for confirmation. Adjusted=age and sex adjusted to 2000 US total population.

