

Supplementary table 1. Results from multiple linear regression fitting age and CMV response as covariates for CD8⁺ T cell subset composition. Table shows the unstandardized coefficient, significance and 95% confidence interval from the output of SPSS software for each CD45RA/CD27 subset. Unit of age is equal to 1 year.

| <i>CD45RA⁺CD27⁺</i> | unstandardized | <i>p</i> -value | 95.0% CI | |
|-------------------------------------------|----------------|-----------------|-------------|-------------|
| | coefficient | | lower limit | upper limit |
| age | -0.59 | <.0001 | -0.7 | -0.49 |
| CMV response | -15.62 | <.0001 | -21.46 | -9.77 |

| <i>CD45RA⁻CD27⁺</i> | unstandardized | <i>p</i> -value | 95.0% CI | |
|-------------------------------------------|----------------|-----------------|-------------|-------------|
| | coefficient | | lower limit | upper limit |
| age | -0.03 | 0.52 | -0.12 | 0.06 |
| CMV response | -6.18 | <.01 | -10.86 | -1.49 |

| <i>CD45RA⁻CD27⁻</i> | unstandardized | <i>p</i> -value | 95.0% CI | |
|-------------------------------------------|----------------|-----------------|-------------|-------------|
| | coefficient | | lower limit | upper limit |
| age | 0.125 | <0.05 | 0.005 | 0.24 |
| CMV response | 9.05 | < 0.01 | 2.58 | 15.53 |

| <i>CD45RA⁺CD27⁻</i> | unstandardized | <i>p</i> -value | 95.0% CI | |
|-------------------------------------------|----------------|-----------------|-------------|-------------|
| | coefficient | | lower limit | upper limit |
| age | 0.385 | <.0001 | 0.27 | 0.5 |
| CMV response | 14.77 | < .0001 | 8.43 | 21.11 |

Supplementary table 2. Statistical comparisons of percentage of CD8⁺ T cell cytokine producing populations following overnight stimulation of PBMCs with anti-CD3 and IL-2. Results from Paired T-test.

| | IFN | TNF | IL2/IFN | IL2/TNF | IFN/TNF | IL2/IFN/TNF |
|--------------|--------|--------|---------|---------|---------|-------------|
| Young | | | | | | |
| IL2 | 0.0089 | 0.0013 | 0.0365 | ns | 0.0223 | ns |
| IFN | - | 0.0124 | 0.0081 | 0.0085 | 0.0491 | 0.0124 |
| TNF | - | - | 0.0012 | 0.0012 | ns | 0.0015 |
| IL2/IFN | - | - | - | ns | 0.0220 | 0.0432 |
| IL2/TNF | - | - | - | - | 0.0219 | ns |
| IFN/TNF | - | - | - | - | - | 0.0229 |
| Old | | | | | | |
| IL2 | 0.0343 | 0.0001 | ns | 0.0250 | 0.0089 | ns |
| IFN | - | ns | 0.0252 | 0.0462 | 0.0157 | ns |
| TNF | - | - | <0.0001 | 0.0002 | 0.0282 | 0.0002 |
| IL2/IFN | - | - | - | 0.0313 | 0.0081 | ns |
| IL2/TNF | - | - | - | - | 0.0096 | ns |
| IFN/TNF | - | - | - | - | - | 0.0079 |

Supplementary table 3. Statistical comparisons of telomere length of CD8⁺ T cell cytokine producing populations following overnight stimulation of PBMCs with anti-CD3 and IL-2. Results from Paired T-test.

| | IFN | TNF | IL2/IFN | IL2/TNF | IFN/TNF | IL2/IFN/TNF | Negative |
|--------------|-------|-------|---------|---------|---------|-------------|----------|
| Young | | | | | | | |
| IL2 | 0.012 | 0.004 | 0.011 | ns | 0.006 | 0.018 | ns |
| IFN | - | ns | ns | ns | 0.016 | ns | .023 |
| TNF | - | - | ns | 0.026 | ns | 0.001 | .006 |
| IL2/IFN | - | - | - | ns | ns | ns | ns |
| IL2/TNF | - | - | - | - | 0.028 | ns | ns |
| IFN/TNF | - | - | - | - | - | 0.003 | .001 |
| IL2/IFN/TNF | - | - | - | - | - | - | .034 |
| Old | | | | | | | |
| IL2 | ns | ns | 0.028 | 0.013 | ns | ns | ns |
| IFN | - | ns | ns | ns | ns | ns | ns |
| TNF | - | - | ns | ns | ns | ns | ns |
| IL2/IFN | - | - | - | ns | ns | ns | ns |
| IL2/TNF | - | - | - | - | ns | 0.048 | ns |
| IFN/TNF | - | - | - | - | - | ns | ns |
| IL2/IFN/TNF | - | - | - | - | - | - | ns |

Supplementary table 4. Statistical comparisons of percentage of CD8⁺ T cell cytokine producing populations following overnight stimulation of PBMCs with NLV peptide. Results from Wilcoxon Signed Rank Test.

| | IFN | TNF | IL2/IFN | IL2/TNF | IFN/TNF | IL2/IFN/TNF |
|--------------|------|------|---------|---------|---------|-------------|
| Young | | | | | | |
| IL2 | .043 | .176 | .018 | .028 | .128 | .686 |
| IFN | - | .612 | .018 | .018 | .866 | .018 |
| TNF | - | - | .018 | .018 | .612 | .018 |
| IL2/IFN | - | - | - | .237 | .028 | .043 |
| IL2/TNF | - | - | - | - | .018 | .018 |
| IFN/TNF | - | - | - | - | - | .018 |
| Old | | | | | | |
| IL2 | .138 | .080 | .225 | .893 | .043 | .176 |
| IFN | - | .345 | .043 | .043 | .345 | .043 |
| TNF | - | - | .043 | .043 | .225 | .043 |
| IL2/IFN | - | - | - | .893 | .043 | .866 |
| IL2/TNF | - | - | - | - | .043 | .500 |
| IFN/TNF | - | - | - | - | - | .043 |

Supplementary table 5. Statistical comparisons of telomere length of CD8⁺ T cell cytokine producing populations following overnight stimulation of PBMCs with NLV peptide. Results from Wilcoxon Signed Rank Test.

| | IFN | TNF | IL2/IFN | IL2/TNF | IFN/TNF | IL2/IFN/TNF | Negative |
|--------------|-----|------|---------|---------|---------|-------------|----------|
| Young | | | | | | | |
| IL2 | ns | .028 | .043 | ns | .028 | ns | ns |
| IFN | - | .028 | ns | ns | .028 | ns | ns |
| TNF | - | - | ns | ns | ns | ns | .028 |
| IL2/IFN | - | - | - | ns | ns | ns | ns |
| IL2/TNF | - | - | - | - | ns | ns | ns |
| IFN/TNF | - | - | - | - | - | ns | .028 |
| IL2/IFN/TNF | - | - | - | - | - | - | ns |
| Old | | | | | | | |
| IL2 | ns | ns | ns | ns | ns | ns | ns |
| IFN | - | ns | ns | ns | .043 | ns | .043 |
| TNF | - | - | ns | ns | ns | ns | .043 |
| IL2/IFN | - | - | - | ns | ns | ns | ns |
| IL2/TNF | - | - | - | - | ns | ns | - |
| IFN/TNF | - | - | - | - | - | ns | .043 |
| IL2/IFN/TNF | - | - | - | - | - | - | ns |

