

**Supplementary Fig. 1** (related to Fig. 2). Common differentially expressed genes (DEGs) between irradiated young male and irradiated young female mice (*left*) and irradiated old male and irradiated old female mice (*right*). Below the Venn diagrams, the total number of unique and common differentially expressed genes in both sexes is depicted and the percentage of common genes is calculated. M: male; F: female. MY-IR: male young-irradiated; FY: female young-irradiated; MO-IR: male-old-irradiated; FO-IR: female old-irradiated.

**Table S1a**. Immune cell death-related functions between young and old female irradiated mice.

| Temale inaciated mice.               |         |        |
|--------------------------------------|---------|--------|
|                                      | Young   | Old    |
| Diseases and Functions Annotation    | z-score |        |
| Cell death of immune cells           | 3.920   | 0.604  |
| Cell death of blood cells            | 3.774   | 0.187  |
| Cell death of B lymphocytes          | 2.706   | 0.000  |
| Apoptosis of leukocytes              | 2.705   | -0.449 |
| Cell death of myeloid cells          | 2.537   | 0.000  |
| Cell death of lymphocytes            | 2.370   | 0.000  |
| Cell death of lymphatic system cells | 2.198   | 0.000  |
| Cell death of lymphoid cells         | 2.182   | 0.000  |
| Cell death of mononuclear leukocytes | 2.168   | -0.033 |
| Production of antibody               | -2.346  | -2.020 |
| Quantity of immunoglobulin           | -2.659  | -2.204 |
| Quantity of IgG1                     | -3.046  | 0.000  |
| Quantity of lymphoid tissue          | -2.378  | -0.637 |
| Quantity of B lymphocytes            | -3.155  | 0.000  |
| Quantity of IgG                      | -2.562  | 0.000  |

**Table S1b**. Immune cell death-related functions between young and old female and male irradiated mice.

|  | Female  |        | Male   |        |
|--|---------|--------|--------|--------|
|  | 2-mo    | 24-mo  | 2-mo   | 21-mo  |
| <b>Diseases and Functions Annotation</b> | z-score |        |        |        |
| Cell death of immune cells               | 3.920   | 0.604  | 2.285  | -0.276 |
| Cell death of blood cells                | 3.774   | 0.187  | 2.152  | -0.271 |
| Cell death of B lymphocytes              | 2.706   | 0.000  | 2.973  | 1.646  |
| Apoptosis of leukocytes                  | 2.705   | -0.449 | 2.700  | 0.411  |
| Cell death of myeloid cells              | 2.537   | 0.000  | 0.000  | 0.000  |
| Cell death of lymphocytes                | 2.370   | 0.000  | 3.063  | 1.051  |
| Cell death of lymphatic system cells     | 2.198   | 0.000  | 2.774  | 1.131  |
| Apoptyosis of lymphoid cells             | 2.056   | 0.000  | 2.620  | 0.000  |
| Apoptosis of mononuclear leukocytes      | 2.008   | -0.033 | 2.632  | 0.697  |
| Quantity of lymphoid tissue              | -2.378  | -0.637 | -3.742 | -2.028 |
| Quantity of B lymphocytes                | -3.155  | 0.000  | -4.166 | -3.120 |
| Quantity of T lymphocytes                | 0.000   | 0.000  | -2.538 | -1.260 |
| Quantity of lymphocytes                  | 0.000   | 0.000  | -3.206 | -1.960 |

**Table S2a**. Phagocytosis-related functions between young and old female mice exposed to 4 Gy x-rays.

| young and old remaie mice exposed to      | •       | _     |
|---|---------|-------|
|   | Young   | Old   |
| <b>Diseases and Functions Annotations</b> | z-score |       |
| Endocytosis                               | 5.861   | 3.029 |
| Cell movement of phagocytes               | 5.113   | 3.382 |
| Engulfment of cells                       | 5.100   | 3.082 |
| Endocytosis by eukaryotic cells           | 5.068   | 2.496 |
| Phagocytosis                              | 5.061   | 2.470 |
| Migration of phagocytes                   | 4.519   | 2.150 |
| Phagocytosis of cells                     | 4.467   | 2.643 |
| Chemotaxis of phagocytes                  | 4.317   |       |
| Recruitment of phagocytes                 | 4.266   | 3.052 |
| Activation of phagocytes                  | 3.958   |       |
| Phagocytosis of myeloid cells             | 3.678   |       |
| Engulfment of blood cells                 | 3.595   |       |
| Engulfment of leukocytes                  | 3.507   |       |
| Phagocytosis of blood cells               | 3.506   |       |
| Phagocytosis of leukocytes                | 3.415   |       |
| Phagocytosis of phagocytes                | 3.269   |       |
| Engulfment of phagocytes                  | 2.971   |       |
| Degranulation of phagocytes               | 2.69    |       |
| Cell death of phagocytes                  | 2.603   |       |
| Cellular infiltration by phagocytes       | 2.497   |       |
| Immune response of phagocytes             | 2.325   |       |
| Cell viability of phagocytes              | 2.259   |       |
| Apoptosis of phagocytes                   | 2.073   |       |
| Binding of professional phagocytic cells  |         | 2.305 |

**Table S2b**. Comparison of phagocytosis-related functions between male and female mice exposed to 4 Gy x-rays.

|  | Young | Old   |
|--|-------|-------|
| Diseases or Functions Annotation         | z-sc  | ore   |
| Phagocytosis                             | 4.372 |       |
| Phagocytosis of cells                    | 4.123 | 2.042 |
| Engulfment of cells                      | 3.943 |       |
| Internalization of cells                 | 3.778 |       |
| Cell movement of phagocytes              | 3.712 |       |
| Endocytosis                              | 3.519 |       |
| Endocytosis by eukaryotic cells          | 3.416 | 2.162 |
| Cell death of phagocytes                 | 3.163 |       |
| Phagocytosis of blood cells              | 3.145 |       |
| Phagocytosis of phagocytes               | 3.127 |       |
| Apoptosis of phagocytes                  | 3.054 |       |
| Phagocytosis of myeloid cells            | 3.020 |       |
| Phagocytosis of leukocytes               | 3.008 |       |
| Engulfment of myeloid cells              | 2.925 |       |
| Migration of phagocytes                  | 2.852 |       |
| Engulfment of blood cells                | 2.790 |       |
| Phagocytosis of antigen presenting cells | 2.700 |       |
| Engulfment of leukocytes                 | 2.648 |       |
| Engulfment of phagocytes                 | 2.548 |       |
| Engulfment by macrophages                | 2.236 |       |
| Recruitment of phagocytes                | 2.112 |       |

 Table S3a.
 Neutrophil-associated functions among irradiated mice of

different sex and age.

|   | female  |       | male  |       |
|---|---------|-------|-------|-------|
|   | 2-mo    | 24-mo | 2-mo  | 21-mo |
| <b>Diseases or Functions Annotation</b> | z-score |       |       |       |
| Recruitment of neutrophils              | 3.829   |       | 2.944 |       |
| Cell movement of neutrophils            | 3.096   | 3.280 |       |       |
| Chemotaxis of neutrophils               | 4.523   |       |       |       |
| Migration of neutrophils                | 2.700   |       |       |       |
| Infiltration by neutrophils             | 2.200   |       |       |       |
| Degranulation of neutrophils            | 2.121   |       |       |       |

**Table S3b**. Reactive oxygen species synthesis and metabolism-associated functions among irradiated mice of different sex and age.

|                                       | female |        | male  |       |
|---------------------------------------|--------|--------|-------|-------|
|                                       | 2-mo   | 24-mo  | 2-mo  | 21-mo |
| Diseases or Functions Annotation      |        | z-scor | e     |       |
| Metabolism of reactive oxygen species | 5.821  | 2.278  | 3.489 |       |
| Synthesis of reactive oxygen species  | 5.645  | 2.267  | 3.291 |       |
| Production of reactive oxygen species | 4.47   | 2.031  | 3.224 |       |
| Production of superoxide              | 2.684  |        |       |       |

**Table S3c**. Myeloid-associated functions among irradiated mice of different sex and age.

| 3                                      | fem     | ماد   | male   |       |
|--|---------|-------|--------|-------|
|  | IGIII   | laie  | IIIali | -     |
|  | 2-mo    | 24-mo | 2-mo   | 21-mo |
| Diseases or Functions Annotation       | z-score |       |        |       |
| Cell movement of myeloid cells         | 5.22    | 3.508 | 2.454  |       |
| Recruitment of myeloid cells           | 4.338   |       |        |       |
| Activation of myeloid cells            | 4.184   |       | 3.665  |       |
| Chemotaxis of myeloid cells            | 4.523   | 3.315 |        |       |
| Binding of myeloid cells               | 3.835   | 2.975 | 3.218  |       |
| Phagocytosis of myeloid cells          | 3.678   |       |        |       |
| Adhesion of myeloid cells              | 3.386   |       |        |       |
| Migration of myeloid cells             | 3.311   |       |        |       |
| Response of myeloid cells              | 3.155   |       | 3.089  |       |
| Cellular infiltration by myeloid cells | 2.885   |       |        |       |
| Engulfment of myeloid cells            |         |       | 3.745  |       |
| Recruitment of myeloid cells           |         |       | 2.713  |       |