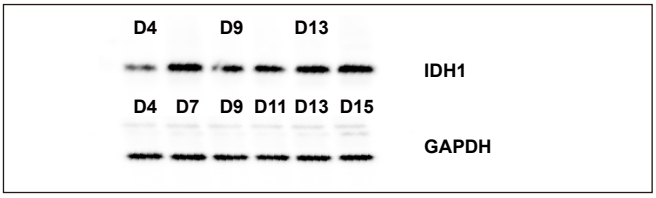**Figure S1A**